The association of COVID-19 incidence with temperature, humidity, and UV radiation – A global multi-city analysis

5 Abstract

4

Background and Aim. The associations between COVID-19 transmission and 6 meteorological factors are scientifically debated. Several studies have been conducted 7 worldwide, with inconsistent findings. However, often these studies 8 had methodological issues, e.g., did not exclude important confounding factors, or had 9 limited geographic or temporal resolution. Our aim was to quantify associations 10 between temporal variations in COVID-19 incidence and meteorological variables 11 globally. 12

Methods. We analysed data from 455 cities across 20 countries from 3 February to 13 31 October 2020. We used a time-series analysis that assumes a quasi-Poisson 14 distribution of the cases and incorporates distributed lag non-linear modelling for the 15 exposure associations at the city-level while considering effects of autocorrelation, 16 long-term trends, and day of the week. The confounding by governmental measures 17 was accounted for by incorporating the Oxford Governmental Stringency Index. The 18 effects of daily mean air temperature, relative and absolute humidity, and UV radiation 19 were estimated by applying a meta-regression of local estimates with multi-level 20 random effects for location, country, and climatic zone. 21

Results. We found that air temperature and absolute humidity influenced the spread
 of COVID-19 over a lag period of 15 days. Pooling the estimates globally showed that
 overall low temperatures (7.5°C compared to 17.0°C) and low absolute humidity (6.0

g/m³ compared to 11.0 g/m³) were associated with higher COVID-19 incidence (RR temp =1.33 with 95%CI: 1.08; 1.64 and RR AH =1.33 with 95%CI: 1.12; 1.57). RH revealed no significant trend and for UV some evidence of a positive association was found. These results were robust to sensitivity analysis. However, the study results also emphasise the heterogeneity of these associations in different countries.

Conclusion. Globally, our results suggest that comparatively low temperatures and
 low absolute humidity were associated with increased risks of COVID-19 incidence.
 However, this study underlines regional heterogeneity of weather-related effects on
 COVID-19 transmission.

Key words: Temperature, Humidity, UV Radiation, COVID-19, Distributed Lag Non Linear Modelling, Global Analysis

37 Abbreviations

| Abbreviation | Meaning |
|--------------|--|
| AC | Autocorrelation |
| AH | Absolute Humidity |
| BLUP | Best Linear Unbiased Prediction |
| CAMS | Copernicus Atmosphere Monitoring Service |
| СВ | Crossbasis |
| CI | Confidence Interval |
| COVID-19 | Coronavirus Disease 2019 |
| DLNM | Distributed Lag Non-Linear Models |
| df | degrees of freedom |
| ERA5 | Earth Reanalysis Dataset 5 |
| GSI | Government Stringency Index |
| JHU | John Hopkins University |
| MERS | Middle East Respiratory Syndrome |
| NS | Natural Spline |
| OxCGRT | Oxford COVID-19 Government Response |
| | Tracker |
| PACF | Partial Autocorrelation Function |
| РМ | Particulate Matter |
| Q-AIC | Quasi Akaike Information Criterium |
| REML | Restricted maximum likelihood method |
| RH | Relative Humidity |
| RR | Risk Ratio or Relative Risk |
| SARS | Severe Acute Respiratory Syndrome |
| SARS-CoV-2 | SARS Coronavirus 2 |
| SIR | Susceptible Infectious Recovered |
| TSR | Time Series Regression |

39 **1. Introduction**

The severe acute respiratory syndrome coronavirus-2 (SARS-CoV-2) pandemic arose 40 in late December 2019 in Wuhan, China. According to the WHO, by 12 July 2021 190 41 million cases and 4 million deaths had been reported globally due to coronavirus 42 disease 2019 (COVID-19).¹ Evidence points towards transmission mainly taking place 43 via airborne transmission (respiration of SARS-CoV-2 containing droplets).² Other 44 modes of transmission, including direct contact through contaminated surfaces, faecal-45 oral transmission and other body fluids are still under investigation regarding the extent 46 to which they influence the infection dynamics.^{3,4} 47

The relationship between COVID-19 incidence and meteorological factors is greatly 48 discussed in the literature and of high public interest. A connection between 49 meteorology and COVID-19 is considered likely as other coronaviruses and respiratory 50 viruses show strong seasonal patterns of disease incidence that can to some extent 51 be explained by meteorological factors in temperate regions.^{5,6} There are several ways 52 in which meteorological factors (e.g. air temperature and humidity) could influence 53 COVID-19 incidence. Extreme climatic conditions (e.g., extreme cold and heat) can 54 result in people spending more time indoors, in closed, poorly ventilated spaces, which 55 can increase the transmission of SARS-CoV-2.7,8 Moreover, lower temperatures 56 enhance the stability of viral lipid envelopes and lower humidity favours droplet nuclei 57 formation which prolong viability and transmissibility of SARS-CoV-2 9,10,11,12,13 Also, 58 cold and dry conditions affect the human innate and adaptive immune response in 59 various ways (e.g., in cold nostrils through inhibited mucociliary clearance and a 60 decrease of phago- and leukocyte activity, which changes the likeliness of infection or 61 symptom severity). ^{14,15,16,17} Altogether, these mechanisms support the hypothesis that 62

colder and drier conditions would favour SARS-CoV-2 transmission and increase
 COVID-19 incidence.

Several spatial ecological and time-series studies have investigated the association 65 between meteorological conditions and COVID-19 cases.^{18,19} However, so far the 66 literature remains mainly inconclusive showing positive, negative, and no associations 67 for temperature, humidity (relative and absolute) and UV radiation in different 68 analyses.^{20–25,26–33} The variation in study results could partially be explained by varying 69 70 spatial scales of analysis, application of different statistical methods with varying degrees of sophistication, and varying levels of consideration of potential confounding 71 factors. Moreover, according to previous systematic reviews, epidemiologic studies 72 assessing the relationship between weather and COVID-19 incidence could have 73 methodologic limitations that may introduce bias and limit causal inference.^{34–36} For 74 example, many studies did not consider the possibility of a non-linear relation and 75 lagged effects of weather and incidence, they did not account for time-varying 76 confounders, and they did not consider location-specific confounders. To address 77 these limitations time-series regression methods could be used. These methods have 78 been used to quantify short-term associations of environmental exposures with health 79 outcomes, notably with infectious diseases.³⁷ Time-series regression methods allows 80 long-term trends, other time-varying cofounding 81 seasonality, factors, and autocorrelation to be controlled for. It also allows us to explore the association with 82 delayed and non-linear exposure effects.³⁸ With the availability of longer time-series 83 several studies have used time-series methods to evaluate the association between 84 meteorological factors and COVID-19 incidence.^{39–50} Among those, three studies were 85 performed on a global scale, ^{40,44} but they considered the country as unit of analysis. 86 City-level studies are more appropriate given the lower measurement error on the 87

outcome and on the exposure. Moreover, they allow accounting for phenomena, like
 high levels of population density or human mobility, which are only observable on a
 small scale.³⁸

The aim of this study is to use city-level time-series models to evaluate the association between meteorological exposures (e.g., temperature, humidity, and UV radiation) and COVID-19 incidence at the global scale.

94

95 2. Methods

96 2.1 Data sources and extraction

The data extraction was performed by members of the Multi-City Multi-Country (MCC) Network, an international research network focused on the study of environmental conditions, climate change, and human health (https://mccstudy.lshtm.ac.uk/). We considered the COVID-19 case time-series data for 455 cities between 3 February and 31 October 2020. Details of the cities and sources can be found in Supplementary Table S1.

We obtained exposure data from the Copernicus ERA5 dataset with a latitudelongitude grid size of $0.25^{\circ} \times 0.25^{\circ}$ (roughly 28x28km).⁵¹ We selected temperature and dew temperature in 2 m above the surface as well as the surface downwelling shortwave radiation (solar UV radiation, J/m²). For these variables daily averages were taken from the closest grid cell for each city or small region.

We calculated the relative humidity (RH) from temperature and absolute humidity (AH)
 using the R "humidity" package.⁵² RH measures the percentage of water molecules in
 the air relative to concentration at full saturation, whereas AH measures the amount of

water vapor in a specific volume of air.⁵³ This is the formula of how AH relates to RH
 and temperature:⁵⁴

113
$$AH(g/m^3) = \frac{6.112 \times e^{\frac{17.67 \times T(^{\circ}C)}{T(^{\circ}C)+243.5}} \times RH(\%) \times 2.1674}{273.15 + T(^{\circ}C)}$$

The following variables were captured as we expected them to be confounders of the 114 associations between weather variables and COVID-19 incidence. We extracted the 115 Government Stringency Index (GSI) from the Oxford COVID-19 Government 116 Response Tracker (OxCGRT) to control for changing governmental public health 117 measures implemented in response to the pandemic.⁵⁵ The GSI scale ranges from 0 118 to 100 points with 100 representing the most strict measures implemented to hinder 119 COVID-19 transmission such as closure policies, movement restrictions, income 120 support, and testing policies. For the purpose of sensitivity analysis, we also used 121 residential mobility from the Google Mobility index which measures the change in 122 average duration of time spent at home compared to the median for the same weekday 123 in a pre-pandemic period (3 January to 6 February 2020).⁵⁶ 124

We considered the long-term mean temperature, demographic information on population size, density, and age proportion above 65 years in the fixed effects of the meta-regression. Demographic variables were collected from the Organisation for Economic Co-operation and Development (OECD) Global Human Settlement Layer Urban Centre Database unless specified otherwise in the results.⁵⁷ This data was available at the city-level from the MCC Network.

132 2.2 Statistical analysis

133 **2.2.1 Descriptive analysis**

For the descriptive data analysis, the daily and cumulative COVID-19 cases of the included cities were summed for each country and the cases per 100.000 inhabitants were calculated using total population size of each city (OECD data).⁵⁷

137 2.2.2 Two-stage design

We used a two-stage design to assess the association between the meteorological factors and COVID-19 incidence. The first stage consists of estimating the city-specific exposure-response association considering time-varying confounding in a time-series regression (TSR). In the second stage, a meta-analytic model is used to combine the city-specific estimates to obtain the pooled exposure-response association curve.

For the first stage of the analysis, independent models for each exposure were 143 formulated for all locations. The city-specific time-series were shortened to start up to 144 15 days (depending on the considered days of lag) before the first time that 10 cases 145 occurred in that city. This aims to exclude first imported cases. The exposures were 146 modelled using distributed lag non-linear models (DLNMs).⁵⁸ The basis function for the 147 exposure dimension (temperature, AH, RH, and UV radiation) was chosen as a 2nd 148 degree polynomial. The lag dimension was modelled with a natural cubic spline 149 containing two equally spaced (at logarithmic scale) internal knots. In the main 150 analysis, a lag of 15 days was considered, since the incubation period was estimated 151 to be around 6 days for COVID-19^{59,60} and there is a delay in testing and reporting. 152 The two bases were then combined to make a bi-dimensional basis called a "cross-153 basis".⁶¹ The residual variation of case counts was assumed to follow a guasi-Poisson 154 distribution. 155

Several confounding factors were considered in the main model. Since the reporting, 156 as well as many other factors (e.g., social behaviour and testing capacities), might vary 157 between weekdays, we included a series of dummy day of the week variable (dow). 158 Two other time-varying confounders were considered. The intra-year trend of COVID-159 19 was considered in the model using a natural spline function of the date with 160 6 degrees of freedom (df) (NS(date, df = 6)) which equals approximately 1.5 df per 161 month. Changing governmental public health measures were modelled with a linear 162 lag association model of GSI considering up to 15 days of lag dependence (CB_{GSI}). 163 The model was built using the R package "dlnm".⁶¹ An autocorrelation term was 164 included to account for transmission dynamics.³⁷ For this purpose, the logarithm of one 165 day lagged cases added to 0.5 was included (AC). 166

In summary, the basic first stage model for each exposure (temperature, RH, AH, or
 UV) which was performed for each city looked like this:

169
$$Ln(y) \sim CB_{exposure, lag=15} + AC + dow + CB_{GSI} + NS(date, df = 6)$$
(I)

For the subsequent second stage meta-analysis, the R package "mixmeta" was used.⁶² 170 The coefficients representing estimated meteorology to COVID-19 associations were 171 cumulated over all lags and their covariance matrices which were obtained at the first 172 step were pooled over all included locations using a random effect meta-analytic 173 model. We used the estimation method of restricted maximum likelihood (REML). In 174 the main model (Model A), we considered groups defined jointly by country and climatic 175 zones as random effects. The same model was used in Model B but only for the subset 176 of locations with complete data in the meta-predictors (GDP, mean temperature, and 177 % of population aged more than 65 years). In the subset of locations with complete 178 data, we then also fitted the meta-regression model with the meta-predictors as fixed 179 effects (Model C). To evaluate the role of country in explaining the heterogeneity in the 180

association curves, we considered models with city as a random effect and country as
 a fixed effect (Model D), or random effect (Model E). We then derived country-level
 Best Linear Unbiased Prediction (BLUP) curves from Model E.

Using the pooled polynomial basis coefficients, we plotted the pooled mean curve (for all included cities) of COVID-19 risk against each exposure (temperature, RH, AH and UV) expressed as relative risk (RR) to the median level which was set as the minimum exposure value.

188 2.3 Sensitivity analysis

We performed a sensitivity analysis of the observed effect on the days of lags 189 accounted in the first stage model by varying the length of the lag period from 15 to 10 190 days. The influence of choice of df used to model intra-year trends was also explored 191 by altering from 6 df to 4 df. Furthermore, we evaluated the possible time-varying 192 confounding of air pollution by considering city-level particulate matter (PM₁₀) data in 193 the first stage model using a distributed linear model (DLM) parametrization and up to 194 15 days of lag. The PM10 data was obtained from the Copernicus Atmosphere 195 Monitoring Service (CAMS) global near-real time service.^{63–65} The hourly modelled 196 values of surface PM10 (0.4 x 0.4 arc degrees grid cell resolution) were averaged daily 197 over the observation period and linked to the city using the city centroid coordinates. 198 The statistical analysis was performed using R 4.1.2 statistical software. 199

200

201 **3. Results**

202 **3.1 Descriptive analysis**

This analysis considered 10.5 million confirmed COVID-19 cases across 455 different cities in 20 countries between 3 February and 31 October 2020. The city locations are

shown below as well as the country-wide aggregated time-series of daily reported 205 COVID-19 cases per 100.000 in the included city populations (Figure 1 and 2). In most 206 of the countries in the northern hemisphere we can recognise two waves in late winter 207 or early spring and in autumn, while countries in the southern hemisphere (e.g., Brazil, 208 Chile, Peru, and South Africa) experienced a single wave during the observation period 209 of this study. Table 1 shows the country-wide cumulative incidence per 100,000 210 inhabitants which varied from 49 in South Korea to 8,350 in the USA. The average 211 minimum and maximum recorded exposures per country within the observation period 212 are reported in Table 1. Daily country averages of the meteorological variables over 213 the observation period are represented in Supplementary Figures S1-4. Countries in a 214 tropical climate or in the southern hemisphere (e.g., Brazil, Chile, Mexico, Peru, 215 Singapore, and South Africa) show less variation of the meteorological variables, 216 especially mean temperature, RH and AH. The correlation between the four exposures 217 is shown in Supplementary Table S4. An overview of the governmental interventions 218 against COVID-19 over time can be seen in Supplementary Figure S5. Most countries 219 started out with stringent restrictions in the beginning of 2020 and loosened them by 220 the middle of the year. Some tightened them again towards the end of October 2020 221 (Supplementary Figure S5). Estonia had the lowest overall level of governmental 222 interventions with an average GSI of 36.9% during the observation period, whereas 223 Peru ranked highest on governmental stringency with an average GSI at 75.8% (Table 224 1). 225

226 3.2 Association between COVID-19 cases and temperature

The pooled association curve, representing overall results across all cities, obtained from the pooled models for temperature exposure (Model A) is represented in Figure 3a. Low temperatures were associated with higher risk of infection. At 7.5°C the relative

risk of COVID-19 incidence is 1.33-fold higher (CI-95%: 1.08;1.64) compared to a 230 reference level at 17.0°C. The exposure-lag association indicated increased RRs with 231 a 3-day lag after temperature exposure, reached a peak at 8-9 days, and decayed by 232 the end of the observed 15 days' lag period (Supplementary Figure S6a). We observed 233 a substantial heterogeneity in the meta-analytic model (I²=67.3%). Investigating the 234 city-level factors which could explain this heterogeneity (Model C), we found that old 235 population (% population aged more than 65 years), the average daily mean 236 temperature, and GDP modified the association between temperature and COVID-19 237 incidence (Supplementary Figure S7a, Supplementary Table S2). Cities with an older 238 population and lower long-term mean temperature seemed to have a higher impact of 239 lower temperature on COVID-19 spread, but overall, these factors explain only 1.1% 240 of heterogeneity. We also investigated the role of country on heterogeneity comparing 241 the meta-analytic model with and without country modelled as fixed effect with an I² 242 decrease equal to 4.3% (Supplementary Table S2). The Figure 4 shows the country 243 specific curves obtained using BLUPs prediction from the Model E with country as a 244 random effect. We observed different patterns of the temperature COVID-19 incidence 245 curve with most countries showing curves with higher COVID-19 incidence with cold 246 temperatures (e.g. Chile, Czech Republic, Estonia, Germany, Italy, Japan, Kuwait, 247 Romania, Spain and UK), some with limited exposure variation had a flat curve (Brazil, 248 Peru, Singapore and South Africa), three had no evidence of an association (France, 249 Canada and US), and three showed a tendency of increased COVID-19 risk with higher 250 temperatures (Finland, South Korea, Mexico). 251

252 3.3 Association between COVID-19 cases and Humidity

Overall, little evidence was found for an association between relative humidity and
 COVID-19 spread (Model A), with a slight tendency of a lower risk of infection for higher

level of RH (Figure 3b). With respect to a reference level set at 65% RH, the RR of 255 observing COVID-19 cases was 0.89 at 85% RH (CI-95%: 0.75; 1.06). This association 256 did not diverge from RR=1.00 when considering different lags (Supplementary Figure 257 S6b). There was substantial heterogeneity in this association (I²=68.3%), but 258 examination of meta-predictors and country specific curves showed no interpretable 259 patterns (Supplementary Figure S7b). Country modelled as a fixed effect explained 260 3.6% of the heterogeneity (Supplementary Table S2). Figure 5 shows the country 261 specific curves obtained using BLUPs prediction from the model with country as 262 random effect (Model E). Adjusting for daily mean temperature gives a tendency of a 263 protective effect at higher levels of RH (Supplementary Figure S8). 264

For AH, we observed an inverse association (Model A in Figure 3c). Compared to the 265 median value of 11.0 g/m³ there was a 1.33-fold increased RR at the AH of 6.0 g/m³ 266 (95%-CI: 1.12; 1.57). The RRs were observed to be increased (RR>1.00) between 3 267 to 15 days of lag (Supplementary Figure S6c). The meta-predictors old population, 268 long-term mean temperature and GDP explained 3.7% of the heterogeneity 269 (Supplementary Figure S7c, Supplementary Table S2). Cities with higher long-term 270 mean temperature show a lower risk of COVID-19 infection associated with high levels 271 of AH. Country modelled as fixed effect explained 4.7% of the heterogeneity. Country 272 BLUPs estimates are presented in Figure 6 (Model E). As observed for temperature, 273 we found different patterns of the association between AH and COVID-19 incidence in 274 different countries. There are countries with higher COVID-19 incidence with low AH 275 (e.g., Chile, Czech Republic, Estonia, France, Japan, Spain and UK), countries with 276 no evidence of an association (Brazil, Kuwait, Mexico, Italy, Romania, Singapore, 277 South Africa and US), and countries showing a tendency of increased COVID-19 risk 278 with higher AH (Canada, Finland, and South Korea). 279

281

3.4 Association between COVID-19 cases and UV

We found some evidence of an association between UV exposure and COVID-19 spread (Model A in Figure 3d). Meta-predictors have little influence on the association curve explaining only 1.2% of the l² (Supplementary Figure S6d, Supplementary Table S2). Country modelled as fixed effect explained 3.4% of the heterogeneity. Country BLUPs estimates are presented in Figure 7, with some countries (Canada, Finland, Kuwait, Mexico, Spain and US) showing lower COVID-19 incidence with lower levels of UV radiation.

289 3.5 Sensitivity analysis

A sensitivity analysis was conducted to assess the robustness of estimates of the 290 previously described models. A decrease to 10 days of lag or lower degrees of freedom 291 of long-term trend (4 df instead of 6 df) in general led to similar association curves 292 (Supplementary Figure S10 and S11). Also, the inclusion of PM₁₀ into the first stage 293 model resulted in no major change of the exposure to COVID-19 associations 294 (Supplementary Figure S12). Stratifying the analysis according to climatic zone, for air 295 temperature all curves show a decreasing trend. For RH tropical cities show a higher 296 COVID-19 spread in dry conditions. More variability was observed for AH and UV 297 radiation (Supplementary Figure S13). 298

299

300 4. Discussion

301 **4.1 Main findings**

Overall, this study supports previous findings that temperature and absolute humidity are environmental factors that potentially influence the spread of COVID-19. Globally, low temperatures and low absolute humidity were associated with higher COVID-19 incidences, but for RH no evidence of an association was found. There was substantial heterogeneity in the associations of the respective environmental exposures and COVID-19 risk between countries.

308 4.2 Possible biological and behavioural mechanisms

Our results can be viewed in light of previous studies investigating the mechanistic 309 principles behind associations between meteorological variables and COVID-19. The 310 observation that low temperatures lead to higher transmission rates of viral disease 311 has been made in many previous studies. Biophysical theory and laboratory results 312 suggest that lower temperatures support the stability and viability of viral particles.^{66,67} 313 Additionally, animal experiments hint towards a connection with lower blood circulation 314 and consequent local impairment of adaptive immunity at low temperatures, thereby 315 affecting the host's immune system's ability to fight respiratory viruses.^{14,68} 316

The association between lower levels of humidity and higher levels of infections could be explained by virus-containing droplets having short ballistic settling characteristics under wet conditions. In contrast at dry conditions, droplets evaporate forming dry nuclei that are able to maintain floating over longer durations of time.^{12,69} Influenzarelated studies also hinted at an impaired immune response under dry conditions (e.g., through impaired mucociliary clearance and other innate responses).¹⁵ A US study found that outdoor AH is a good predictor for indoor AH while this is not true for RH.⁷⁰

Hence, it could be that AH is a more useful predictor for COVID-19 incidence than RH. Previous studies came to that conclusion regarding AH as predictor for influenza transmission rates as well.⁷¹ However, the high correlation between AH and temperature (r=0.88, average of all cities in our dataset) implies that it is difficult to disentangle effects of the two exposures, with one of the associations possibly merely reflecting confounding by the other.

There was some evidence of a positive association between COVID-19 cases and UV radiation. This was unexpected, as one hypothesis is that UV light could cause inactivation of viruses in the air and on surfaces. Also, there is a theory that more solar radiation could lead to less vitamin D deficiency (contributing to better functioning immune system).⁷²

4.3 *Comparison to other modelling studies*

Due to the extensively growing literature in this field, the state of scientific knowledge 336 on this topic is constantly evolving. A review from late 2020 reporting on about 60 337 studies on associations between COVID-19 and weather identified a variety of findings 338 for temperature and humidity.²¹ The included studies that reported a linear trend mostly 339 showed a negative association bewteen COVID-19 cases and temperature as well as 340 humidity (33 vs. 6 studies and 13 vs. 3, respectively). Global analyses support these 341 local findings for temperature and humidity. Using different methodologies Sarkodie et 342 al., Wu et al., Yuan et al., and Zhang et al. all found a negative association between 343 temperature and RH with COVID-19 case rates in 20 countries, 166 countries, 127 344 countries, and 1236 regions globally with data until April, March, August, and May 345 2020, respectively.^{73–76} The study from Yuan et al. also widened their analysis to 346 include 188 countries with data through December 2020, and also analysed the 347 non-linear associations, showing similar exposure response associations as found in 348

our study using generalized additive model and as well DLNM methods.44,75 The 349 temperature of minimum COVID-19 risk in both Yuan et al. studies was around 20°C 350 and for RH the risk was highest at humidity around 70%. Another global study using 351 DLNM from Guo et al. including 190 countries showed a similar association for 352 temperature (highest RR at 5°C and lowest at 20°C) but exhibited a different exposure-353 risk association for RH (risk maximum at 72% RH).77 Two studies that also used DLNM 354 models on US counties only also found elevated infection risks (increased Rt levels) at 355 lower temperatures and one of them as well for lower specific humidities.^{29,32} Fontal et 356 al. analysed the transitory associations of temperature and AH until October 2020 in 357 10 world regions and obtained negative associations for both.³³ One global study did 358 find only a small effect of temperature in 3739 global locations (Xu et al.) and two global 359 studies did not find a statistically relevant effect for temperature (Carleton et al., Islam 360 et al.) and RH (Islam et al.) in 206 countries and 3235 regions, respectively.^{30,27,78} 361 However, Guo et al., Xu et al., Carleton et al., and Islam et al. all had a comparatively 362 short study period reaching until April 2020.77,30,27,78 363

We recently performed a different global city-level analysis of meteorological factors 364 and SARS-CoV-2 transmission.²² This used an ecological approach comparing 365 effective reproduction number (Re) and meteorological variables between cities in the 366 early phase of the pandemic, and identified a non-linear (though primarily downward) 367 association between mean temperature, and absolute humidity with Re, and a 368 tendency of a negative association between RH and Re. Non-pharmaceutical 369 interventions had a greater effect on Re. The results of the current study complement 370 our previous results that showed higher Re at lower mean temperature, lower absolute 371 humidity, and a negative association between RH and Re. 372

Regarding UV exposure, out of the 60 studies analysed in the previously mentioned 373 systematic review, only six analysed solar radiation and among those there was no 374 consensus of whether there is an association and if so what type of association.²¹ Two 375 of the previously mentioned global analyses also included UV variables. Carleton et al. 376 reported in contrast to our study that higher UV radiations were associated with lower 377 COVID-19 growth rates, whereas Islam et al. concluded the relationship to be 378 inconclusive within the same time period.^{27,78} 379

380 4.4

Strength and Weaknesses

381 Our study has several important strengths. It considered a multitude of locations globally with smaller spatial units of analysis and longer observation periods than most 382 published studies. Lagged effects of exposure were considered, as were potential non-383 linear relationships of the exposure with COVID-19 incidence. Ecological and time-384 varying confounders were analysed and incorporated. 385

Possible short comings of this study are that the case definitions differed from country 386 to country, that GSI might not adjust sufficiently for changes in governmental measures 387 over time, and that the distribution of cities included is not equally distributed around 388 the globe, with some regions underrepresented and only few locations close to the 389 equator. Thus, while this study is one of the most detailed global analyses to date, the 390 pooled estimates provide insights into the associations in the included cities but are 391 not fully representative for everywhere around the globe. Also, a global estimate itself 392 might be of limited use due to the heterogeneity amongst locations that was 393 encountered. We considered factors explaining this heterogeneity and we found that 394 long-term mean temperature (a proxy of the city climate) and the percentage of the 395 population older than 65 years modify the association found. There was a tendency in 396 cities with lower long-term temperature and older population to have higher COVID-19 397

incidence in colder and drier conditions, but these factors explain only a small amount
of the observed heterogeneity leading to some difference among countries. These
differences could be due to limited sample size in some countries (e.g., Estonia,
Finland, South Africa), and different and narrower ranges of exposure experience in
countries (e.g. Brazil, Mexico, Chile, Peru and Singapore). Moreover, the observed
differences could also be due to different adaptation of local populations to various
weather conditions.

405

406 **5. Conclusion**

This study indicates that there is a tendency of a higher risk of COVID-19 cases at low 407 temperature or absolute humidity levels, which aligns to an extent with available 408 mechanistic explanations and previous literature basis. The between country 409 heterogeneity of weather-related effects on COVID-19 when applying our uniform 410 modelling framework in a global analysis shows the importance of determining location 411 specific estimates of meteorological effects on COVID-19 spread. As more data 412 accumulates, studies using longer observational periods will help elucidate weather-413 sensitivity and seasonal patterns of COVID-19 transmission. 414

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Table 1: Summary Table of observed COVID-19 cases, meteorological exposures,
 and governmental stringency index in the different countries.

| Country | Number of included cities | Cumulative cases per day [# per 100.000] | Daily mean temperature [°C] | Daily mean RH [%] | Daily mean AH [g/m³] | Daily mean UV [J/m²] | OxCGRT GSI [%] |
|-------------------|---------------------------------|--|-----------------------------------|-------------------------|----------------------------|----------------------------|----------------------|
| Brazil | 13 | 1677 | 24.3 (4.7, 31.7) | 76.5 (30.2, 99.0) | 17.5 (5.9, 23.6) | 210.7 (20.3, 353.7) | 65.0 (51.6, 69.8) |
| Canada | 15 | 578 | 12.6 (-21.0, 28.9) | 67.3 (23.7, 95.8) | 8.3 (0.7, 20.3) | 207.8 (10.9, 368.1) | 60.4 (6.7, 65.1) |
| Chile | 4 | 8,052 | 11.9 (1.85, 23.7) | 74.1 (34.8, 96.7) | 7.9 (4.0, 13.2) | 169.5 (3.8, 357.6) | 68.7 (0.0, 78.4) |
| Czech Republic | 1 | 7,390 | 13.8 (-1.5, 25.7) | 65.4 (33.6, 93.6) | 8.2 (2.2, 14.8) | 176.7 (9.7, 316.0) | 52.1 (10.9, 80.2) |
| Estonia | 1 | 410 | 10.9 (-3.2, 22.5) | 75.9 (46.0, 96.2) | 8.1 (2.3, 14.6) | 165.0 (6.7, 336.1) | 40.8 (0.0, 63.5) |
| Finland | 1 | 915 | 10.9 (-2.6, 23.2) | 75.2 (45.9, 97.9) | 7.90 (2.2, 13.9) | 168.6 (6.4, 341.9) | 44.4 (16.2, 57.8) |
| France | 17 | 477 | 15.9 (0.5, 30.0) | 69.0 (24.1, 96.9) | 9.6 (2.4, 20.3) | 202.0 (11.2, 352.2) | 60.0 (12.0, 75.0) |
| Germany | 12 | 575 | 14.1 (-1.4, 29.2) | 66.6 (30.8, 97.5) | 8.4 (2.2, 16.0) | 177.1 (4.5, 338.7) | 56.2 (14.6, 69.8) |
| Italy | 23 | 2,013 | 18.6 (1.0, 30.8) | 67.5 (26.9, 97.7) | 11.21 (2.1, 22.0) | 222.9 (9.6, 345.0) | 68.4 (49.7, 81.0) |
| Japan | 10 | 163 | 19.8 (-5.9, 32.6) | 74.2 (34.2, 97.8) | 14.0 (2.4, 25.4) | 182.6 (11.3, 342.9) | 43.9 (16.2, 49.0) |
| Kuwait | 1 | 2,949 ⁷⁹ | 30.2 (7.8, 41.3) | 40.6 (18.6, 84.1) | 12.4 (2.6, 29.4) | 274.7 (114.8, 336.0) | 58.5 (5.2, 79.2) |
| Mexico | 8 | 1,367 | 20.2 (9.5, 31.1) | 59.5 (9.8, 96.9) | 10.5 (2.2, 20.3) | 269.6 (40.1, 371.4) | 51.9 (0.0, 62.5) |
| Peru | 18 | 3,489 80 | 15.0 (1.1, 30.1) | 67.0 (5.0, 96.3) | 9.8 (0.5, 24.1) | 236.7 (36.2, 383.8) | 76.3 (13.0, 81.8) |
| Romania | 8 | 1,006 81 | 18.5 (3.3, 30.1) | 63.7 (22.9, 97.8) | 10.4 (2.1, 19.2) | 211.1 (14.6, 338.2) | 52.0 (42.2, 71.4) |
| Singapore | 1 | 2,879 | 27.6 (26.0, 29.3) | 80.6 (71.2, 86.5) | 21.9 (20.0, 23.6) | 196.3 (49.1, 304.8) | 60.7 (31.7, 78.7) |
| South Africa | 1 | 1,998 82 | 15.1 (9.9, 19.8) | 78.6 (56.8, 95.2) | 10.30 (6.2, 13.5) | 172.7 (30.4, 350.1) | 68.9 (14.1, 80.2) |
| South Korea | 6 | 49 | 18.5 (-6.0, 29.8) | 73.3 (24.8, 98.0) | 13.0 (1.1, 24.7) | 185.1 (134.8, 332.1) | 53.3 (22.9, 72.9) |
| Spain | 52 | 6,210 | 17.9 (0.3, 34.2) | 63.9 (17.0, 97.3) | 9.9 (1.7, 22.1) | 232.2 (12.9, 368.6) | 56.8 (2.1, 72.9) |
| United Kingdom | 54 | 1,254 | 13.4 (2.3, 26.2) | 75.4 (41.9, 99.4) | 9.0 (3.7, 16.9) | 170.6 (5.9, 344.0) | 65.4 (8.3, 71.9) |
| United States | 209 | 8,350 | 19.4 (-14.1, 41.0) | 64.4 (5.8, 100.0) | 11.7 (0.7, 26.0) | 225.7 (7.1, 384.3) | 63.0 (8.3, 66.2) |

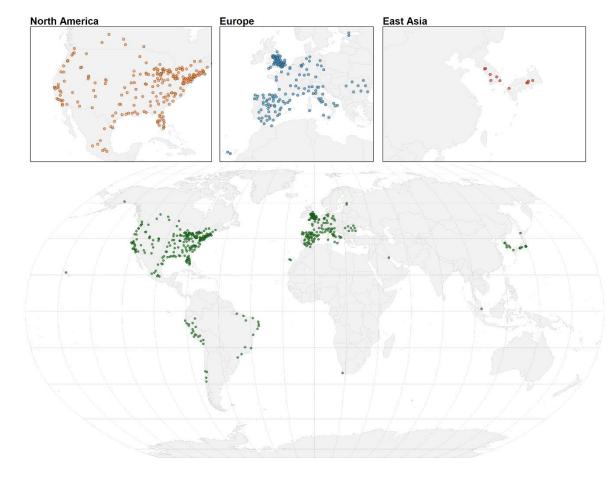


Figure 1. World map showing the included cities colour-coded by region.

Figure 2. Time-series of COVID-19 cases per 100.000 inhabitants aggregated by
country over the period form 3 February to 31 October 2020.

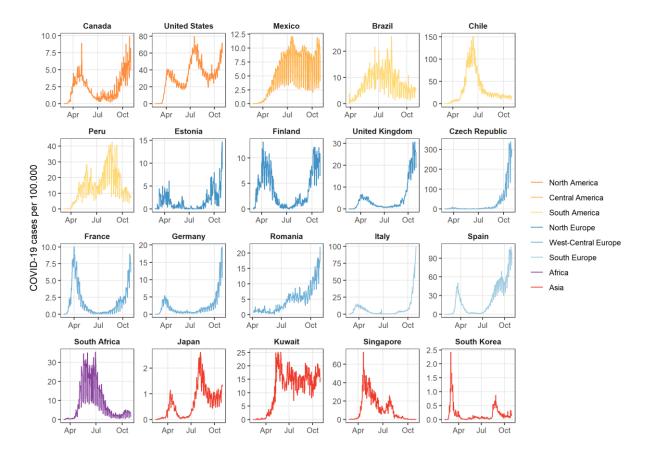


Figure 3: Association between meteorological variables and COVID-19 incidence. Association curves were obtained with meta-regression Model A with random effect defined by country and climatic zones.

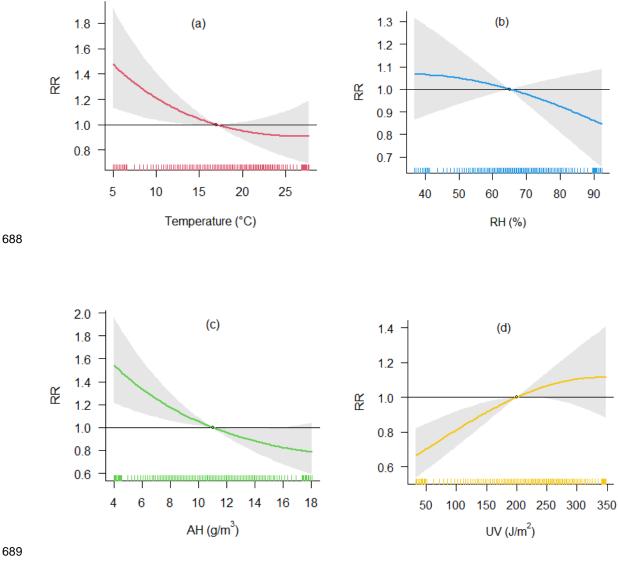
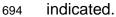


Figure 4. Country specific association between temperature and COVID-19 incidence. For each country the number (n) of cities included in the analysis is



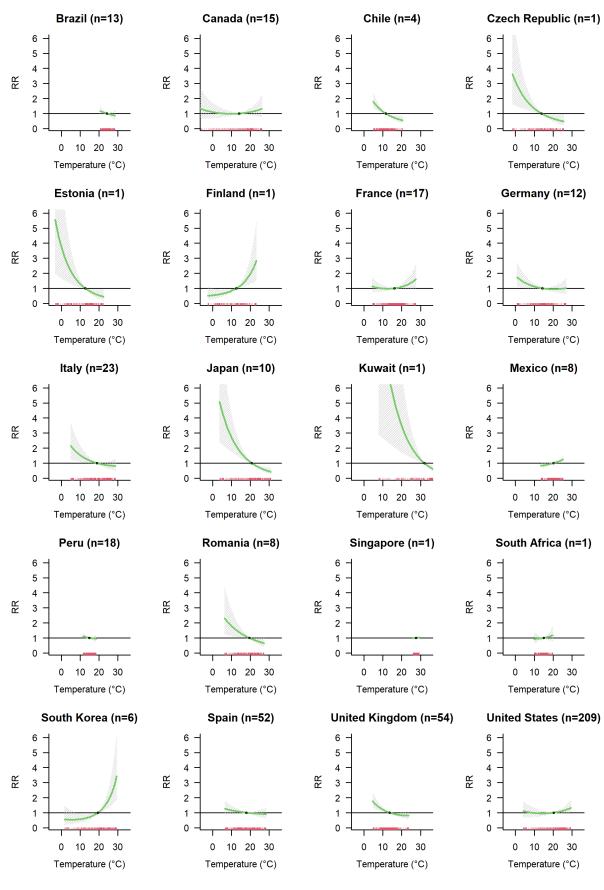


Figure 5. Country specific association between relative humidity and COVID-19 incidence. For each country the number (n) of cities included in the analysis is indicated.

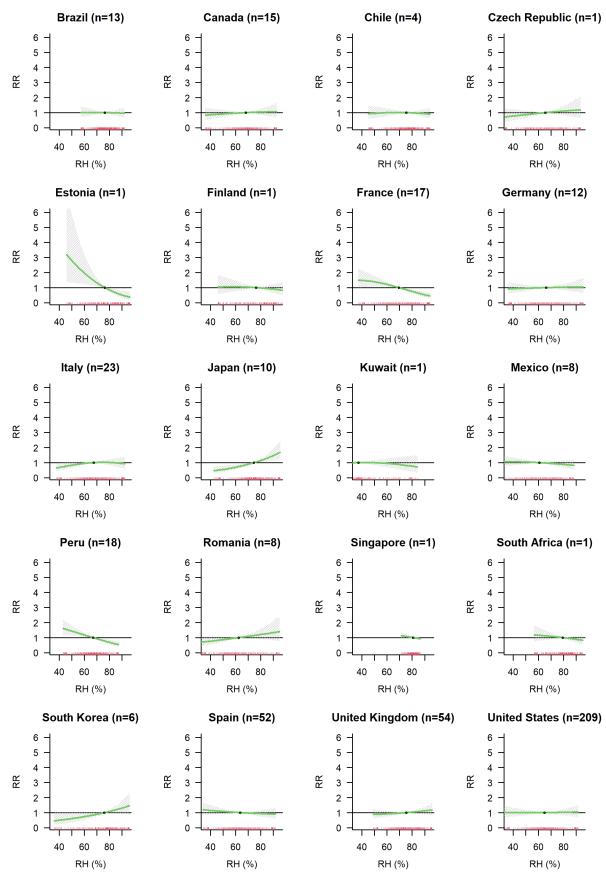


Figure 6. Country specific association between absolute humidity and COVID-19 incidence. For each country the number (n) of cities included in the analysis is

indicated.

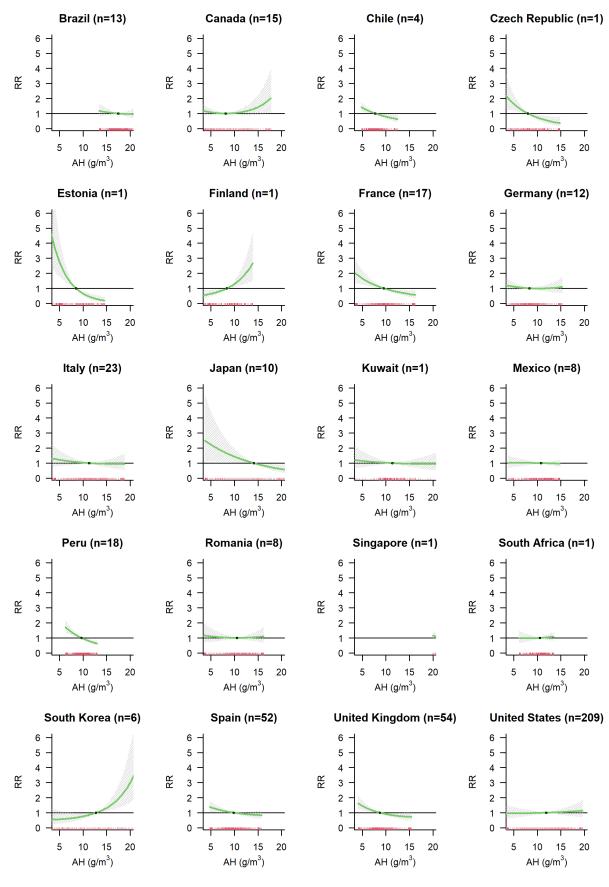


Figure 7. Country specific association between UV radiation and COVID-19

incidence. For each country the number (n) of cities included in the analysis is indicated.

