Prevalence of SARS-CoV-2 RNA on inanimate surfaces: a systematic review and meta-analysis

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Received: 25 February 2021 / Accepted: 29 June 2021 / Published online: 27 July 2021
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Abstract
Coronavirus disease (COVID-19) is a respiratory disease affecting many people and able to be transmitted through direct and perhaps indirect contact. Direct contact transmission, mediated by aerosols or droplets, is widely demonstrated, whereas indirect transmission is only supported by collateral evidence such as virus persistence on inanimate surfaces and data from other similar viruses. The present systematic review aims to estimate SARS-CoV-2 prevalence on inanimate surfaces, identifying risk levels according to surface characteristics. Data were obtained from studies in published papers collected from two databases (PubMed and Embase) with the last search on 1 September 2020. Included studies had to be papers in English, had to deal with coronavirus and had to consider inanimate surfaces in real settings. Studies were coded according to our assessment of the risk that the investigated surfaces could be contaminated by SARS-CoV-2. A meta-analysis and a metaregression were carried out to quantify virus RNA prevalence and to identify important factors driving differences among studies. Thirty-nine out of forty retrieved paper reported studies carried out in healthcare settings on the prevalence of virus RNA, five studies carry out also analyses through cell culture and six tested the viability of isolated viruses. Overall prevalences of SARS-CoV-2 RNA on high-, medium- and low-risk surfaces were 0.22 (CI95 [0.152–0.296]), 0.04 (CI95 [0.007–0.090]), and 0.00 (CI95 [0.00–0.019]), respectively. The duration surfaces were exposed to virus sources (patients) was the main factor explaining differences in prevalence.

Keywords Coronavirus · COVID-19 · Event rate · Epidemiology · Fomites · Risk · SARS-CoV-2

Introduction
Coronavirus disease 19 (COVID-19) is caused by the Severe Acute Respiratory Syndrome coronavirus type 2 (SARS-CoV-2). To date, the SARS-CoV-2 ongoing pandemic has caused more than 85 million cases in 191 countries, reaching around 2 million deaths worldwide by January 2021 [1].

COVID-19 case fatality rate was estimated to be 0.25% to 10% [2] with important differences among countries due to disparities in the number of people tested, demographics, characteristics of the healthcare system and other factors [3]. Symptoms caused by COVID-19 are variable and can include fever, cough, loss of sense of smell, and breathing difficulty. Most people are asymptomatic or mildly symptomatic, but some individuals can develop an acute respiratory distress syndrome, requiring medical support in intensive care units [4].

The main transmission route for SARS-CoV-2 to a susceptible individual appears to be direct contact with an infected subject who releases the virus into the air by sneezing, coughing, or speaking [5]. It has been hypothesized that SARS-CoV-2 could be also transmitted by indirect contact through fomites, defined as contaminated porous and non-porous surfaces or objects [6, 7]. For example, in many bacterial infectious diseases, the role of fomites is recognized as determinant in their transmission, especially in the nosocomial environment [7, 8]. Although this transmission route is well-known for some respiratory and enteric pathogens, its role in SARS-CoV-2 transmission is still largely unknown [9].
To establish SARS-CoV-2 ability to be transmitted through fomites, we need multiple examples of indirect evidence; otherwise, evaluation of this transmission route is hardly demonstrable. Moreover, we have to consider viral persistence in the environment, which not only depends on structural characteristics of the virus, for example, the presence of the envelope [10], but also on many environmental factors such as temperature, moisture, exposure to UV and surface characteristics [7]. The persistence of human and veterinary coronaviruses on different surfaces was recently reviewed by Marzoli et al. [11].

Since the beginning of the pandemic, besides the wearing of face masks, several public guidelines recommended cleaning and disinfection, use of gloves and hand hygiene as preventive measures, reverting to the precautionary principle due to the scarcity of available evidence [12]. Indeed, understanding the role of fomites in SARS-CoV-2 transmission is imperative, not only to define the appropriate measures of prevention, but also to improve their management under an ecological perspective. High levels of fomite transmission could make social distancing and wearing face masks insufficient actions to prevent the virus spreading among people. Alternatively, if fomite transmission were a negligible route of infection for this virus, we could try to limit the prodigious and widespread use of chemical disinfectant products or to reduce the amount of waste caused by unnecessary personal protective equipment (PPE), which is negatively affecting our environment [13].

In the present work, we collected, evaluated and discussed all existing data on the role of inanimate surfaces in the indirect transmission of SARS-CoV-2.

Materials and methods

We applied systematic review methodology using a specific, non-extensive approach in order to rapidly retrieve and screen relevant records from multiple databases.

Details of the systematic reviews carried out were the following:

**Review question** What is the prevalence of coronaviruses on surfaces in real settings?

**Population** surfaces (objects and environment).

**Outcome** prevalence of coronavirus RNA and/or infective viruses (Effect size: event rate).

We considered all papers published in peer-reviewed journals in English language. No time limits were imposed. We searched PUBMED and EMBASE (Title/Abstract and Title, Abstract, Author keywords, respectively) with the search terms reported in Table 1. The last date searched was 01/09/2020. To supplement the search process, we also used the final list of papers selected to carry out a backward reference search in order to identify potential missing evidence.

We applied several criteria to select eligible papers: (1) had to be written in English (2) had to report data belonging to primary research; (3) had to deal with coronavirus; (4) had to consider inanimate surfaces in real settings (not experimental contamination).

We carried out the screening process using EPPI-4 Reviewer software [14].

In the case of a poorly explicative abstract or in the case of doubt about the available data, the paper was included and evaluated at the full-text level. Thereafter, four reviewers (SB, FM, MF, AP) screened all papers (in duplicate) obtained via the initial literature search according to Title/Abstract and full text, independently (parallel method). Disagreements were resolved through consensus. For each relevant paper, one reviewer collected data and a second reviewer checked the collected data against the original paper (sequential method). We coded all papers according to the previously chosen parameters, and we recorded all data.

We collected data in pre-defined forms for a number of variables belonging to three categories (characteristics, design, and methodology) useful to describe the studies and to report results, together with relevant variables.

The methodology of included studies can have an influence on the results. To describe such possibilities, we collected information on the following parameters: analytical method/s, viability assessment, compliance statement to WHO guidelines concerning sampling [15] and/or detection [16], control of air contamination with droplets containing virus, PCR target genes, and size of the sampling area. According to WHO guidelines for sampling [15], a blank control swab should be included to assess the risk of aerosol/air contamination at the time of sampling. For this reason, air sampling was included as a relevant criterion.
Synthesis of results

We used event rate (prevalence) as effect size. We collected relevant data from primary studies, but we excluded data referring to surfaces that had been cleaned and/or disinfected immediately before sampling, and surfaces of clinical tools that were used inside COVID-19 patients (i.e. endotracheal tube). From each study on SARS-CoV-2, we extrapolated three parameters: the virus source (spreading subject or fomites), the location of surface where the sampling was performed and the duration of exposure. Briefly, the virus source, defined as the origin of viral contamination, distinguished among: direct contamination from a confirmed positive patient; direct contamination from a suspected positive patient; indirect contamination (this means without direct contact with a positive patient) and; an unknown viral source. The location of surface permitted us to evaluate the sampling site for each study and to group sampled objects and surfaces: in strict contact with the patient; in close proximity to the patient; in the healthcare environment and; in areas not related to the healthcare environment. Finally, we assigned, according to the available information, to each investigated surface, the possible duration of exposure to a viral source. Those parameters are described in detail in Table 2 and were subsequently scored from 1 to 4, as reported in Fig. 1.

For each study, we evaluated the parameters and assigned each one a relative score. The addition of those scores allowing us to synthesize the likelihood of the presence of SARS-CoV-2 RNA on a given surface as high, medium, low or very low. In metaregression, this scoring system was also used to investigate the influence of each parameter (used as moderator) on the risk of viral contamination.

We defined as a “study” an investigation performed on a group of surfaces which shared the same possibility of contamination. We defined as a “cluster” a group of studies belonging to the same paper. Papers reporting studies on coronavirus other than SARS-CoV-2 were not included in our categorization or in the meta-analyses; however, their results are discussed in the text.

Pooled prevalence

A cumulative variable “contamination risk” was defined according to Fig. 1. Meta-analyses and meta regressions were performed using the metafor package [17] of the statistical software R (version 3.6.0) [18]. Meta-analysis is a statistical method that combines outcomes of primary studies with a weight assigned according to the inverse of the variance. For this reason, the variance is a critical parameter, which has to be calculated when studies reporting zero prevalences are included. The Freeman-Tukey (double arcsine) transformation for proportion [19] was used to obtain a variance stabilizing transformation without applying continuity corrections or removing studies from the meta-analysis, and to appropriately weight studies with zero prevalence and high number of samples [17, 20]. We combined transformed prevalence estimates in meta-analysis using a multilevel random-effect model and later back-transformed in the original metrics [21]. We applied a multilevel structure to take into account the multiple estimated prevalences included in the same paper (cluster) referring to different studies (rma.mv function from metafor package; [17]).

We carried out several metaregression analyses to account for the applied study categorization and single factor scoring:

- a metaregression with “contamination risk” as moderator;
- a metaregression with virus source as moderator;
- a metaregression with location of surface as moderator;
- a metaregression with duration of exposure as moderator;
- a multivariate metaregression analysis including each of three risk factors (virus source, location of surface and duration of exposure) and their interactions as different moderators. The amount of heterogeneity was estimated using the Q, $T^2$ and $I^2$ [22, 23] statistics obtained by Restricted Maximum Likelihood (REML), which is considered approximately unbiased and relatively efficient [24]. In the multilevel model, the $T^2$ value for the total heterogeneity can be divided into two variance components, one for the between-cluster (papers) heterogeneity ($\sigma^2_1$) and one for the within-cluster (studies within paper) heterogeneity ($\sigma^2_2$).

Heterogeneity was explored through univariate and multivariate metaregression using the multilevel models [25, 26]. Moderator significance was assessed through the Likelihood Ratio Test (LRT) by comparing the proportional reduction for heterogeneity ($T^2$ value) of the full and reduced models.

The percentage of reduction in the total variance due to the moderator was evaluated using pseudo $R^2$ statistics. Maximum Likelihood (ML) estimate instead of REML was used to compare two nested models and to evaluate the importance of the moderators [17]. We made several attempts to explain heterogeneity through moderators describing the levels of exposure of different surfaces to viral contamination. In cases of moderator significance, determined according to the Likelihood Ratio Test, Tukey’s all-pairwise comparisons were performed using the False Discovery Rate correction [27] by glht function from multicomp package [28].
To account for the risk of bias within studies, we evaluated the influence of sampling procedure and analytical methods on prevalence, and we ran two additional meta-regressions with two moderators potentially linked with quality, according to data availability. The chosen factors were: the presence of a blank control or an alternative method to ascertain the absence of SARS-CoV-2 in air and; the use of two or more RNA targets for PCR. In the case of a significant moderator, we tested the interaction with the moderator “contamination risk”.

We took into consideration the risk of bias across studies with a mixed strategy. We applied a number of a priori exclusion criteria (eligibility criteria), and in addition, studies had to deal with SARS-CoV-2 (criterion for inclusion in meta-analysis). Additional factors potentially accounting for differences among studies were considered through the scoring system. The choice of surfaces that were sampled is the main difference in study design able to affect the results of each investigation, and thus, this could be an important source of bias. The risk of a surface becoming contaminated with virus genetic material depends on several factors. We considered as particularly relevant: the presence and the characteristics of the virus source; the proximity of the sampled surface to the viral source, and; the duration of exposure of the surface to the virus source. All these factors were considered when we scored and categorized the studies, as explained above, and were included in the metaregression to explain heterogeneity. Details about the criteria we used to assign different scores within moderators are reported in Table 2. Researchers (FB, AB, and EM) with expertise in virology and direct knowledge of SARS-CoV-2 experimental behavior assessed the risk of bias issue. In particular, we addressed this issue by categorizing the studies according to the scheme described above. Note that although we chose factors that we considered useful, this is inevitably a subjective procedure and alternative categorizations could be equally valid.

In addition, to account for bias, we ran sensitivity analyses in each model in order to evaluate the presence of outliers or leverage studies and their potential influence on each model. Four parameters were examined: the externally studentized residuals; the DFBETAS values; the Cook’s distance; and
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the hat function. Influence was defined according to meta-for package criteria (DFBETAS value is larger than 1, the lower tail area of a chi-square distribution with p degrees of freedom cut-off by the Cook’s distance is larger than 50% OR hat value > 3(p/k)). Furthermore, studies were excluded one by one from the model to evaluate relevant changes in heterogeneity (T² and Q) and pooled estimates. Finally, publication bias was evaluated through Egger’s regression test, obtained by including the standard error or the variance of the effect sizes as a moderator in the model [29]. The significance of the moderator suggests the presence of publication bias. A p-value < 0.05 (p) was considered as significant in the statistical analysis.

Results

Study selection

After literature searches in the selected databases, we retrieved 2105 records, 1142 remained after duplicate removal and finally, after title/abstract and full text screening, 40 papers were selected for inclusion (Fig. 2).

Study characteristics

After the screening process, we included 40 papers reporting data about coronaviruses’ presence on surfaces. Thirty-two papers (all published in 2020) investigated the presence of SARS-CoV-2 RNA and reported data for 88 studies, whereas eight papers (all published before 2020) investigated other coronaviruses. Out of 40 papers, 23 were full articles, while the others were short communications, letters, dispatches or other types of publications. Twenty-two papers described studies carried out in Southeast Asia. Twenty-nine out of 32 studies were fully carried out in healthcare facilities where a known source of contamination (infected patients) was present. Emergency departments, isolation rooms and intensive care units were the most frequently investigated areas. Six studies considered high contact surfaces in the public environment, although four of them dealt with common human coronaviruses (HCoVs) (Table 3). A wide number of different surfaces was investigated, but the surface material was not reported (with only one exception), and thus, it was impossible for us to investigate the role of surface materials on virus presence. Full details on the 40 papers’ characteristics are reported in "Appendix" Table 6.

Risk of bias within studies (quality evaluation)

The parameters influencing the papers’ characteristics are summarized in Table 4. We observed high variability regarding the sampling procedure; in particular, the swabbed area ranged from 9 cm² to 100 cm², with most studies not reporting this information. Some studies referred to the WHO guidelines (published in February 2020). WHO guidelines recommend swabbing a 25 cm² surface area, and to put the swab in 1–3 ml of Viral Transport Medium. Multiple sampling for each surface is suggested, as is the inclusion of a blank control swab (to determine whether aerosol/air contamination at the time of sampling could interfere with study results) [15]. In our current analysis, the surfaces selected for sampling purposes by the different studies were highly varied, but most of them were listed as suggested surfaces in WHO guidelines [15]. It should be emphasized that most studies,
according to the WHO suggestion, investigated mainly high contact surfaces, with a higher risk of contamination, and not randomly chosen surfaces. All studies except one carried out PCR analysis on swabs. Five studies (four targeting SARS-CoV-2) also carried out virus viability assays in cell culture for PCR positive samples.

As regards the risk of bias assessment, the moderators tested were not significant. In particular, both, the presence of a blank control or an alternative method to ascertain the absence of SARS-CoV-2 in air and the use of two or more RNA target for PCR were non significant with p-values of, respectively, 0.1665 and 0.2636.

**Results of individual studies**

We observed high variability in the prevalence of RNA contamination on surfaces as reported by individual studies. Study characteristics could, in part, explain such differences and were taken into account in our synthesis of results. The results of individual studies dealing with SARS-CoV-2 are summarized in Fig. 3.

With regards to the eight papers dealing with coronaviruses other than SARS-CoV-2, four studies investigated the presence of H-CoV. Out of these, two studies evaluated the frequency of H-CoV contamination on airport surfaces and reported prevalences of 3.3% [70] and 7.5% [51]. One study investigating the frequency of contamination by respiratory pathogens of areas and surfaces in places associated with worship reported an overall HCoV prevalence of 1.4% [42], whereas in a university classroom, desktops and the door-knob were the surfaces most commonly contaminated with H-CoV [34]. The prevalence of SARS-CoV contamination
Table 4 methodology characteristics of the 40 included papers: type of diagnostic test, viability assessment (conducted on PCR positive samples), compliance statement to WHO guidelines, control of air contamination, number of PCR target detected per virus, sampled area. Dashes are reported only for studies to which it was not possible to assign a dot in the characteristics column.

| References     | Detected virus | Diagnostic test | Viability assessment | WHO guidelines | Air control sample | PCR target | Area (cm²) |
|----------------|----------------|-----------------|---------------------|----------------|--------------------|------------|------------|
|                |                | PCR             | Cell culture        | Sampling       | Detection          | Swab       | Other method | l ≥ 2     | NR       |
| Ahn et al. [30]| SARS-CoV-2     | •               | •                   | •              | •                  | •          | •          | –         |          |
| Aytogan et al. [31]| SARS-CoV-2 | •               | –                   | •              | •                  | •          | •          | –         |          |
| Bin et al. [32]| MERS-CoV       | •               | •                   | •              | •                  | •          | •          | –         |          |
| Bloise et al. [33]| SARS-CoV-2 | •               | –                   | •              | •                  | •          | •          | –         |          |
| Bonny et al. [34]| HCoV          | •               | –                   | •              | •                  | •          | •          | –         |          |
| Booth et al. [35]| SARS-CoV       | •               | •                   | •              | •                  | •          | •          | –         |          |
| Cheng et al. [36]| SARS-CoV-2   | •               | •                   | •              | •                  | •          | •          | 9/WS      |          |
| Chia et al. [37]| SARS-CoV-2     | •               | –                   | •              | •                  | •          | •          | –         |          |
| Colaneri et al. [38]| SARS-CoV-2 | •               | •                   | •              | •                  | •          | •          | –         |          |
| Colaneri et al. [39]| SARS-CoV-2 | •               | •                   | •              | •                  | •          | •          | –         |          |
| Dowell et al. [40]| SARS-CoV      | •               | –                   | •              | •                  | •          | •          | 20        |          |
| Guo et al. [41]| SARS-CoV-2     | •               | –                   | •              | •                  | •          | •          | –         |          |
| Hoang et al. [42]| HCoV          | •               | –                   | •              | •                  | •          | •          | 25        |          |
| Ikonen et al. [43]| HCoV        | •               | –                   | •              | •                  | •          | •          | 100       |          |
| Jerry et al. [44]| SARS-CoV-2     | •               | •                   | •              | •                  | •          | •          | –         |          |
| Jiang et al. [45]| SARS-CoV-2     | •               | –                   | •              | •                  | •          | •          | –         |          |
| Jiang et al. [46]| SARS-CoV-2     | •               | –                   | •              | •                  | •          | •          | –         |          |
| Kim et al. [47]| MERS-CoV       | •               | –                   | •              | •                  | •          | •          | –         |          |
| Lee et al. [48]| SARS-CoV-2     | •               | –                   | •              | •                  | •          | •          | 700       |          |
| Lei et al. [49]| SARS-CoV-2     | •               | –                   | •              | •                  | •          | •          | –         |          |
| Lv et al. [50]| SARS-CoV-2     | •               | •                   | •              | •                  | •          | •          | 25-100    |          |
| Memish et al. [51]| HCoV        | •               | –                   | •              | •                  | •          | •          | 25        |          |
| Mouchtouri et al. [52]| SARS-CoV-2 | •               | –                   | •              | •                  | •          | •          | 25        |          |
| Nelson et al. [53]| SARS-CoV-2   | •               | –                   | •              | •                  | •          | •          | 25        |          |
| Ong et al. [54]| SARS-CoV-2     | •               | –                   | •              | •                  | •          | •          | –         |          |
| Pasquarella et al. [55]| SARS-CoV-2 | •               | –                   | •              | •                  | •          | •          | –         |          |
| Peyrony et al. [56]| SARS-CoV-2 | •               | •                   | •              | •                  | •          | •          | 25-50     |          |
| Razzini et al. [57]| SARS-CoV-2 | •               | •                   | •              | •                  | •          | •          | –         |          |
| Ryu et al. [58]| SARS-CoV-2     | •               | •                   | •              | •                  | •          | •          | –         |          |
| Santarpia et al. [59]| SARS-CoV-2 | •               | •                   | •              | •                  | •          | •          | –         |          |
| Shin et al. [60]| SARS-CoV-2     | •               | –                   | •              | •                  | •          | •          | –         |          |
| Tan et al. [61]| SARS-CoV-2     | •               | –                   | •              | •                  | •          | •          | –         |          |
| Wang et al. [62]| SARS-CoV-2     | •               | –                   | •              | •                  | •          | •          | ≤ 100     |          |
| Wang et al. [63]| SARS-CoV-2     | •               | –                   | •              | •                  | •          | •          | –         |          |
was investigated in two studies on hospital surfaces, and both revealed the high frequency of SARS-CoV contamination on surfaces exposed to SARS-CoV positive patients [35, 71]. The frequency of MERS-CoV contamination was investigated on hospital surfaces, and the frequency of contamination on the sampled surfaces was over 20% in both the retrieved studies [32, 47].

Only 11 studies reported data about the viability of coronaviruses. In particular, six studies were conducted on SARS-CoV-2 [30, 36, 38, 59, 66, 69], two on SARS-CoV [35, 40], two on MERS-CoV [32, 47], and one on H-CoV [34]. Among them, five studies conducted the viability assessment only on RT-PCR positive samples, while six studies performed cell culture analysis on all the sampled surfaces (Table 4). Only three studies detected viable SARS-CoV-2 on tested surfaces [30, 36, 59]; prevalences were 8.2% [30] and 5% [36] of total sampled surfaces. Viability was also reported for MERS-CoV on 4.3% [32] and 22.1% [47] of total sampled surfaces, and for HCoV on 37.5% of total sampled surfaces [34].

Synthesis of results

We included 88 studies dealing with SARS-CoV-2 in the metaregression. Across all studies overall, SARS-CoV-2 RNA was detected on surfaces with a prevalence of 0.085 (CI95 [0.042–0.138]). When we assigned surfaces into different risk categories (high, medium and low), SARS-CoV-2 RNA was detected with a prevalence of 0.22 (CI95 [0.152–0.296]), 0.04 (CI95 [0.007–0.090]) and 0.00 (CI95 [0.00–0.019]), respectively.

The moderator “contamination risk” was significant (p < 0.0001), and the proportional reduction in the total variance was R² = 43%. Multiple pairwise comparison of means (Tukey contrasts) resulted in a statistically significant difference of studies investigating high-risk surfaces vs studies investigating medium-risk surfaces (p = 2.56 e-06) and of studies investigating high-risk surfaces vs studies investigating low-risk (p = 2.91 e-10) surfaces. Also the comparison between medium- and low-risk surfaces was significant (p = 0.011) (Fig. 3).

To account for the importance of different factors used to assign the category “contamination risk”, the scores of each factor (virus source, location of surface and duration of exposure) were also taken into account separately.

The three moderators, taken individually, were all statistically significant as reported in Table 5. Heterogeneity was always significant according to Q statistics and around 90% when estimated as I². The multiple comparison of means was run in each model between all paired variables. In the model considering virus source, only the comparison
Fig. 3  forest plot with individual study results and result of metaregression with “contamination risk” as moderator
between “direct contact-patients confirmed positive” and “indirect contact-fomites” showed a significant difference ($p = 0.035$). In the model with location of surface as the moderator, four pairwise comparisons resulted in significant differences: “objects and surfaces in proximity with patient/s” vs “healthcare surfaces and areas” ($p = 2.67 \times 10^{-4}$), “objects in strict contact with patient/s” vs “healthcare surfaces and areas” ($p = 4.07 \times 10^{-4}$), “objects and surfaces in proximity with patient/s” vs “non-healthcare related areas” ($p = 0.046$) and “objects in strict contact with patient/s” vs “non-healthcare related areas” ($p = 0.027$).

In the case of duration of exposure, the significant pairwise comparisons were: “less than 20 min or unspecified” vs “1 to 12 h” ($p = 5.36 \times 10^{-5}$), “up to 1 h post disinfection” vs “1 to 12 h” ($p = 2.90 \times 10^{-5}$), “more than 12 h” vs “less than 20 min or unspecified” ($p = 1.26 \times 10^{-4}$) and “up to 1 h post disinfection” vs “more than 12 h” ($p = 9.65 \times 10^{-4}$).

Even when all moderators were included in the multivariate model, heterogeneity was high, with $I^2 = 87\%$. According to this model, about 53% of the total variance was due to between-cluster differences, with the remaining 34% due to within-cluster differences. The remaining 13% was due to sampling variance. The proportional reduction in the total variance was $R^2 = 44\%$. The estimated prevalence rates for each score are shown in Fig. 4.

In the multiple comparison test, only differences due to the moderator “duration of exposure” were significant: “less than 20 min or unspecified” vs “1 to 12 h” ($p = 0.003$), “up to 1 h post disinfection” vs “1 to 12 h” ($p = 4.4 \times 10^{-5}$), “more than 12 h” vs “less than 20 min or unspecified” ($p = 8.13 \times 10^{-4}$) and “up to 1 h post disinfection” vs “more than 12 h” ($p = 8.82 \times 10^{-4}$).

In this last model, interactions among moderators were also evaluated, but they were not significant.

Sensitivity analysis was carried out in each model according to the definition of influence previously described. In the model without moderators, two studies [54, 59] were influential. In the model with “contamination risk” as the moderator, one study was influential [59]. We ran the model excluding the studies that proved to be influential, and in both cases, we obtained an overall prevalence of 0.073 (CI95 [0.036–0.118]). In the model with moderators of “low” and “medium” contamination risk, small changes (0.000 CI95 [0.000; 0.013] and 0.036 CI95 [0.006; 0.080], respectively) in prevalence occurred, whereas in the “high” risk category, prevalence decreased to 0.194 CI95 [0.133; 0.262]. In the models with the other moderators, the analysis of influence was affected by the non-homogeneous number of studies in different groups with the same scores. Full results of the sensitivity analysis are reported in Supplementary materials (Figures 1 to 5).

### Table 5 results of the three models including a single moderator each

| Virus source | Q $p < 0.0001$ | $T^2 = 0.053$ | $I^2 = 91\%$ | Mod $p = 0.02$ | $R^2 = 14\%$ |
|--------------|----------------|--------------|-------------|---------------|-------------|
| Patients confirmed positive | 0.130 CI95 [0.072–0.197] | | | | |
| Patients suspected positive | 0.061 CI95 [0.00–0.220] | | | | |
| Indirect contact–fomites | 0.023 CI95 [0.00–0.089] | | | | |
| Unknown or SARS-CoV-2 negative | 0.000 CI95 [0.00–0.118] | | | | |
| Location of surface | Q $p < 0.0001$ | $T^2 = 0.049$ | $I^2 = 90\%$ | Mod $p = 0.0002$ | $R^2 = 20\%$ |
| Objects in strict contact with patient/s | 0.177 CI95 [0.084–0.289] | | | | |
| Objects and surfaces in proximity with patient/s | 0.132 CI95 [0.069–0.206] | | | | |
| Healthcare related areas | 0.005 CI95 [0.00–0.049] | | | | |
| Non healthcare related areas | 0.000 CI95 [0.00–0.110] | | | | |
| Duration of exposure | Q $p < 0.0001$ | $T^2 = 0.040$ | $I^2 = 89\%$ | Mod $p < 0.0001$ | $R^2 = 36\%$ |
| More than 12 h | 0.428 CI95 [0.158–0.721] | | | | |
| Up to 12 h | 0.164 CI95 [0.104–0.234] | | | | |
| Up to 1 h | 0.017 CI95 [0.000–0.059] | | | | |
| Unknown/less than 20 min | 0.000 CI95 [0.000–0.020] | | | | |
In the model without moderators, the null-hypothesis of no relationship between the primary study’s effect size and standard error was not rejected by Egger’s test ($p = 0.106$). This implies that there is no evidence for small-study effects in the meta-analysis [72]. In contrast, Egger’s test identified significant asymmetry in the funnel plots of all models with moderators ($p < 0.01$; Figures 6 to 9 in the Supplementary material).

**Discussion**

Since the beginning of the current pandemic, the unknown role of indirect transmission has resulted in restrictive preventive measures such as cleaning, disinfection, glove use and hand hygiene in addition to use of face masks [12]. Moreover, there is scarcity of data about the frequency of contamination of different surfaces that could drive any refinement of preventive measures.

**Summary of evidence**

SARS-CoV-2 RNA was detected on surfaces with an overall prevalence of 8.5%. Most studies ("Appendix" Table 6) were carried out in healthcare settings, and often in emergency department units, isolation rooms or intensive care units, with exposure to a direct or indirect virus source. In addition, surfaces were often chosen according to the extent of contact with patients. Following these considerations, in these areas, the prevalences detected are relatively low, possibly because of the disinfection protocols applied in such settings, as supported by the importance of the moderator “duration of exposure”.

The moderator “contamination risk” built up from the score deriving from the categorization system was significant, and was able to summarize different risk factors potentially explaining surface contamination. The prevalences of SARS-CoV-2 RNA on low-risk surfaces were between 0 and 1.9%. If we consider non-health care areas, such as public
areas or the outside environment, we suggest the risk of a surface being contaminated is very low.

Duration of exposure greatly influenced the prevalence rate, whereas virus source and location of surface were significant in the single moderator models, losing significance in the multivariate model accounting for all moderators.

The moderator “virus source” was aimed at demonstrating the influence of a virus source, namely an infected individual actively shedding the respiratory pathogen, on surface contamination. The absence of such significance could be due to several factors: the presence of unrecognized or not reported infected individuals in the surrounding areas, the wide circulation of virus through indirect contact (i.e. fomites or air ventilation systems) and; the high variability in some categories, as for example, in direct contact-patients suspected positive, where SARS-CoV-2 RNA prevalences ranged between 0 and 25.8%. This last category accounted for a lack of information in the primary research papers that were included in our study. Another reason for lack of significant results could be due to disinfection procedures that keep the level of surface contamination below certain levels, reducing our ability to highlight differences. As a matter of fact, the only pairwise difference observed in the univariate model was lost in the multivariate model.

The moderator “location of surface” was aimed at identifying a risk of surface contamination due to proximity to the virus source. The starting point was that surfaces in the same room as a spreading patient and surfaces in the same room and in close proximity to a patient, could be contaminated with higher frequency since virus-laden respiratory droplets tend to settle and contaminate surfaces within a 1.8 m distance from the virus source [73]. The failure to confirm proximity to a virus source was associated with SARS-CoV-2 RNA surface contamination could also be due to the disinfection procedures that keep the level of contamination below certain levels, deterring our analysis from highlighting differences. For this moderator too, some pairwise differences observed in the univariate model were lost in the multivariate one.

The moderator “duration of exposure” was significant in both univariate and multivariate models, suggesting that reduced duration of exposure of a surface to a virus source is the main factor explaining the variability of prevalence rates on surfaces and in healthcare settings. A reduced duration of exposure could be due to the presence of the virus source for a short time only or to frequently applied disinfection procedures, as occurs in healthcare settings.

If virus RNA is detectable around patients, on surfaces, and in the surrounding air, this likely relates only poorly to the possibility of transmitting the disease, since the presence of virus RNA does not hold or imply any information on virus viability. Among the retrieved studies, only six assessed SARS-CoV-2 viability and only three detected viable virus [30, 36, 59]. It is important to stress that in those studies that detected viable SARS-CoV-2, the analysed surfaces were all classified as high-risk. Only one study analyzing high-risk surfaces did not detect viable SARS-CoV-2 on any surfaces [69]. Studies analyzing low- and medium-risk surfaces did not report any viable SARS-CoV-2, despite the fact that some surfaces were RT-PCR positive [38, 66, 69].

Given the low number of studies testing virus viability, we cannot correlate the RNA prevalence with the possibility of fomite transmission. However, the potential for a virus to survive and to be transmitted via fomites also depends on virus type and characteristics. For example, the presence of viral envelope, as in SARS-CoV-2, is associated with relatively low virus resistance on surfaces [10]. Although prolonged survival of SARS-CoV-2 on surfaces has been proven (until 28 days), evidence of transmission from contaminated dry surfaces in real settings is still lacking, and direct person-to-person transmission remains the main route [11].

If droplets are inhaled, the minimal infectious dose must be reached [74]. To date, a minimal infectious dose has not been defined for SARS-CoV-2, although it is suspected to be as low as 50 particles [75].

Finally, we should also consider the efficiency of transmission. Data for bacteriophages showed the transmission efficiency ranges from <0.01% to 80% for fomite to hand
transmission, with smooth surfaces showing the highest efficiency, and around 33.90% for hand to lip transmission [76]. No data on the efficiency of transmission for SARS-CoV-2 are available at the present date.

Limitations

The present work has some limitations. Data used to estimate prevalences were based on RNA presence and neither quantitative information about genomic copies nor information about the presence of viable virus particles were available. Few papers assessed the viability of virus in positive samples, and only two tested all the samples with cell culture. Thus, the reported prevalence rates have a risk of overestimation as, in some cases, the virus RNA detected could be non-infective. Studies assessing virus viability failed in most cases to demonstrate it. However, previous investigations suggested that virus culturability is possible only with virus loads greater than 10–100 TCID₅₀/ml [77–79].

Another limitation is linked to the choice of samples within each study. Sampled surfaces were in most cases chosen based on risk of contact (i.e. on the risk the surfaces could be contaminated or could be able to transfer contamination). This is another characteristic of the included studies that could have led to an overestimation of real prevalence. To account for this, we categorized surfaces according to variables that were likely to increase the risk the surfaces could be contaminated. However, our categorization has some limitations due to the difficulty, in some cases, of us categorizing the studies without introducing a risk of subjectivity.

Each category has specific limitations. We used virus source to take into account patient status, but were unable to account for disease stage and amount of virus shed by patients. This is partly because not all studies reported these data and partly because in some cases it was not possible to directly pair the virus source with the surface. The location of surface was classified according to the descriptions in the included studies, but, especially for objects in the same room, a risk of misclassification exists. Moreover, any analysis of duration of exposure suffers from potential poor reporting of cleaning and disinfection procedures.

Conclusions

The risk a surface will be contaminated with SARS-CoV-2 RNA depends mainly on the duration of exposure to a virus source. Irrespective of the amount of viable virus, potential measures to reduce contamination need to include timely and appropriate cleaning and disinfection protocols for at-risk surfaces. The risk of surface contamination is of particular importance in healthcare settings. According to our scoring system, the risk can be considered very low where exposure to a virus source is limited in duration, as in most public areas, and high in areas where there is prolonged exposure to infected patients. Data for coronaviruses other than SARS-CoV-2 show that RNA from these viruses can sometimes be found in public areas, but no data specific for SARS-CoV-2 were retrieved. These results suggest adoption of a cautious approach as regards the widespread use of disinfectants in low-risk areas. This is due to the potential contribution of disinfectants to the increase of antimicrobial resistance [80, 81], which is a global challenge for human and animal health.

Appendix

See Table 6.
| References | Type of publication | Country | Virus | Setting | Place | Investigated surfaces |
|------------|---------------------|---------|-------|---------|-------|-----------------------|
| 1 Ahn et al. [30] | Article | South Korea | SARS-CoV-2 | HCF | Isolation rooms | Tables, blood pressure cuffs, pillows, bed sheets, nasal prongs, outside surface of the ventilator circuit, tubing, masks, telephones, thermometers, keyboards, and fixed structures in the room (such as the doorknob, bed rails, floor, walls, window, and faucet handles), and grills of the ventilation exits |
| 2 Aytogan et al. [31] | BR | Turkey | SARS-CoV-2 | HCF | Ophthalmology examination room | Bio-microscope stage, slitlamp shield, phoropter, tonometer, desk, and door handles |
| 3 Bin et al. [32] | Article | South Korea | MERS-CoV | HCF | Isolation rooms | Bedrails, Bed tables, Bed controllers, Shelves, Door buttons, Bathroom door knobs, Patient room floor, Patient monitor buttons, Thermometers, IV fluid hangers, Portable X-rays,Computed radiography cassette, Anteroom floors, Anteroom tables, Entrances of air- ventilating equipment |
| 4 Bloise et al. [33] | Letter | Spain | SARS-CoV-2 | HCF | Lab. for SARS-CoV-2 diagnosis | Landline, Barcode scanner, Mobile phone, Mouse, Keyboard, equipment |
| 5 Bonny et al. [34] | BR | USA | CoV-229E | Public areas | University Classrooms | Seat-backs made of hard polyvinylchloride, (2) laminate desktops, (3) a wooden podium, and (4) a stainless steel doorknob |
| References | Type of publication | Country | Virus | Setting | Place | Investigated surfaces |
|------------|---------------------|---------|-------|---------|-------|-----------------------|
| 6 Booth et al. [35] | Article | Canada | SARS-CoV | HCF | Room/wards | (Hand-rails, call buttons, telephones, televisions, television remote controls, light switches, carts, and bed tables) in patient rooms; bathrooms (soap dispensers, faucet handles, safety bars, toilet handles, and toilet seats); ventilation-system components (air vents); and specific patient care equipment (blood pressure cuffs and oxygen-administration equipment). Hallways adjacent to patient rooms, personal protective equipment carts and hand sanitizer stations, and at nurses’ stations. Other potentially contaminated areas and “control” (i.e., non-SARS) areas were also included |
| 7 Cheng et al. [36] | Article | China | SARS-CoV-2 | HCF | Isolation rooms | Mobile phone, bed rails, toilet door handle, bed table, locker |
| 8 Chia et al. [37] | Communication | Singapore | SARS-CoV-2 | HCF | Rooms (ICU) and Isolation wards | Floor, bed rail, locker handle, cardiac table, electric switch, chair, toilet seat and flush, air exhaust vent |
| 9 Colaneri et al. [38] | Research note | Italy | SARS-CoV-2 | HCF | Infectious Disease Emergency Unit and subintensive care ward | Exit buffer zone, Computer keyboards, Bed rails, Infusion pump, Multi-parameter monitor, Nurse buzzer, CPAP helmet, flow oxygen therapy, Table, Staff personal protective, Liquid-repelling gowns after 1 h of use, Face shield and eye goggles, Staff gloves, computer keyboards, Telephones, door knobs and water taps in patient toilets, Staff equipment, Portable X-ray machine, Electrocardiograph machine, Medication cart, Staff mobile phone |
| References | Type of publication | Country          | Virus         | Setting | Place                                                                 | Investigated surfaces                                                                 |
|------------|---------------------|------------------|---------------|---------|----------------------------------------------------------------------|----------------------------------------------------------------------------------------|
| 10 Colaneri et al. [39] | Practice point | Italy          | SARS-CoV-2   | HCF     | Buffer zone of patients rooms, doctors' and nurses' lounge, staff personnel belonging | Door handles, Waste container covers, Sink handles, Wall surfaces, Kitchen table and sink Desks, Computer keyboards, Medical charts and parameters Tabs, Door handles, Therapy trolleys, Mobile phones |
| 11 Dowell et al. [40] | Article | Thailand and Taiwan | SARS-CoV   | HCF     | Patients' room, nursing ward, emergency dept. and public area                                                                 | Nursing station (Sterile cotton swab, Air in nursing station, Telephone, Garbage can, Outside doorknob, Patient room (Doorknob, Cardiopulmonary resuscitation tray, Intravenous imipenem bag, Ventilator control panel, Intravenous line pole, Defibrillator paddle, Bed sheet, Chest tube site, Endotracheal tube, Nasopharynx, Skin hand, Conjunctivae) |
| 12 Guo et al. [41] | Dispatch | China          | SARS-CoV-2   | HCF     | Intensive care unit (ICU) and a general COVID-19 ward (GW) at Huoshenshan Hospital in Wuhan, China | Contaminated area, semicontaminated area, clean area |
| 13 Hoang et al., [42] | Article | Saudi Arabia  | H-CoV        | Public areas | Pilgrimage crowded place during 2016 and 2018 Haji in Medina, Mecca, Arafat, Mina | High contact surfaces in buses, hotels, commercial store, tents, restaurants |
| 14 Ikonen et al. [43] | Article | Finland        | HCoVOC43     | Public areas | Airport Helsinki-Vantaa | High contact surfaces at the airport (Toilet surfaces, luggage, handrails, plastic toy, elevator button, desk, button of payment terminal) |
| 151 Jerry et al. [44] | Article | Ireland        | SARS-CoV-2   | HCF     | Emergency dept, intensive care unit, high-dependency unit and wards | COVID-19 patient room and patient room post terminal clean (Bed rail, bedside table, call bell, patient chair-arm, remote for bed, toilet door handle); Nurses ward (desk, keyboard, telephone) |
| 61 Jiang et al. [45] | Dispatch | China          | SARS-CoV-2   | HCF     | Quarantine hotel rooms of presymptomatic patients | Door handle, light switch, faucet handle, thermometer, television remote, pillow cover, duvet cover, sheet, towel, bathroom door handle, and toilet seat and flushing button |
| References  | Type of publication | Country       | Virus          | Setting          | Place                                 | Investigated surfaces                                                                 |
|-------------|---------------------|---------------|----------------|------------------|---------------------------------------|----------------------------------------------------------------------------------------|
| 71 Jiang et al. [46] | Letter   | China     | SARS-CoV-2 | HCF             | Fomites in patient's room, PPE, nurses carts | Goggles, masks, nurses carts, nurses tables, mobile phones, water cups |
| 81 Kim et al. [47] | Article   | South Korea | MERS-CoV    | HCF       | Hospital A and B in Korea            | Fomites as stethoscopes, bag valve masks, blood pressure cuffs, nasal prongs, pillows and keyboards; fixed-structure in the elevator as buttons, guardrails, doors and in the rooms and restrooms as doorknobs, bed guardrails, toilet seats, hand soap dispenser; ventilator exits |
| 9 Lee et al. [48]    | SC        | South Korea | SARS-CoV-2 | HCF & mass facilities | Highly touched surfaces in mass facilities | Telephones, beds, chairs, door handle in wards, communal facilities |
| 20 Lei et al. [49]   | Article   | China      | SARS-CoV-2  | HCF             | ICU and isolation wards surfaces and fomites | Patient specific areas (floor, clothes, bedsheet, control panel of the ventilator, ventilator outlet valve). Other areas (changing room door handle, floor of the changing room, faucets at the handwashing station, keyboard of shared computer) |
| 21 Lv et al. [50]    | Article   | China      | SARS-CoV-2  | HCF             | Surfaces in a COVID lab of a grade A hospital in China | Sample transportation and reception related facilities (door handle, elevator buttons, handle and inner wall of sample transport boxes and of transfer window), testing related instruments (BSC button, door handle, countertop, wall, pipette, centrifuge, refrigerator, autoclave, eye washer, faucet), PPE (outer and inner gloves, goggles, safety shoes, protective clothing and mask, surgical hats), other facilities (floor, table, chair in contaminated and semi-contaminated area) |
| 22 Memish et al. [51] | Article  | Saudi Arabia | H-CoV OC43/HKU1 | Public areas | Hajj terminal of King Abdul Aziz International Airport, Saudi Arabia | Passport inspection table, chair handles, bathroom handles, check in counter table |
| References | Type of publication | Country | Virus | Setting | Place | Investigated surfaces |
|------------|---------------------|---------|-------|---------|-------|-----------------------|
| Mouchtouri et al. [52] | SC | Greece | SARS-CoV-2 | HCF, mass facilities and public area | Ferry boat, nursing home, isolation facility for asymptomatic people, covid-hospital Isolation room | Ferry boat: ship hospital and cabins, air duct, doorknobs in toilets, passenger bar counter, light switch, handrail, dishes, refrigerator door handle; in nursing home: bed side rail, patient room and toilet doorknobs, air conditioner filter, toilet bowl, table, floor outside patient room; isolation facility: table, floor, window, nurses benches, wall, TV; hospital: bed side surface, washbasin, toilet bowl button and doorknob, food table, light switch |
| Nelson et al. [53] | BR | Canada | SARS-CoV-2 | HCF | Long term care facilities | 89 sampling sites: high-touch surfaces, communal sites, and mobile medical equipment |
| Ong et al. [54] | Letter | Singapore | SARS-CoV-2 | HCF | Hospital-quarantine room | Patient’s room: cardiac table, bed rail, control panel on bed, call bell, locker, chair, light switches, stethoscope, floor, sink, glass window and door PPE storage area, air fan; toilet area: door handle, toilet bowl, hand rail, sink; anteroom: sink, floor, glass doors; corridor outside room floor, staff PPE |
| Pasquarella et al. [55] | Article | Italy | SARS-CoV-2 | HCF | 1 Hospital room | Bed rails and table, head and foot of the bed, floor near the bed, door handle, call button, chair, wall behind the bed, trapeze bar, air intake grill, head of the wheelchair, stethoscope, inner surgical mask |
| Peyrony et al. [56] | Article | France | SARS-CoV-2 | HCF | Emergency dept. patient care and non patient care areas, PPE from health care workers | Direct contact surfaces and non contact surfaces in patient care area and non patient care area, before and after decontamination |
| References | Type of publication | Country | Virus | Setting | Place | Investigated surfaces |
|------------|---------------------|---------|-------|---------|-------|-----------------------|
| 28 Razzini et al. [57] | Article | Italy | SARS-CoV-2 | HCF | Hospital: contaminated, semicontaminated and clean area | Surfaces in contaminated, semicontaminated and clean areas: bedrails, benches, computer keyboards, door handles, glove boxes, hand sanitizer dispensers, medical equipment, medical equipment touch screens, shelves for medical equipment, staff lockers, walls, waste containers, water taps, and windows |
| 29 Ryu et al. [58] | Article | South Korea | SARS-CoV-2 | HCF | Hospital A and B in South Korea—rooms of infected subjects in hospital A. Intensive Care Unit (ICU) and patient rooms in hospital B | Anteroom, corridor, nursing station, patient monitor, pillow, Ambu bag, infusion pump, fluid stand door button or knob, bed side rail, head of the bed, top of the television, air exhaust damper, anteroom, corridor, nursing station, centre of room floor, seat of toilet, bed side rail |
| 30 Santarpia et al. [59] | Article | USA | SARS-CoV-2 | HCF | Hospital: Nebraska Biocontainment Unit (NBU) and quarantine rooms in National Quarantine Unit (NQU) | Common room surfaces, high-touch personal items, and toilets |
| 31 Shin et al. [60] | Observation | South Korea | SARS-CoV-2 | HCF | Isolation room of 2 patients | Mobile phones (both), tablet (daughter), bedside tables (both), entire length of bed rail (daughter), call bell attached to the bed (mother), wall panel/door handle (in room), floor, and sink/toilet bowl (bathroom) |
| 32 Tan et al. [61] | Article | China | SARS-CoV-2 | HCF | Wuhan, China Isolation wards and ICU | Patient mobile phones, bedrails, door handles, light switches, side tables, and medical instruments in the patient wards, as well as low-touch surfaces including floors and chairs in the corridor. PPE |
| 33 Wang et al. [62] | Article | China | SARS-CoV-2 | HCF | Isolation wards and ICU | Several objects in rooms |
| 34 Wang et al. [63] | Article | China | SARS-CoV-2 | HCF | ICU and isolation ward | Nightstands, monitor screens, bed rail, intercoms, sinks, door handles, personal digital assistant, door handles, keyboards, N95 mask, gloves |
| References  | Type of publication | Country   | Virus     | Setting | Place                      | Investigated surfaces                                                                 |
|------------|---------------------|-----------|-----------|---------|----------------------------|----------------------------------------------------------------------------------------|
| 35 Wei et al. [64] | Article             | China     | SARS-CoV-2 | HCF     | Isolation ward             | Patient mobile phones, bedrails, door handles, light switches, side tables, and medical instruments in the patient wards, as well as low-touch surfaces including floors and chairs in the corridor |
| 36 Wu et al. [65] | Article             | China     | SARS-CoV-2 | HCF     | Living quarter and medical area | Beeper, keyboard, computer mouse, telephone, door handle, desktop, medical equipment, bedrail, bedside table, oxygen cylinder valve, gloves |
| 37 Yamagishi et al. [66] | BR                  | Japan     | SARS-CoV-2 | Public areas | Cruise cabins and common areas | Cabin surfaces and common areas |
| 38 Ye et al. [67] | Article             | China     | SARS-CoV-2 | HCF     | Medical and public area     | Self-service printer, table top, keyboard, doorknob, phone, medical equipment, public facilities, wall, floor, Hand sanitizer Dispensers, Gloves, Eye Protection or Face Shield |
| 39 Yung et al. [68] | BR                  | Singapore | SARS-CoV-2 | HCF     | Isolation Room of an infected child | Infant bed, cot side, table, HCW's face shield, mask and gown |
| 40 Zhou et al. [69] | Article             | UK        | SARS-CoV-2 | HCF     | 7 clinical areas and a public area | |
Supplementary Information

The online version contains supplementary material available at https://doi.org/10.1007/s10654-021-00784-y.

Declarations

Conflict of interest

The authors have no competing interests to declare.

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