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Effectiveness of contact tracing apps for SARS-CoV-2: a rapid systematic review

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Title

Effectiveness of contact tracing apps for SARS-CoV-2: a rapid systematic review

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Competing interest statement

All authors have completed the Unified Competing Interest form (available on request from the corresponding author) and declare: no support from any organisation for the submitted work [or describe if any]; no financial relationships with any organisations that might have an interest in the submitted work in the previous three years, no other relationships or activities that could appear to have influenced the submitted work

Transparency declaration

The lead author affirms that the manuscript is an honest, accurate, and transparent account of the study being reported; that no important aspects of the study have been omitted.

Ethical approval

No ethical approval was required for this study

Patient and public involvement statement

No patients or public were involved in the conception and execution of this study

Dissemination statement

We do not plan to further disseminate the results of this study apart from its publication

Abstract

Objective – To systematically review evidence on effectiveness of contact tracing apps (CTAs) for SARS-CoV-2 on epidemiological and clinical outcomes

Design – Rapid systematic review

Data sources - EMBASE (OVID), MEDLINE (PubMed), BioRxiv, and MedRxiv were searched up to October 28th

Study selection – Studies, both empirical and model-based, assessing effect of CTAs for SARS-CoV-2 on reproduction number (R), total number of infections, hospitalization rate, mortality rate, and other epidemiologically and clinically relevant outcomes, were eligible for inclusion.

Data extraction – Empirical and model-based studies were critically appraised using separate checklists. Data on type of study (i.e. empirical or model-based), sample size, (simulated) time horizon, study population, CTA type (and associated interventions), comparator, and outcomes assessed, were extracted. The most important findings were extracted and narratively summarized. Specifically for model-based studies, characteristics and values of important model parameters were collected.

Results – 2140 studies were identified, of which 17 studies (two empirical, 15 model-based studies) were eligible and included in this review. Both empirical studies were observational (non-randomized) studies and at high risk of bias, most importantly due to risk of confounding. Risk of bias of model-based studies was considered low for 12 of 15 studies. Most studies demonstrated beneficial effects of CTAs on R, total number of infections, and mortality rate. No studies assessed effect on hospitalization. Effect size was dependent on model parameters values used, but in general a beneficial effect was observed at CTA adoption rates of 20% or higher.

Conclusions – Contact tracing apps have the potential to be effective in reducing SARS-CoV-2 related epidemiological and clinical outcomes, though effect size depends on other model parameters (e.g. proportion of asymptomatic individuals, or testing delays), and interventions after CTA notification. Methodologically sound comparative empirical studies on effectiveness of CTAs are required to confirm findings from model-based studies.

Strengths and limitations of this study

- This is the first paper to provide a comprehensive overview and critical appraisal of studies assessing the effectiveness of contact tracings apps for SARS-CoV-2 on clinical and epidemiological outcomes
- Studies were retrieved using a large repository that is developed by a specific search string dedicated to identify studies on SARS-CoV-2 published in various underlying databases
- Critical appraisal was performed by reviewers from diverse backgrounds (i.e. mathematical modelling, epidemiology, medicine, systematic reviews) using predefined customized templates for both empirical and model-based effectiveness studies
- Given the rapid execution and (preprint) publication of studies on effectiveness of contact tracing apps for SARS-CoV-2, this review is unlikely to include the most recent studies that published after the search date
- Due to too much heterogeneity across studies, it was not feasible to provide a
 pooled meta-analysis estimate of the effectiveness of contact tracing apps for
 SARS-CoV-2 on the clinical and epidemiological outcomes

Introduction

The severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) outbreak has dominated worldwide news and scientific research throughout 2020. Since the outbreak in Wuhan (People's Republic of China) in early December 2019, reducing transmission of SARS-CoV-2 has been a worldwide priority. Digital technology could be applied for efficient contact tracing. Contact tracing applications (CTAs) are able to identify individuals who have recently been in close contact with infected individuals (and may have acquired infection as a consequence). After identification, the contact person can be instructed to go in self-quarantine, preventing further transmission and spread of the virus.

A substantial amount of research on CTAs for SARS-CoV-2 has been performed since the start of the pandemic. Summarizing all evidence, including results from research that has not yet, or is currently undergoing peer-review, is warranted to provide an overview of what is known regarding CTA effectiveness. Research that has not yet undergone peer-review is often published by authors through so-called preprint databases. However, identifying these articles, extracting data, and drawing conclusions can be a challenge, as this requires knowledge on epidemiology, mathematical modelling, systematically appraising evidence, and summarizing that evidence.

A few overviews of evidence on effectiveness of CTAs have been published in recent time. Anglemyer *et al.* provided an overview of study characteristics and quality appraisal of studies on effectiveness of CTAs and other digital contact tracing technologies. (1) However, their data are based on both SARS-CoV-2 infections and other infections (e.g. Ebola), and lack a quantitative effectiveness measure of CTAs on clinically relevant outcomes. Other systematic reviews focused only on user experience in using a CTA for SARS-CoV-2 (2), or only studied manual as opposed to digital contact tracing (3). One systematic review did look into studies on automated and semi-automated CTAs for SARS-CoV-2, but lacked reporting on CTA effectiveness on total number of infections, and hospitalization or mortality rates. (4)

In this rapid systematic review, we aim to evaluate all (empirical and model based) studies addressing effectiveness of CTAs for SARS-CoV-2 on relevant, i.e. epidemiological and

clinical, outcomes. We will provide descriptive characteristics, critical appraisal, and a narrative summary of evidence of included studies.

Methods

Search strategy

The *Bern COVID-19 Open Access Project (COAP)* database was used for identification of relevant research. The COAP database is comprised of research from EMBASE (OVID), MEDLINE (PubMed), BioRxiv en MedRxiv databases, specifically focused on SARS-CoV-2. On October 28th 2020 the COAP database was searched for scientific literature evaluating the effectiveness of CTAs for SARS-CoV-2 on epidemiological and clinical outcomes. The complete search strategy, as well as background information on the COAP database provided by Bern University, are provided in Supplementary File 1.

Eligibility criteria

Empirical (both observational and experimental) and model-based studies evaluating effectiveness of CTAs for SARS-CoV-2 were eligible for inclusion. Peer-reviewed publications as well as preprint papers were considered.

CTAs were considered when they provided feedback about potential recent exposure to an infected individual, based on proximity measurements (e.g. Bluetooth or GPS). Feedback should be provided directly to the individual through a CTA, although other feedback mechanisms, such as personal devices (e.g. a smartwatch), were also considered. National emergency warning systems using SMS were also included, provided they used proximity data to inform individuals.

All epidemiologically or clinically relevant outcomes quantifying the impact of CTAs were considered, which include but are not limited to: the reproduction number (R), total number of infections, hospitalization rate, and mortality rate related to SARS-CoV-2. Studies investigating other relevant outcomes, such as prevention of outbreaks or a second infection wave of SARS-CoV-2, were also included. Studies assessing (determinants affecting) adoption rate of CTAs, temporal change in incidence SARS-CoV-2, or other non-epidemiological or clinical outcomes were excluded.

Study selection

Studies identified in the search were first screened independently on title and abstract by two reviewers. Relevant studies were included for full text screening, and further selection of articles was performed by two independent reviewers. Any discrepancies were discussed and resolved. When consensus was not reached, a third reviewer was consulted to provide the final judgement.

Critical appraisal

Risk of bias was systematically assessed by two researchers using separate checklists for empirical and model-based studies. Discrepancies between researchers were discussed, and a final verdict was provided by a third reviewer if consensus was not reached. Empirical studies were appraised using a formal scoring method based on the Critical Appraisal Skills Programme (CASP) and Cochrane's Effective Practice and Organisation of Care (EPOC) checklists (5, 6) (Supplementary file 4). Risk of bias in model-based research was evaluated by assessing use of empirical input data for the model, number of scenarios analyzed, and transparency of model reporting. (Supplementary file 5)

Data extraction

Data extraction was performed by one reviewer, and checked by a second reviewer. Descriptive characteristics on type of research, i.e. empirical or model-based, sample size, (simulated) time horizon, study population, CTA properties and intervention, comparator, and epidemiological and clinical outcomes studied, were extracted from all included studies.

Specifically for model-based research, model characteristics (i.e. type of model and distributions used) and values used for important model parameters were collected. Furthermore, CTA specific properties were extracted, such as the method of contact tracing used by these apps. Forward tracing CTAs can only detect the 'offspring', i.e. individuals the index case has infected, of an infected individual. Bidirectional tracing CTAs also detect the 'parents', i.e., the individual that infected the index case of an infected individual. Models were considered to use bidirectional (as opposed to forward) tracing when, after the index case is detected and registered, all contacts within a period of at least the incubation time are identified, such that the parent of the index case could be found.

Another CTA specific property included the use of 1-step-tracing or sequential tracing. When a CTA-identified individual could only notify their contacts after testing positive themselves, this was considered 1-step-contact tracing. When notified contacts could subsequently also notify their own contacts, creating a cascade, even before that individual has shown symptoms or received a positive test result for SARS-CoV-2, this was considered sequential tracing.

The most important findings regarding effectiveness of CTAs for SARS-CoV-2 on epidemiological and clinical outcomes were extracted, synthesized, and reported narratively. These outcomes were pooled quantitatively whenever it was feasible to do so.

Results

Study selection

A total of 2140 potential studies were identified by the search. After selection based on title and abstract, 2059 articles were excluded. Full texts of the 81 remaining studies were assessed, after which 17 articles were included for critical appraisal and data extraction (Supplementary file 2). The 64 excluded studies with their reasons for exclusion are summarized in Supplementary file 3.

Characteristics of included studies

Seventeen primary studies were included, of which two were empirical observational (non-randomized) studies, and 15 were model-based studies (Table 1).

Six of the 17 studies were published preprints, meaning they had not (yet) gone through the peer review process at the time of data extraction (7-12). Included studies focused predominantly on the general population, although some analyzed the effectiveness of CTAs for specific populations such as hospital personnel, or school children (8, 9, 11, 13-16). Especially in model-based studies, results were often presented graphically. Consequently, the effectiveness of CTAs on epidemiological and clinical outcomes was only partly, or not at all, reported in key numerical figures.

The model-based studies typically assessed the effectiveness of CTAs by simulating one or more scenarios based on certain baseline or input values (e.g. proportion of asymptomatic infections). Table 2 provides an overview of characteristics and the most important input parameters used in models of the 15 included articles. Nine of the 15 model-based studies evaluated forward tracing CTAs (8, 9, 11, 13-18), four studies

analyzed bidirectional tracing CTAs (7, 10, 12, 19), and one used an alternative method (20). Four studies used a CTA that used sequential tracing (7, 10, 12, 19). All of these also used bidirectional CTAs, which are more effective than forward tracing CTAs in reducing R, but require quarantining many more contact persons. This is especially the case when a significant number of infections come from asymptomatic individuals (i.e. transmission from a case who does not (yet) have symptoms), who are unaware they have SARS-CoV-2. (19)

The percentage of CTA adoption was varied in almost all studies, allowing for assessment of the impact of CTAs on epidemiological and clinical outcomes. Average incubation time, i.e. the mean time between infection and symptom onset of SARS-CoV-2, was estimated to be 5 to 6 days for SARS-CoV-2 (9, 11-21). The proportion of asymptomatic SARS-CoV-2 infections, used as input parameter in model-based studies, was estimated at 20% to 50% based on empirical data (8, 9, 16, 18), but could vary between 18% to 86% (9). The baseline R value chosen in the model-based studies varied between 1.2 and 4.0. (7-10, 12, 14-21)

Furthermore, so-called superspreaders (i.e. individuals that infect numerous other individuals, and consequently have a high individual R) were discussed in context of the SARS-CoV-2 pandemic. Tracing these superspreaders is key in containing outbreaks. Hence, it is warranted to use bidirectional CTAs to trace these superspreaders, and advise them to immediately enter quarantine on identification. (14, 22)

Critical appraisal

Risk of bias in the two empirical studies was judged to be high (Table 3) (23, 24). Confounding variables (such as smoking, work status, and income) were insufficiently taken into account given the explanatory and observational nature of these empirical studies. It was also unclear how missing (outcome) data were dealt with.

Most model-based research was judged to have a low risk of bias (Table 4). Three of the 15 studies had a high risk of bias due to the lack of use of empirical distributions for variables, the limited number of scenarios analyzed, and insufficient transparency regarding reporting of the model. (11, 20, 21)

Synthesis of results

Evidence from empirical studies

Two empirical comparative observational studies assessed the effectiveness of CTAs compared to a control group that did not use CTAs (Table 1). (23, 24) One study looked at effectiveness of a text warning system used in 627,386 individuals who came in contact with an exposed population, and compared it to the general population of Taiwan who did not use such a warning system. (17) They showed a reduction in incidence of respiratory syndrome from 19.23 to 16.87 per 1000 individuals. They also showed a reduction in pneumonia incidence from 3.81 to 2.36 per 1000 individuals. (17) The second observational study investigated the introduction and adoption of a 'Test and Trace' app by 34,000 individuals living on the Isle of Wight (UK), and compared the estimated value of R in that region to that in the general UK population. (24) The CTA marked individuals as positive based on self-reporting of symptoms. Individuals that came in contact with an individual marked as positive were provided with social distancing advice. The study found that R was reduced from 1.3 to 0.5 after implementation of the CTA. Within 2 to 3 weeks after implementation, incidence of SARS-CoV-2 diagnoses declined by around 90%. (24)

Evidence from model-based studies

Effect on R

Effectiveness of a 1-step-contact tracing in reducing R can be approached using the following formula:

$$R_c = R * (1 - p^2 * f)$$

Here, R_c is the reproduction number when a CTA is used, R is the reproduction number without the use of a CTA, p is the proportion of the population using the CTA, and f is the combination of other factors that affect effectiveness of notification by the CTA. Such factors include, but are not limited to, delay between CTA notification and testing, delay between testing and test result, delay between reception of test result and entry of that result in the CTA, compliance to interventions (e.g. self-quarantine), and the proportion of infections that occur pre- or asymptomatically. Note that p occurs as a quadratic term, which reflects the fact that both infector and infectee have to use the CTA for the transmission to get traced.

Nine of the 15 model-based studies assessed the effect of CTAs on reduction of R. (8, 11, 14-16, 18, 19, 21) CTAs were able to control an ongoing outbreak or epidemic through quicker and more efficient feedback of a positive test result, and by notifying close contacts of a positively tested individual. (15, 16, 19) This speed and efficiency were not feasible using traditional manual contact tracing. (16) New outbreaks could be controlled (i.e. R_c <1.0) by CTAs, by combining them with quarantine or self-isolation interventions, provided that hygiene and social distancing measures are maintained. (8, 14, 18, 21) CTAs were able to reduce R by 0.3 more than traditional manual contact tracing, provided that feedback about contact with a positively tested individual is given to all contacts of the index case of the preceding 7 days. (19) Another model-based study demonstrated that a CTA with 20% adoption rate reduces R by 17.6% compared to no contact tracing, whereas traditional manual contact tracing reduced R by 2.5% compared to no contact tracing. (15) This study also demonstrated that a CTA is able to reduce the R further, even when social distancing has already reduced R to 1.2. In this situation, R can be reduced further by 30% to 0.8 when CTA adoption rate is 80%. (15) Another model-based study determined that 60% adoption rate of a CTA could result in an R below 1.0. (11) In one study, adoption rate of 53% resulted in a 47% reduction in R when the complete household of an individual with a positive test result is advised to be guarantined. (14) The last study looking at effect of CTA on R showed that only at 60% adoption rate of the app a significant beneficial effect on R would become apparent. (12) When R is high (e.g. 3.0), and a considerable proportion of individuals is asymptomatic (e.g. 40% of all infections), CTAs need to be combined with other interventions (such as social distancing and random testing) to be able to lower the R below 1.0. (12) Potential for CTAs to reduce R is not only dependent on the adoption rate of the app, but also on (effectiveness of) various other measures that are provided after a positive notification, the delay between positive notification and opportunity for testing, and delay between receiving a positive test result and sharing that result through the CTA. (5, 6, 10) One study found that the percentage of preventable infections by one individual strongly depends on the time delay between CTA notification and the ability to be tested. (15) When there was no delay (i.e. 0 days) 79.9% of infections could be prevented, compared to 41.8% and 4.9% for 3 and 7 days delay respectively.

Effect on total number of infections

Eight of the 15 model-based studies assessed the effect of CTAs on reducing the total number of infections. (8-11, 13, 17, 18, 20) Two studies indicated that the success of CTAs

in reducing the total number of infections could only be ensured with a high adoption rate of that app. (8, 13) Another study showed that with a high CTA adoption rate of 75%, there would be no more new infections occurring within three months after implementation. (11) It was found that adequate hygiene and social distancing measures are needed to enable CTAs to reduce the total number of infections. (8, 9, 17, 18) Especially in areas where there is low compliance to social distancing, a sufficiently high adoption rate of a CTA is essential to maintain control of an outbreak. (9)

The height of the peak number of new infections can, according to one study, be reduced by half with a 50% adoption rate of a CTA (18), whereas another study showed that this could be achieved with an adoption rate as low as 20%. (20) Another study demonstrated that at 27% CTA adoption rate, a quarter of all new infections can be prevented. (17) However, according to another study that used a similar adoption rate, the number of infections would stabilize, but the epidemic would be maintained by core groups in densely populated areas. (18) There may be a period of time of more than two months between implementation of interventions (such as CTAs) and the effect of that implementation on the total number of SARS-CoV-2 infections. (13)

Effect on number of hospitalizations

None of the 15 model-based studies assessed the effect of CTAs on the number of hospitalizations due to SARS-CoV-2 infection, possibly because the number of hospitalizations is expected to be proportional to the number of infections, only with a time-delay. A German study did look into the effect of a CTA on the number of days that intensive care unit (ICU) capacity was exceeded. (9) They found in their simulations that – based on the German population, and assuming an ICU capacity of 24.000 beds – a CTA adoption rate of 20% would prevent exceedance of ICU capacity at any point in time. In contrast, if no contact tracing (either manual or digital) would be used, ICU capacity would be exceeded on a quarter of days.

Effect on mortality rate

Three of the 15 model-based studies assessed the effect of CTAs on mortality rate. (9, 18, 20) One study demonstrated that a high adoption rate (80%) of a CTA would result in an 85% reduction in mortality rate, over a period of 500 days (9). Another study found that a low CTA adoption rate (25%) is associated with a 10% decrease in mortality rate, an average adoption rate (50%) with 25% decrease, and a high adoption rate (75%) with 40-

60% decrease. (18) A third study showed that at 40% adoption rate, during the peak of an outbreak, a reduction in number of deaths by 97% could be achieved. (20)

Discussion

Empirical evidence regarding the effectiveness of using CTAs for detection of SARS-CoV-2 is still limited. Currently, no randomized studies have been performed, and only two observational comparative studies were identified in this systematic review. Although some benefits of using CTAs for detection of SARS-CoV-2 were observed, both studies were deemed to be of low methodological quality. However, the results of these studies were in accordance with the 15 included, higher quality, model-based studies assessing effectiveness of CTAs. These studies showed that CTAs can be effective and a valuable addition to manual contact tracing. CTA use resulted in a lower R, lower total number of infections, and lower mortality rate. These reductions were already observed at relatively low adoption rates (e.g. 20%), though higher adoption rates of CTAs resulted in greater reductions. Shortening delays between CTA notification and diagnostic testing may increase its effectiveness.

This rapid systematic review assesses key features, quality, and main clinical and epidemiological outcomes of a set of studies, both empirical and model-based, on effectiveness of CTAs for SARS-CoV-2. To our knowledge, no such systematic review has been published, assessing these specific properties. Methodological quality of empirical studies was assessed using standardized tools. No such tool was available in literature for model-based studies, and as such a set of key features used in other systematic reviews on this topic was used. This set was validated by experts in mathematical modelling.

To fully appreciate the findings from this systematic review, some considerations should be taken into account. First, the studies found through the literature search may not be a comprehensive set. Studies on SARS-CoV-2 are published at a rapid, almost daily, basis in various online repositories. Although we cannot ensure that all studies on the effectiveness of CTAs for SARS-CoV-2 have been identified, we believe that the set of included studies that we have identified represents a representative sample.

Furthermore, effectiveness of CTAs for SARS-CoV-2 described in model-based studies is complex. Numerous input variables used in the models interact with one another, and

consequently affect effectiveness of, for example, adoption rate of CTAs on clinical or epidemiological outcomes. Summarizing these findings into a general effectiveness is difficult, and will always suffer from simplification of a system of complex interactions. Though we feel that providing some (conditional) findings from these studies will help provide some general insight in the impact CTAs can have on clinical and epidemiological outcomes for SARS-CoV-2.

Current evidence on the effectiveness of CTAs for SARS-CoV-2 is predominantly based on modelling studies, which indicate that there is potential in beneficially affecting key clinical and epidemiological outcomes. High quality empirical evidence, either from experimental or methodologically sound observational studies, is needed in order to be able to draw more robust conclusions regarding effectiveness of CTAs for SARS-CoV-2.

References

- 1. Anglemyer A, Moore TH, Parker L, Chambers T, Grady A, Chiu K, et al. Digital contact tracing technologies in epidemics: a rapid review. Cochrane Database Syst Rev. 2020;8:CD013699.
- 2. Davalbhakta S, Advani S, Kumar S, Agarwal V, Bhoyar S, Fedirko E, et al. A Systematic Review of Smartphone Applications Available for Corona Virus Disease 2019 (COVID19) and the Assessment of their Quality Using the Mobile Application Rating Scale (MARS). J Med Syst. 2020;44(9):164.
- 3. Nussbaumer-Streit B, Mayr V, Dobrescu AI, Chapman A, Persad E, Klerings I, et al. Quarantine alone or in combination with other public health measures to control COVID-19: a rapid review. Cochrane Database Syst Rev. 2020;4:CD013574.
- 4. Braithwaite I, Callender T, Bullock M, Aldridge RW. Automated and partly automated contact tracing: a systematic review to inform the control of COVID-19. Lancet Digit Health. 2020;2(11):e607-e21.
- 5. CASP Checklist [Internet]. Critical appraisal skills programme. 2018 [cited 15-10-2020]. Available from: https://casp-uk.net/wp-content/uploads/2018/01/CASP-Cohort-Study-Checklist 2018.pdf.
- 6. EPOC Resources for review authors [Internet]. Cochrane Effective Practice and Organisation of Care (EPOC). 2017 [cited 15-10-2020]. Available from: https://epoc.cochrane.org/resources/epoc-resources-review-authors.
- 7. Bulchandani VB, Shivam S, Moudgalya S, Sondhi SL. Digital Herd Immunity and COVID-19. medRxiv. 2020:2020.04.15.20066720.
- 8. Cencetti G, Santin G, Longa A, Pigani E, Barrat A, Cattuto C, et al. Digital Proximity Tracing in the COVID-19 Pandemic on Empirical Contact Networks. medRxiv. 2020:2020.05.29.20115915.
- 9. Grimm V, Mengel F, Schmidt M. Extensions of the SEIR Model for the Analysis of Tailored Social Distancing and Tracing Approaches to Cope with COVID-19. medRxiv. 2020:2020.04.24.20078113.
- 10. Guttal V, Krishna S, Siddharthan R. Risk assessment via layered mobile contact tracing for epidemiological intervention. medRxiv. 2020:2020.04.26.20080648.
- 11. Shamil MS, Farheen F, Ibtehaz N, Khan IM, Rahman MS. An Agent Based Modeling of COVID-19: Validation, Analysis, and Recommendations. medRxiv. 2020:2020.07.05.20146977.
- 12. Pollmann TR, Pollmann J, Wiesinger C, Haack C, Shtembari L, Turcati A, et al. The impact of digital contact tracing on the SARS-CoV-2 pandemic a comprehensive modelling study. medRxiv. 2020:2020.09.13.20192682.
- 13. Scott N, Palmer A, Delport D, Abeysuriya R, Stuart RM, Kerr CC, et al. Modelling the impact of relaxing COVID-19 control measures during a period of low viral transmission. Med J Aust. 2021;214(2):79-83.
- 14. Kucharski AJ, Klepac P, Conlan AJK, Kissler SM, Tang ML, Fry H, et al. Effectiveness of isolation, testing, contact tracing, and physical distancing on reducing transmission of SARS-CoV-2 in different settings: a mathematical modelling study. Lancet Infect Dis. 2020;20(10):1151-60.
- 15. Kretzschmar ME, Rozhnova G, Bootsma MCJ, van Boven M, van de Wijgert J, Bonten MJM. Impact of delays on effectiveness of contact tracing strategies for COVID-19: a modelling study. Lancet Public Health. 2020;5(8):e452-e9.
- 16. Ferretti L, Wymant C, Kendall M, Zhao L, Nurtay A, Abeler-Dörner L, et al. Quantifying SARS-CoV-2 transmission suggests epidemic control with digital contact tracing. Science. 2020;368(6491):eabb6936.
- 17. Currie DJ, Peng CQ, Lyle DM, Jameson BA, Frommer MS. Stemming the flow: how much can the Australian smartphone app help to control COVID-19? Public Health Res Pract. 2020;30(2).
- 18. Ferrari A, Santus E, Cirillo D, Ponce-de-Leon M, Marino N, Ferretti MT, et al. Simulating SARS-CoV-2 epidemics by region-specific variables and modeling contact tracing app containment. NPJ Digit Med. 2021;4(1):9.

- 19. Bradshaw WJ, Alley EC, Huggins JH, Lloyd AL, Esvelt KM. Bidirectional contact tracing could dramatically improve COVID-19 control. Nat Commun. 2021;12(1):232.
- 20. Nuzzo A, Tan CO, Raskar R, DeSimone DC, Kapa S, Gupta R. Universal Shelter-in-Place Versus Advanced Automated Contact Tracing and Targeted Isolation: A Case for 21st-Century Technologies for SARS-CoV-2 and Future Pandemics. Mayo Clin Proc. 2020;95(9):1898-905.
- 21. Kurita J, Sugawara T, Ohkusa Y. Estimated effectiveness of school closure and voluntary event cancellation as COVID-19 countermeasures in Japan. J Infect Chemother. 2021;27(1):62-4.
- 22. Endo A, Leclerc QJ, Knight GM, Medley GF, Atkins KE, Funk S, et al. Implication of backward contact tracing in the presence of overdispersed transmission in COVID-19 outbreak. medRxiv. 2020:2020.08.01.20166595.
- 23. Chen CM, Jyan HW, Chien SC, Jen HH, Hsu CY, Lee PC, et al. Containing COVID-19 Among 627,386 Persons in Contact With the Diamond Princess Cruise Ship Passengers Who Disembarked in Taiwan: Big Data Analytics. J Med Internet Res. 2020;22(5):e19540.
- 24. Kendall M, Milsom L, Abeler-Dorner L, Wymant C, Ferretti L, Briers M, et al. Epidemiological changes on the Isle of Wight after the launch of the NHS Test and Trace programme: a preliminary analysis. Lancet Digit Health. 2020;2(12):e658-e66.



Table 1. Descriptive characteristics of included studies

Characteristics of empirical epidemiological and model-based studies looking at effectiveness of contact- and tracing apps for SARS-CoV-2. N/R = 1 not reported, R = 1 reproduction number, $R_0 = 1$ baseline reproduction number

8 9 10	Study	Country (of first author)	Study type	Sample size / # of simulations	Time horizon	Population	Specific setting(s)	Intervention	Comparison	Outcome(s)	Main findings
11 12 13 14 15 16 17 18 19	Bradshaw 2020 (peer reviewed)	Germany	Modelling	500 or 1000 simulations	52 weeks or 10,000 cases	General population		Contact tracing app (Bluetooth) with quarantine	- Manual contact tracing - Current practice	- R - Outbreak control	- Bidirectional tracing will enable more effective control of COVID-19 - Switching from forward to bidirectional tracing can reduce R by 0.3 if the tracing time window is sufficiently wide - High adoption of bidirectional manual and digital contact tracing is 3x more effective at outbreak control compared to current practice
21 22 23 24 25	Bulchandani 2020 (preprint)	USA	Modelling	4000 simulations	N/R	Susceptible population (i.e. no immunity)	- 101	Contact tracing app (not specific) with quarantine	-	- R - Outbreak control	- Outbreak control is possible regardless of proportion of asymptomatic transmission - Outbreak control requires a contact tracing app adoption of 75%-95%
26 27 28 29 30 31 32 33 34 35 36 37 4	Cencetti 2020 (preprint)	Italy	Modelling	20 simulations	50 days	General population	- University Campus - High school - Workplace	Contact tracing app (Bluetooth) with quarantine	0/1/	- R - Outbreak control	- Reduction of R and outbreak control is dependent on contact tracing efficiency, isolation efficiency, and R ₀ - Outbreak control can be achieved through tracing and isolation, provided that hygiene and social distancing measures limit R ₀ to 1.5 - Outbreak control not feasible if contact tracing app adoption is insufficient or if R ₀ is >2
38 39 40	Chen 2020 (peer reviewed)	Taiwan	Empirical	3000 individuals	40 days	General population (Taiwan)	-	SMS warning (GPS) with quarantine &	Current practice	RespiratorysyndromePneumonia	- Contact tracing and SMS feedback resulted in less cases of respiratory syndrome (16.87 vs 19.23 per 1000) and

							symptom monitoring			pneumonia (2.36 vs 3.81 per 1000) compared to the general population
										- Resource requirements for manual contact tracing could be reduced by using contract tracing apps combined with big data analytics
Currie 2020 (peer reviewed) 3 4 5 6 6 7 8 9 0 1	Australia	Modelling	Not reported	12 months	General population (Australia)	- - - -	Contact tracing app (Bluetooth) with quarantine	No contact tracing app	- Outbreak control - Cumulative incidence SARS-CoV-2	- Outbreak control by a contact tracing app can be achieved when adoption is sufficient, and is combined with testing and social distancing - Cumulative incidence of SARS-CoV-2 can within 8 months (depending social distancing and testing intensity) be reduced to: 13-24% at an app adoption of 27% 17-35% at an app adoption of 40% 36-59% at an app adoption of 61% 47-76% at an app adoption of 80%
Ferrari 2020 (peer reviewed) 7 8 9 0 1 2 3 4 5 6 7 8 9 0 0 1 2 3 4 5 6 7 8 9 0 0	Italy	Modelling	simulations (per scenario)	50 days 300 days 400 days	General population (Italy)	-	Contact tracing app (not specified) with quarantine & symptom monitoring	O/7/	- R - Outbreak control - Cumulative incidence SARS-CoV-2 (symptomatic) - Mortality	- Reduction of R below 1.0 can be achieved when contact tracing apps have sufficient adoption, efficacy of case identification, and compliance to quarantine - Outbreak control can be achieved using contact tracing apps combined with voluntary self-quarantine and efficient case isolation, depending population density and transportation - Outbreak control was achieved with 75% app adoption rate - Cumulative incidence can be suppressed with 25% app adoption rate, but outbreaks will be sustained by districts with high population density

2											
3 4 5 6 7											 Mortality was reduced by: 10% at 25% app adoption rate 25% at 50% app adoption rate 40-60% at 75% app adoption rate
8 9 10 11 12 13 14 15	Ferretti 2020 (peer reviewed)	China	Modelling	40 simulations (pairs)	12 days 20 days	General population (China)	- Home - Train - Work	Contact tracing app (Bluetooth) with quarantine	Manual contact tracing	R	- Manual contact tracing is not able to stop outbreaks due to delays (~ 3 days), whereas contact tracing apps are able to prevent outbreaks - Reduction of R below 1.0 is feasible using instantaneous (red. without delays) contact tracing apps
16 17 18 19 20 21 22 23 24 25 26	Grimm 2020 (preprint)	Germany	Modelling	N/R	500 days	General population (Germany)	- High risk of severe course of infection - Low risk of severe course of infection	Contact tracing app (not specified) with quarantine	- No intervention - Uniform social distancing - Group specific social distancing	- Cumulative incidence SARS-CoV-2 - # of days ICU capacity exceeded - Mortality	- ICU capacity and mortality can be kept low by using contact tracing apps combined with tailored social distancing and personal protection measures - ICU capacity was not exceeded at any point with a contact tracing app adoption of 20% or more - Mortality was reduced by 85% when a high (80%) adoption rate of the contact tracing app was achieved
27 28 29 30 31 32 33 34 35 36	Guttal 2020 (preprint)	N/R	Modelling	N/R	150-200 days	General population	-	Contact tracing app (Bluetooth) with quarantine	0/1/	Cumulative incidence SARS-CoV-2	- Peak cumulative incidence can be flattened significantly even when a small fraction of cases are identified using contact tracing apps, tested and isolated - Peak cumulative incidence can strongly be reduced even if contact tracing app testing is only performed in the most probable individuals (p > 0.8)

2											
3 4 5 6 7 8 9	Kendall 2020 (peer reviewed)	United Kingdom	Empirical	Population- size Isle of Wight Population- size UK (except Wales)	<2 months	General population (Isle of Wight and UK (except Wales))	-	Contact tracing app (Bluetooth) with social distancing	-	- R - Cumulative incidence SARS-CoV-2	- Reduction of R from 1.3 to 0.5 was achieved after implementation of a contact tracing app - Cumulative incidence of SARS-CoV-2 reduced by 87% in 2-3 weeks after implementation of a contact tracing app
10 11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 30 31	Kretzschmar 2020 (peer reviewed)	Netherlan ds	Modelling	1,000 simulations		General		Contact tracing app (Bluetooth) with quarantine	Social distancing without contact tracing app	R	- Contact tracing apps, with short delays and high coverage for testing and tracing, could substantially reduce the R, alleviating more stringent control measures - Reduction of the R from 1.2 with social distancing alone to 0.8 (95% CI 0.7–1.0) by adding a contact tracing app with an adoption of 80% - Reduction of the R through contact tracing apps is more effective compared to manual contact tracing, with respectively 17.6% and 2.5% reduction of R compared to no contact tracing - Reduction in transmission rate (reflective of R) depends on tracing delay 0 79.9% with 0-day testing delay 0 41.8% with 3-day testing delay 0 4.9% with 7-day testing delay
32 33 34 35 36 37 38 39 40 41 42	Kucharski 2020 (peer reviewed)	United Kingdom	Modelling	25,000 simulations	N/R	General population (UK)	- Household - Work - School - Other	Contact tracing app (Bluetooth) with quarantine	-	- R - Outbreak control	- Combining contact tracing app with quarantine and reduce transmission more than mass testing or self-isolation alone - Reduction in transmission rate (reflective of R) was 47% when contact tracing app was used at 53% adoption rate - Maintaining an R < 1.0 requires a combination of self-isolation, contact tracing, and physical distancing

2											
3 4 5 5											- Outbreak control in a scenario where incidence is high requires a considerable number of individuals to be quarantined after contact tracing
7 8 9 10 11 12 13	Kurita 2020 (peer reviewed)	Japan	Modelling	N/R	5 months	General population (Japan)	-	Contact tracing app (Bluetooth) with quarantine	-	R	- Reduction of R < 1.3 using a contact tracing app is not feasible if there are no voluntary restrictions - Reduction of R < 1.0 is feasible if contact tracing app adoption is 10% combined with 15% compliance for voluntary restrictions against going out
15 16 17 18 19 20 21 22 23 24 25 26 27 28	Nuzzo 2020 (peer reviewed)	USA	Modelling	N/R	400 days 150 days	Susceptible individuals	10	Contact tracing app (GPS,WiFi, and/or Bluetooth) with quarantine	Shelter in place	- Cumulative incidence SARS-CoV-2 - Mortality	- Contact tracing apps can mitigate infection spread similar to universal shelter-in-place, but with considerably fewer individuals isolated - Cumulative peak incidence can be reduced by 49% at 20% app adoption rate - Cumulative peak incidence can be reduced by 90% at 50% app adoption rate (similar to 40% compliance to shelter in place) - Mortality can be reduced by 23% at 20% app adoption rate
29 30 31 32 33 34 35 36 37 38 39	Pollmann 2020 (preprint)	Germany	Modelling	100 simulations	500 days	General population	-	Contact tracing app (Bluetooth) with quarantine		- R - Outbreak control - Cumulative incidence SARS-CoV-2	- Recursive tracing by contact tracing apps is more efficient than 1-step-tracing - Contact tracing apps alone cannot bring R below 1.0, unless 100% adoption is approached, and app notifications are strictly followed by quarantining and testing - Reducing an R _o of >3.0, in which 40% are asymptomatic SARS-CoV-2 carriers, below 1.0, can only be achieved by a contact tracing app if combined with other
41 42											

2											
2 3 4 5 6 7 8 9 11 11 11 11 11 11	Scott 2020 (peer reviewed)	Australia	Modelling	N/R	3.5 months	Susceptible population (Victoria,	Various*	Contact tracing app (Bluetooth) with quarantine	-	Cumulative incidence SARS-CoV-2	interventions such as social distancing and/or random testing - Reducing R significantly requires a contact tracing app adoption rate of at least 60% - Cumulative incidence is reduced at any percentage of contact tracing app adoption - Impact of policy changes on cumulative incidence can take >2 months to become apparent
116 117 118 119 220 221 222 23 224 225 226 227 228 229						Australia)		ich	0		 Opening pubs/bars was identified as the greatest risk for increasing incidence of SARS-CoV-2. This could be mitigated by either of these measures: 30% app adoption rate is achieved Transmission within venues was reduced by >40% through physical distancing policies Manual contact tracing was used that enabled >60% of contacts to be traced Cumulative incidence is unlikely to be significantly impacted when app adoption
30 31 32 33 34 35 36 37 38 39 40	Shamil 2020 (preprint)	Banglades h	Modelling	N/R	60 days (Ford County) 120 days (New York city)	Susceptible population	- Healthcare workers - Students - Service holders - Unemployed people	Contact tracing app (not specified) with quarantine	- Lockdown - Extra personal protection	- R - Cumulative incidence SARS-CoV-2	- Reduction of R below 1.0 can be achieved within 3 weeks at 60% app adoption rate - Cumulative incidence approach zero within 3 months when 75% app adoption rate is achieved - Cumulative incidence is reduced by 3.5% when using a contact tracing app compared to not using one

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2					
3 [- Cumulative incidence is reduced by 4.6%
4					after 90 days when either:
5					
5					 All doctors, nurses, healthcare
7					workers and 50% of service holders
3					are using a contact tracing app for 2
9					days
10					o 75% of the population are using a
11					contact tracing app for 2 days

beer telien only

^{12 *} Household, school, work, community, church, professional sports, community sports, beaches, entertainment, cafés / restaurants, pubs / bars, public transport, national parks, public parks, large events, child care, social networks, and aged care

Table 2. Properties of model-based studies

Model-specific characteristics of model-based studies looking at effectiveness of contact- and tracing apps for SARS-CoV-2. Hyphens (-) indicate a continuous range between numbers, semicolons indicate separate distinct values. R = Reproductionnumber, N/A = not applicable, N/R = not reported, ODE = ordinary differential equations, PDE = partial differential equations, HH = household

	Model-related	l properties	Contact- an	d tracing app relat	ted properties		Disease-rel	ated properties	5	Mo	difyable pro	perties
Study	Modeltype	Input parameter properties	Tracing direction	# of sequential generations	Adoption rate app	R	Incubation time	Infectious period	Probability of disease transmission	Delay symptom onset and testing	Delay testing and feedback app	Quarantine effectiveness
Bradshaw 2020 (peer reviewed)	Branching- process model	Distributions	Bidirectional	Infinite generations	53;80%	2.5	5.5 days	Fitted to curve, value not specified	Fitted to curve, value not specified	1 days	0 days	90%
Bulchandani 2020 (preprint)	Branching- process model	Based on exponential distributions	Bidirectional	3-infinite generations	0-100%	3.0	N/A ¹	N/A	N/R	N/A ²	0 days	100%
Cencetti 2020 (preprint)	Continuous weighted temporal network	Distributions	Forward	1 generation	60;80;100%	1.2;1.5;2.0	Fitted to curve, value not specified	Fitted to curve, value not specified	Fitted to curve, value not specified	2 days	0 days	0-100%
Currie 2020 (peer reviewed)	ODE compartmental model	Based on exponential distributions	Forward	1 generation	0;27;40;61;80%	2.5	2.0 days	11 days	N/R	3 days	N/R	90%
Ferrari 2020 (peer reviewed)	ODE compartmental model	Based on exponential distributions	Forward	1 generation	0;25;50;75%	1.5	5.1 days	10 days	10%	2 days	N/R	90%
Ferretti 2020 (peer reviewed)	PDE compartmental model	Distributions	Forward	1 generation	0-100%	2.0	5.5 days	12 days	Fitted to curve, value not specified	1.6 days	0 days	0-100%
Grimm 2020 (preprint)	ODE compartmental model	Based on exponential distributions	Forward	1 generation	20-80%	2.2;3.0	5.0 days	10;12.5;14;20 days	N/R	N/R	N/R	100%³

		Model-related properties							lated propertie	<u>es</u>	Modifyable properties		
	Study	Modeltype	Input parameter properties	Tracing direction	# of sequential generations	Adoption rate app	R	Incubation time	Infectious period	Probability of disease transmission	Delay symptom onset and testing	Delay testing and feedback app	Quarantine effectiveness
0 1 2	Guttal 2020 (preprint)	Individual- based network model	Based on exponential distributions	Bidirectional	>1 generation	100%	3.0;4.0	N/A	20 days	0.2%	N/R	N/R	100%
3 4 5	Kretzschmar 2020 (peer reviewed)	Branching- process model	Distributions	Forward	1 generation	20;40;60;80;100%	2.5	6.4 days	10 days	2-12%	0 days	0 days	0;20;40; 60;80;100%
6 7 8	Kucharski 2020 (peer reviewed)	Individual- based network model	Distributions	Forward	1 generation	53%	2.6	5.0 days	5 days	20% within HH 6% outside HH 50% less for asymptomatic	0 days	0 days	90%
0 1	Kurita, 2020 (peer reviewed)	ODE compartmental model	Based on exponential distributions	N/R	1 generation	0;10;20;30;40;50; 60;70;80;90;100%	1.5	6.6 days	N/R	N/R	2 days	0 days	N/R
2 3 4 5	Nuzzo, 2020 (peer reviewed)	ODE compartmental model	Based on exponential distributions	N/A ⁴	N/A ⁴	0;10;20;30;40;50; 60;70;80;90%	3.02	5.1 days	N/R	Fitted to curve, value not specified	N/R	N/R	100%
6 7 8 9	Pollmann 2020 (preprint)	ODE compartmental model	Based on exponential distributions & distributions	Bidirectional	>1 generation	60;75;90;100%	2.0-3.0- 4.0	4.0;7.4 days	10 days	7% ⁵	0;2;4;6 days	N/R	100%
1 2 3	Scott 2020 (peer reviewed)	Agent-based model	Distributions	Forward	1 generation	0-50%	Fitted to curve, value not specified	4.6 days	8-14 days	Fitted to curve, value not specified	1 day	1 day	0% in HH 80-100% in other settings
4 5 6 7	Shamil 2020 (preprint)	Agent-based model	Distributions	Forward	1 generation	60;75%	Fitted to curve, value not specified	6.0 days	10 days	N/R	0 days	0 days	100%

¹ Fraction of infections before symptoms is relevant

² Isolation based on positive notification, not a positive test

³ Changing app coverage coveres imperfect isolation

⁴ No true tracing, fixed proportion cases will self-isolate

⁵ Time-dependent, maximum value reported in table

Table 3. Critical appraisal of empirical studies

Table - Critical appraisal empirical epidemiological studies looking at effectiveness of contact- and tracing apps for SARS-CoV-2

Study	Confounding?	Selection bias: participants?	Selection bias: missing data?	Information bias: intervention misclassification / non- compliance?	Information bias: Misclassification of the outcome?	Other concerns?	Overall risk of bias
Chen 2020 (peer reviewed)	Yes*	No	Unclear	No	Unclear	None	High
Kendall 2020 (peer reviewed)	Yes	No	Unclear	No	No	Competing interests and funding not reported	High
Only adjusted fo	or age			erien			

^{*} Only adjusted for age

Table 4. Critical appraisal of model-based studies

Table - Critical appraisal model based studies looking at effectiveness of contact- and tracing apps for SARS-CoV-2

Study	Were empirical distributions used for a varying infectiousness since time of infection?	Were various different scenarios evaluated for important model assumptions and parameter values?	Were models reported transparently? (i.e. no black box)	Other concerns?	Overall study validity
Bradshaw 2020 (peer reviewed)	Yes	Yes	Yes	External funding ¹	High
Bulchandani 2020 (preprint)	No	Yes	Yes	Competing interests & funding not reported	High
Cencetti 2020 (preprint)	Yes	Yes	Yes	No	High
Currie 2020 (peer reviewed)	Yes	Yes	Yes	No	High
Ferrari 2020 (peer reviewed)	No	Yes	Yes	Competing interests ²	High
Ferretti 2020 (peer reviewed)	Yes	Yes	Yes	No	High
Grimm 2020 (preprint)	No	Yes	Yes	No	High
Guttal 2020 (preprint)	Yes	Yes	Yes	Competing interests and funding not reported	High
Kretzschmar 2020 (peer reviewed)	Yes	Yes	Yes	No	High

Kucharski 2020 (peer reviewed)	Yes	Yes	Yes	Funding ³ , though no influence of funder on study results	High
Kurita, 2020 (peer reviewed)	No	No*	Unclear	Type of model used unclear	Low
Nuzzo, 2020 (peer reviewed)	No	No*	Yes	Potential competing interests ⁴	Low
Pollmann 2020 (preprint)	Yes	Yes	Yes	Competing interests and funding not reported	High
Scott 2020 (peer reviewed)	Yes	Yes	Yes	Funding ⁵	High
Shamil 2020 (preprint)	No	Yes	Unclear	No	Low

^{*} Scenarios were limited only to variation in rate of adoption of the contact- and tracing app and voluntary quarantine

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² E.S. works for Bayer, is collaborating to COVID Safe Paths app, by MIT, and advising LEMONADE tracing app, by Nuland. A.S.C. works for Roche Pharma. M.T.F is consultant for Ely Lilly.

³ Wellcome Trust, UK Engineering and Physical Sciences Research Council, European Commission, Royal Society, Medical Research Council.

⁴ Dr Raskar is the founder of a non-profit to facilitate digital contact tracing. The other authors report no potential competing interests.

⁵ Funding by the Burnet Institute



Supplementary file 1. Search strategy

Search strategy

On October 28th 2020 the comprehensive set of studies included in the COAP database (available on https://ispmbern.github.io/covid-19/living-review/collectingdata.html) was loaded in Endnote X9.

The dataset consisted of 82,401 references related to research on COVID-19. The following search was performed within this dataset:

(contact OR tracing OR track OR tracking OR warn OR warning) AND (smartphone OR app OR smartwatch OR device OR mobile OR smart phone OR bluetooth OR wearable OR iphone OR cell phone)

Background COAP database

The COAP database is a repository provided by Bern University, in which studies related to COVID-19 are incorporated. (available on https://ispmbern.github.io/covid-19/living-review/collectingdata.html)

Studies included in this repository are extracted on a daily basis from <u>EMBASE</u> (OVID), MEDLINE (PubMed), BioRxiv, and MedRxiv. References that are not yet available in the repository are added based on the date of publication provided by the aforementioned databases. The date on which the reference is added to the COAP database is included under the heading 'strategydate'.

Search strategies used for the COAP database are updated on a regular basis. An overview of these updates can be found below.

Initial search: 01.01.2020

MEDLINE

("Wuhan coronavirus" [Supplementary Concept] OR "COVID-19" OR "2019 ncov"[tiab] OR (("novel coronavirus"[tiab] OR "new coronavirus"[tiab]) AND (wuhan[tiab] OR 2019[tiab])) OR 2019-nCoV[All Fields] OR (wuhan[tiab] AND coronavirus[tiab])))))

EMBASE

ncov OR (wuhan AND corona) OR COVID

BioRxiv/MedRxiv

ncov or corona or wuhan or COVID

Update #1: 26.03.2020

MEDLINE

("Wuhan coronavirus" [Supplementary Concept] OR "COVID-19" OR SARS-CoV-2 OR "2019 ncov"[tiab] OR (("novel coronavirus"[tiab] OR "new coronavirus"[tiab]) AND (wuhan[tiab] OR 2019[tiab])) OR 2019-nCoV[All Fields] OR (wuhan[tiab] AND coronavirus[tiab]))

EMBASE

(nCoV or 2019-nCoV or ((new or novel or wuhan) adj3 coronavirus) or covid19 or covid-19 or SARS-CoV-2).mp.

BioRxiv/MedRxiv

ncov or corona or wuhan or COVID or SARS-CoV-2

With the kind support of the <u>Public Health & Primary Care Library PHC</u>, and following guidance of the Medical Library Association

Update #2: 01.04.2020

From 01.04.2020, we retrieve the currate BioRxiv/MedRxiv dataset Link

Update #3: 29.04.2020

MEDLINE

("coronavirus"[MH] OR "coronavirus infections"[MH] OR "coronavirus"[TW] OR "corona virus"[TW] OR "HCoV"[TW] OR "nCov"[TW] OR "covid"[TW] OR "covid19"[TW] OR "Severe Acute Respiratory Syndrome Coronavirus 2"[TW] OR "SARS-CoV2"[TW] OR "SARS-CoV 2"[TW] OR "MERS-CoV"[TW]) AND (2019/1/1:3000[PDAT])

Update #4: 01.05.2020

EMBASE

(SARS coronavirus/ or middle east respiratory syndrome/ or severe acute respiratory syndrome/ or (coronavirus* or corona virus* or HCoV* or ncov* or covid or covid19 or sars-cov* or sarscov* or Sars-coronavirus* or Severe Acute Respiratory Syndrome Coronavirus*).mp.) and 20191201:20301231.(dc).

Update #5: 30.10.2020

<u>EMBASE</u>

(exp SARS-related coronavirus/ or severe acute respiratory syndrome/ or coronavirus disease 2019/ or (coronavir* or corona virus* or HCoV* or ncov* or 2019 cov or covid or covid19 or sars-cov* or sars-coronavirus* or Severe Acute Respiratory Syndrome Coronavirus* or nCoV).mp.) and 20191101:20301231.(dc).

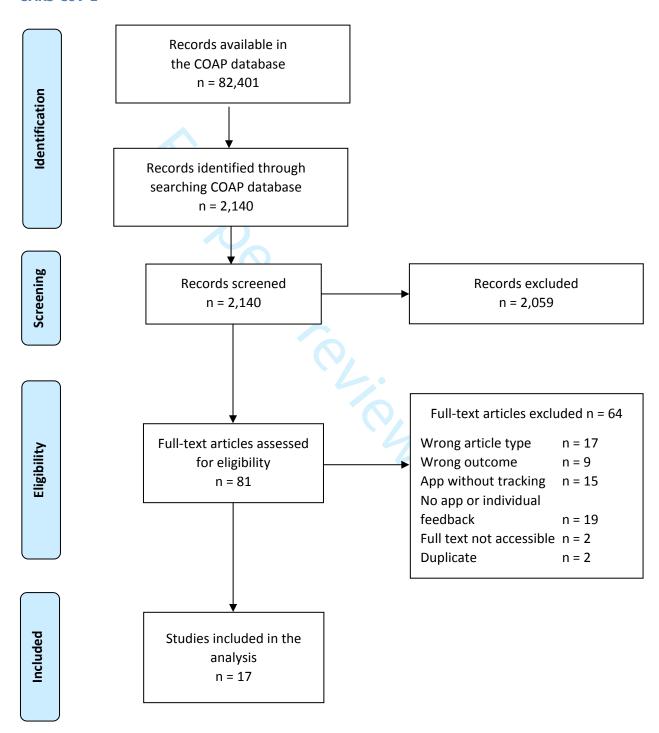
MEDLINE

("severe acute respiratory syndrome coronavirus 2"[Supplementary Concept] OR "COVID-19" [Supplementary Concept] OR "coronavirus" OR "corona virus" OR "HCoV" OR "nCoV" OR "2019 CoV" OR "covid" OR "covid19" OR "Severe Acute Respiratory Syndrome Coronavirus 2" OR "SARS-CoV2" OR "SARS-CoV 2" OR "SARS Coronavirus 2") AND (2019/11/01:3000/12/31[PDAT])



Supplementary file 2. Flowchart study selection

Flowchart regarding selection of studies looking at effectiveness of contact- and tracing apps for SARS-CoV-2



Supplementary file 3. Excluded studies

Studies not meeting inclusion criteria after full text screening, and excluded from analyses (n=64)

Reference	Reason for exclusion
Aleta 2020	No app or individual feedback
Aleta 2020	No app or individual feedback
Ayres 2020	Wrong outcome
Bian 2020	Wrong article type
Bianconi 2020	Full text not accessible
Braithwaite 2020	Wrong article type
Braithwaite 2020	Duplicate
Braun 2020	Full text not accessible
Brooks-Pollock 2020	No app or individual feedback
Chan 2020	Wrong article type
Chen 2020	No app or individual feedback
Di Domenico 2020	No app or individual feedback
Drake 2020	Wrong article type
Drew 2020	App without tracking
Fateh-Moghadam 2020	App without tracking
Fenton 2020	Wrong outcome
Firth 2020	No app or individual feedback
Gozzi 2020	App without tracking
Grantz 2020	Wrong outcome
Güemes 2020	App without tracking
Haller 2020	Wrong article type
Huang 2020	Wrong outcome
Hussein 2020	No app or individual feedback
Jian 2020	Wrong outcome
Kassaye 2020	App without tracking
Kendall 2020	Duplicate
Khataee 2020	Wrong article type
Kogan 2020	Wrong outcome
Kretzschmar 2020	Duplicate
Lambert 2020	Wrong article type
Leith 2020	Wrong article type
Liu 2020	No app or individual feedback
Maghdid 2020	Wrong article type
Marín-García 2020	Wrong article type
Menni 2020	App without tracking
Menni 2020	App without tracking
L	

Milenkovic 2020	No app or individual feedback
Mishra 2020	App without tracking
Morley 2020	No app or individual feedback
Nagarajan 2020	No app or individual feedback
Ni Lochlainn 2020	App without tracking
Pépin 2020	Wrong outcome
Petrellis 2020	Wrong article type
Ranjan 2020	App without tracking
Ruediger 2020	No app or individual feedback
Salathe 2020	Wrong outcome
Sattler 2020	Wrong article type
Serafino 2020	App without tracking
Sun 2020	App without tracking
Sun 2020	No app or individual feedback
Szocska 2020	No app or individual feedback
Unwin 2020	No app or individual feedback
Vannoni 2020	No app or individual feedback
Varsavsky 2020	No app or individual feedback
Vinceti 2020	App without tracking
Wallentin 2020	Wrong article type
Whaiduzzaman 2020	Wrong article type
Wilson 2020	Wrong article type
Wong 2020	Wrong article type
Yabe 2020	No app or individual feedback
Yap 2020	Wrong outcome
Yasaka 2020	Wrong article type
Zens 2020	App without tracking
Zhan 2020	No app or individual feedback

Supplementary file 4. Method for critical appraisal of empirical studies

Method used for critical appraisal of empirical epidemiologic studies

Confounding

Have the authors identified all important confounding factors? Yes / No / Unclear

Were the identified confounding factors <u>adjusted</u> for in the design and/or analysis? Yes / No

/ Unclear

- Model-based adjustment of confounders
- Stratification
- Matching
- No adjustment required (randomization)

Selection bias

Was patient exposure / intervention status at inclusion likely to result in bias? Yes / No /

Unclear

- Non-randomized study
- Randomized study with issues regarding allocation concealment or non-random sequencing
- Stringent exclusion criteria

Was missing data or loss to follow-up during the study likely to result in bias? Yes / No /

Unclear

- Missingness likely not completely at random (i.e. not MCAR or % of missingness different between groups)
- No methods described for handling missingness (i.e. imputation)
- Other methods explored to prevent missingness (i.e. cross checking data sources)

Informationbias

Was measurement of exposure / administration of the intervention likely to result in bias? Yes

/ No / Unclear

- Blinding
- Standardization
- Objective
- Non-compliance
- Breaking protocol

Was measurement of outcome likely to result in bias? Yes / No / Unclear

- Blinding
- Standardization
- Objective (note: if this is the case item should be scored 'No')

Other concerns? FREE TEXT

Items to consider (but not limited to)

- Reporting bias
- Conflict of interest

Supplementary file 5. Method for critical appraisal of model-based studies

Method used for critical appraisal of model based studies

Were empirical distributions used for a varying infectiousness since time of infection?

Yes / No / Unclear

Keywords indicating distributions were used

- Weibull
- Log-normal
- Exponential distribution

Were various different scenarios evaluated for important model assumptions and parameter values? Yes / No / Unclear

Keywords indicating uncertainty was taken into account

- Sensitivity analysis
- Scenario analysis

Were models reported transparently? (i.e. no black box) Yes / No / Unclear

Key elements indicating that model can be reproduced

- (differential) Equation specified
- Behavior of agents specified
- Graphic representation of model
- All variables and distributions specified

Other concerns? FREE TEXT

Items to consider (but not limited to)

- Reporting bias
- Conflict of interest
- Illogical properties of the model not captured by the criteria above



PRISMA 2009 Checklist

Section/topic	#	Checklist item	Reported on page #
TITLE			
Title	1	Identify the report as a systematic review, meta-analysis, or both.	1
ABSTRACT			
Structured summary	2	Provide a structured summary including, as applicable: background; objectives; data sources; study eligibility criteria, participants, and interventions; study appraisal and synthesis methods; results; limitations; conclusions and implications of key findings; systematic review registration number.	3
INTRODUCTION			
Rationale	3	Describe the rationale for the review in the context of what is already known.	4
Objectives	4	Provide an explicit statement of questions being addressed with reference to participants, interventions, comparisons, outcomes, and study design (PICOS).	4-5
METHODS			
Protocol and registration	5	Indicate if a review protocol exists, if and where it can be accessed (e.g., Web address), and, if available, provide registration information including registration number. Given the rapid nature of this systematic review, no protocol was registered beforehand	-
Eligibility criteria	6	Specify study characteristics (e.g., PICOS, length of follow-up) and report characteristics (e.g., years considered, language, publication status) used as criteria for eligibility, giving rationale.	5
Information sources	7	Describe all information sources (e.g., databases with dates of coverage, contact with study authors to identify additional studies) in the search and date last searched.	5, 28
Search	8	Present full electronic search strategy for at least one database, including any limits used, such that it could be repeated.	6, 28-30
Study selection	9	State the process for selecting studies (i.e., screening, eligibility, included in systematic review, and, if applicable, included in the meta-analysis).	6
Data collection process	10	Describe method of data extraction from reports (e.g., piloted forms, independently, in duplicate) and any processes for obtaining and confirming data from investigators.	6-7
Data items	11	List and define all variables for which data were sought (e.g., PICOS, funding sources) and any assumptions and simplifications made.	6-7
Risk of bias in individual studies	12	Describe methods used for assessing risk of bias of individual studies (including specification of whether this was done at the study or outcome level), and how this information is to be used in any data synthesis.	6, 34, 35
Summary measures	13	State the principal summary measures (e.g., risk ratio, difference in means).	6-7

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8

PRISMA 2009 Checklist

4 5	Synthesis of results	14	Describe the methods of handling data and combining results of studies, if done, including measures of consistency (e.g., I²) for each meta-analysis.	7	
٠ ١				1	

Page 1 of 2 Reported Section/topic # **Checklist item** on page # Risk of bias across studies 15 Specify any assessment of risk of bias that may affect the cumulative evidence (e.g., publication bias, selective reporting within studies). Additional analyses 16 Describe methods of additional analyses (e.g., sensitivity or subgroup analyses, meta-regression), if done, indicating which were pre-specified. **RESULTS** Give numbers of studies screened, assessed for eligibility, and included in the review, with reasons for exclusions at 7. 31-33 Study selection each stage, ideally with a flow diagram. Study characteristics For each study, present characteristics for which data were extracted (e.g., study size, PICOS, follow-up period) and 7, 8, provide the citations. 16-24 Risk of bias within studies Present data on risk of bias of each study and, if available, any outcome level assessment (see item 12). 8, 25-27 Results of individual studies For all outcomes considered (benefits or harms), present, for each study: (a) simple summary data for each 9-12 intervention group (b) effect estimates and confidence intervals, ideally with a forest plot. Present results of each meta-analysis done, including confidence intervals and measures of consistency. Not Synthesis of results applicable, as only qualitative assessment was possible Risk of bias across studies Present results of any assessment of risk of bias across studies (see Item 15). Give results of additional analyses, if done (e.g., sensitivity or subgroup analyses, meta-regression [see Item 16]). Additional analysis DISCUSSION Summary of evidence 24 Summarize the main findings including the strength of evidence for each main outcome; consider their relevance to 12 key groups (e.g., healthcare providers, users, and policy makers). ³⁶ Limitations Discuss limitations at study and outcome level (e.g., risk of bias), and at review-level (e.g., incomplete retrieval of 12. 13 identified research, reporting bias). Conclusions Provide a general interpretation of the results in the context of other evidence, and implications for future research. 13 **FUNDING Funding** Describe sources of funding for the systematic review and other support (e.g., supply of data); role of funders for the 2 systematic review.

For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml
From: Moher D, Liberati A, Tetzlaff J, Altman DG, The PRISMA Group (2009). Preferred Reporting Items for Systematic Reviews and Meta-Analyses: The PRISMA Statement. PLoS Med 6(7): e1000097.

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Effectiveness of contact tracing apps for SARS-CoV-2: a rapid systematic review

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Title

Effectiveness of contact tracing apps for SARS-CoV-2: a rapid systematic review

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Competing interest statement

All authors have completed the Unified Competing Interest form (available on request from the corresponding author) and declare: no support from any organisation for the submitted work [or describe if any]; no financial relationships with any organisations that might have an interest in the submitted work in the previous three years, no other relationships or activities that could appear to have influenced the submitted work

Transparency declaration

The lead author affirms that the manuscript is an honest, accurate, and transparent account of the study being reported; that no important aspects of the study have been omitted.

Ethical approval

No ethical approval was required for this study

Patient and public involvement statement

No patients or public were involved in the conception and execution of this study

Dissemination statement

We do not plan to further disseminate the results of this study apart from its publication

Data sharing statement

All data relevant to the study are included in the article or uploaded as supplementary information

Abstract

Objective – To systematically review evidence on effectiveness of contact tracing apps (CTAs) for SARS-CoV-2 on epidemiological and clinical outcomes

Design – Rapid systematic review

Data sources - EMBASE (OVID), MEDLINE (PubMed), BioRxiv, and MedRxiv were searched up to October 28th 2020

Study selection – Studies, both empirical and model-based, assessing effect of CTAs for SARS-CoV-2 on reproduction number (R), total number of infections, hospitalization rate, mortality rate, and other epidemiologically and clinically relevant outcomes, were eligible for inclusion.

Data extraction – Empirical and model-based studies were critically appraised using separate checklists. Data on type of study (i.e. empirical or model-based), sample size, (simulated) time horizon, study population, CTA type (and associated interventions), comparator, and outcomes assessed, were extracted. The most important findings were extracted and narratively summarized. Specifically for model-based studies, characteristics and values of important model parameters were collected.

Results – 2140 studies were identified, of which 17 studies (two empirical, 15 model-based studies) were eligible and included in this review. Both empirical studies were observational (non-randomized) studies and at high risk of bias, most importantly due to risk of confounding. Risk of bias of model-based studies was considered low for 12 of 15 studies. Most studies demonstrated beneficial effects of CTAs on R, total number of infections, and mortality rate. No studies assessed effect on hospitalization. Effect size was dependent on model parameters values used, but in general a beneficial effect was observed at CTA adoption rates of 20% or higher.

Conclusions – Contact tracing apps have the potential to be effective in reducing SARS-CoV-2 related epidemiological and clinical outcomes, though effect size depends on other model parameters (e.g. proportion of asymptomatic individuals, or testing delays), and interventions after CTA notification. Methodologically sound comparative empirical studies on effectiveness of CTAs are required to confirm findings from model-based studies.

Strengths and limitations of this study

- This is the first paper to provide a comprehensive overview and critical appraisal of studies assessing the effectiveness of contact tracings apps for SARS-CoV-2 on clinical and epidemiological outcomes
- Studies were retrieved using a large repository that is developed by a specific search string dedicated to identify studies on SARS-CoV-2 published in various underlying databases
- Critical appraisal was performed by reviewers from diverse backgrounds (i.e. mathematical modelling, epidemiology, medicine, systematic reviews) using predefined customized templates for both empirical and model-based effectiveness studies
- Given the rapid execution and (preprint) publication of studies on effectiveness of contact tracing apps for SARS-CoV-2, this review is unlikely to include the most recent studies that published after the search date
- Due to too much heterogeneity across studies, it was not feasible to provide a
 pooled meta-analysis estimate of the effectiveness of contact tracing apps for
 SARS-CoV-2 on the clinical and epidemiological outcomes

Introduction

The severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) outbreak has dominated worldwide news and scientific research throughout 2020. Since the outbreak in Wuhan (People's Republic of China) in early December 2019, reducing transmission of SARS-CoV-2 has been a worldwide priority. Digital technology could be applied for efficient contact tracing. Contact tracing applications (CTAs) are able to identify individuals who have recently been in close contact with infected individuals (and may have acquired infection as a consequence). After identification, the contact person can be instructed to go in self-quarantine, preventing further transmission and spread of the virus.

A substantial amount of research on CTAs for SARS-CoV-2 has been performed since the start of the pandemic. Summarizing all evidence, including results from research that has not yet, or is currently undergoing peer-review, is warranted to provide an overview of what is known regarding CTA effectiveness. Research that has not yet undergone peer-review is often published by authors through so-called preprint databases. However, identifying these articles, extracting data, and drawing conclusions can be a challenge, as this requires knowledge on epidemiology, mathematical modelling, systematically appraising evidence, and summarizing that evidence.

A few overviews of evidence on effectiveness of CTAs have been published in recent time. Anglemyer *et al.* provided an overview of study characteristics and quality appraisal of studies on effectiveness of CTAs and other digital contact tracing technologies. (1) However, their data are based on both SARS-CoV-2 infections and other infections (e.g. Ebola), and lack a quantitative effectiveness measure of CTAs on clinically relevant outcomes. Other systematic reviews focused only on user experience in using a CTA for SARS-CoV-2 (2), or only studied manual as opposed to digital contact tracing (3). One systematic review did look into studies on automated and semi-automated CTAs for SARS-CoV-2, but lacked reporting on CTA effectiveness on total number of infections, and hospitalization or mortality rates. (4)

In this rapid systematic review, we aim to evaluate all (empirical and model based) studies addressing effectiveness of CTAs for SARS-CoV-2 on relevant, i.e. epidemiological and

clinical, outcomes. We will provide descriptive characteristics, critical appraisal, and a narrative summary of evidence of included studies.

Methods

Search strategy

The *Bern COVID-19 Open Access Project (COAP)* database was used for identification of relevant research. The COAP database is comprised of research from EMBASE (OVID), MEDLINE (PubMed), BioRxiv en MedRxiv databases, specifically focused on SARS-CoV-2. On October 28th 2020 the COAP database was searched for scientific literature evaluating the effectiveness of CTAs for SARS-CoV-2 on epidemiological and clinical outcomes. The complete search strategy, as well as background information on the COAP database provided by Bern University, are provided in Supplementary File 1.

Eligibility criteria

Empirical (both observational and experimental) and model-based studies evaluating effectiveness of CTAs for SARS-CoV-2 were eligible for inclusion. Peer-reviewed publications as well as preprint papers were considered.

CTAs were considered when they provided feedback about potential recent exposure to an infected individual, based on proximity measurements (e.g. Bluetooth or GPS). Feedback should be provided directly to the individual through a CTA, although other feedback mechanisms, such as personal devices (e.g. a smartwatch), were also considered. National emergency warning systems using SMS were also included, provided they used proximity data to inform individuals.

All epidemiologically or clinically relevant outcomes quantifying the impact of CTAs were considered, which include but are not limited to: the reproduction number (R), total number of infections, hospitalization rate, and mortality rate related to SARS-CoV-2. Studies investigating other relevant outcomes, such as prevention of outbreaks or a second infection wave of SARS-CoV-2, were also included. Studies solely assessing (determinants affecting) adoption rate of CTAs (i.e. the proportion of citizens using, and following recommendations provided by, the CTA), temporal change in incidence SARS-CoV-2, or other non-epidemiological or clinical outcomes were excluded.

Study selection

Studies identified in the search were first screened independently on title and abstract by two reviewers. Relevant studies were included for full text screening, and further selection of articles was performed by two independent reviewers. Any discrepancies were discussed and resolved. When consensus was not reached, a third reviewer was consulted to provide the final judgement.

Critical appraisal

Risk of bias was systematically assessed by two researchers using separate checklists for empirical and model-based studies. Discrepancies between researchers were discussed, and a final verdict was provided by a third reviewer if consensus was not reached. Empirical studies were appraised using a formal scoring method based on the Critical Appraisal Skills Programme (CASP) and Cochrane's Effective Practice and Organisation of Care (EPOC) checklists (5, 6) (Supplementary file 2). Risk of bias in model-based research was evaluated by assessing use of empirical input data for the model, number of scenarios analyzed, and transparency of model reporting. (Supplementary file 3)

Data extraction

Data extraction was performed by one reviewer, and checked by a second reviewer. Descriptive characteristics on type of research, i.e. empirical or model-based, sample size, (simulated) time horizon, study population, CTA properties and intervention, comparator, and epidemiological and clinical outcomes studied, were extracted from all included studies.

Specifically for model-based research, model characteristics (i.e. type of model and distributions used) and values used for important model parameters were collected. Furthermore, CTA specific properties were extracted, such as the method of contact tracing used by these apps. Forward tracing CTAs can only detect the 'offspring', i.e. individuals the index case has infected, of an infected individual. Bidirectional tracing CTAs also detect the 'parents', i.e., the individual that infected the index case of an infected individual. Models were considered to use bidirectional (as opposed to forward) tracing when, after the index case is detected and registered, all contacts within a period of at least the incubation time are identified, such that the parent of the index case could be found.

Another CTA specific property included the use of 1-step-tracing or sequential tracing. When a CTA-identified individual could only notify their contacts after testing positive themselves, this was considered 1-step-contact tracing. When notified contacts could subsequently also notify their own contacts, creating a cascade, even before that individual has shown symptoms or received a positive test result for SARS-CoV-2, this was considered sequential tracing.

The most important findings regarding effectiveness of CTAs for SARS-CoV-2 on epidemiological and clinical outcomes were extracted, synthesized, and reported narratively. These outcomes were pooled quantitatively whenever it was feasible to do so.

Results

Study selection

A total of 2140 potential studies were identified by the search. After selection based on title and abstract, 2059 articles were excluded. Full texts of the 81 remaining studies were assessed, after which 17 articles were included for critical appraisal and data extraction (Supplementary file 4). The 64 excluded studies with their reasons for exclusion are summarized in Supplementary file 5.

Characteristics of included studies

Seventeen primary studies were included, of which two were empirical observational (non-randomized) studies, and 15 were model-based studies (Table 1).

Six of the 17 studies were published preprints, meaning they had not (yet) gone through the peer review process at the time of data extraction (7-12). Included studies focused predominantly on the general population, although some analyzed the effectiveness of CTAs for specific populations such as hospital personnel, or school children (8, 9, 11, 13-16). Especially in model-based studies, results were often presented graphically. Consequently, the effectiveness of CTAs on epidemiological and clinical outcomes was only partly, or not at all, reported in key numerical figures.

The model-based studies typically assessed the effectiveness of CTAs by simulating one or more scenarios based on certain baseline or input values (e.g. proportion of asymptomatic infections). Table 2 provides an overview of characteristics and the most important input parameters used in models of the 15 included articles. Nine of the 15 model-based studies evaluated forward tracing CTAs (8, 9, 11, 13-18), four studies

analyzed bidirectional tracing CTAs (7, 10, 12, 19), and one used an alternative method (20). Four studies used a CTA that used sequential tracing (7, 10, 12, 19). All of these also used bidirectional CTAs, which are more effective than forward tracing CTAs in reducing R, but require quarantining many more contact persons. This is especially the case when a significant number of infections come from asymptomatic individuals (i.e. transmission from a case who does not (yet) have symptoms), who are unaware they have SARS-CoV-2. (19)

The percentage of CTA adoption was varied in almost all studies, allowing for assessment of the impact of CTAs on epidemiological and clinical outcomes. Average incubation time, i.e. the mean time between infection and symptom onset of SARS-CoV-2, was estimated to be 5 to 6 days for SARS-CoV-2 (9, 11-21). The proportion of asymptomatic SARS-CoV-2 infections, used as input parameter in model-based studies, was estimated at 20% to 50% based on empirical data (8, 9, 16, 18), but could vary between 18% to 86% (9). The baseline R value chosen in the model-based studies varied between 1.2 and 4.0. (7-10, 12, 14-21)

Furthermore, so-called superspreaders (i.e. individuals that infect numerous other individuals, and consequently have a high individual R) were discussed in context of the SARS-CoV-2 pandemic. Tracing these superspreaders is key in containing outbreaks. Hence, it is warranted to use bidirectional CTAs to trace these superspreaders, and advise them to immediately enter quarantine on identification. (14, 22)

Critical appraisal

Risk of bias in the two empirical studies was judged to be high (Table 3) (23, 24). Confounding variables (such as smoking, work status, and income) were insufficiently taken into account given the explanatory and observational nature of these empirical studies. It was also unclear how missing (outcome) data were dealt with.

Most model-based research was judged to have a low risk of bias (Table 4). Three of the 15 studies had a high risk of bias due to the lack of use of empirical distributions for variables, the limited number of scenarios analyzed, and insufficient transparency regarding reporting of the model. (11, 20, 21)

Synthesis of results

Evidence from empirical studies

Two empirical comparative observational studies assessed the effectiveness of CTAs compared to a control group that did not use CTAs (Table 1). (23, 24) One study looked at effectiveness of a text warning system used in 627,386 individuals who came in contact with an exposed population, and compared it to the general population of Taiwan who did not use such a warning system. (17) They showed a reduction in incidence of respiratory syndrome from 19.23 to 16.87 per 1000 individuals. They also showed a reduction in pneumonia incidence from 3.81 to 2.36 per 1000 individuals. (17) The second observational study investigated the introduction and adoption of a 'Test and Trace' app by 34,000 individuals living on the Isle of Wight (UK), and compared the estimated value of R in that region to that in the general UK population. (24) The CTA marked individuals as positive based on self-reporting of symptoms. Individuals that came in contact with an individual marked as positive were provided with social distancing advice. The study found that R was reduced from 1.3 to 0.5 after implementation of the CTA. Within 2 to 3 weeks after implementation, incidence of SARS-CoV-2 diagnoses declined by around 90%. (24)

Evidence from model-based studies

Effect on R

Effectiveness of a 1-step-contact tracing in reducing R can be approached using the following formula:

$$R_c = R * (1 - p^2 * f)$$

Here, R_c is the reproduction number when a CTA is used, R is the reproduction number without the use of a CTA, p is the proportion of the population using the CTA, and f is the combination of other factors that affect effectiveness of notification by the CTA. Such factors include, but are not limited to, delay between CTA notification and testing, delay between testing and test result, delay between reception of test result and entry of that result in the CTA, compliance to interventions (e.g. self-quarantine), and the proportion of infections that occur pre- or asymptomatically. Note that p occurs as a quadratic term, which reflects the fact that both infector and infectee have to use the CTA for the transmission to get traced.

Nine of the 15 model-based studies assessed the effect of CTAs on reduction of R. (8, 11, 14-16, 18, 19, 21) CTAs were able to control an ongoing outbreak or epidemic through quicker and more efficient feedback of a positive test result, and by notifying close contacts of a positively tested individual. (15, 16, 19) This speed and efficiency were not feasible using traditional manual contact tracing. (16) New outbreaks could be controlled (i.e. R_c <1.0) by CTAs, by combining them with quarantine or self-isolation interventions, provided that hygiene and social distancing measures are maintained. (8, 14, 18, 21) CTAs were able to reduce R by 0.3 more than traditional manual contact tracing, provided that feedback about contact with a positively tested individual is given to all contacts of the index case of the preceding 7 days. (19) Another model-based study demonstrated that a CTA with 20% adoption rate reduces R by 17.6% compared to no contact tracing, whereas traditional manual contact tracing reduced R by 2.5% compared to no contact tracing. (15) This study also demonstrated that a CTA is able to reduce the R further, even when social distancing has already reduced R to 1.2. In this situation, R can be reduced further by 30% to 0.8 when CTA adoption rate is 80%. (15) Another model-based study determined that 60% adoption rate of a CTA could result in an R below 1.0. (11) In one study, adoption rate of 53% resulted in a 47% reduction in R when the complete household of an individual with a positive test result is advised to be guarantined. (14) The last study looking at effect of CTA on R showed that only at 60% adoption rate of the app a significant beneficial effect on R would become apparent. (12) When R is high (e.g. 3.0), and a considerable proportion of individuals is asymptomatic (e.g. 40% of all infections), CTAs need to be combined with other interventions (such as social distancing and random testing) to be able to lower the R below 1.0. (12) Potential for CTAs to reduce R is not only dependent on the adoption rate of the app, but also on (effectiveness of) various other measures that are provided after a positive notification, the delay between positive notification and opportunity for testing, and delay between receiving a positive test result and sharing that result through the CTA. (5, 6, 10) One study found that the percentage of preventable infections by one individual strongly depends on the time delay between CTA notification and the ability to be tested. (15) When there was no delay (i.e. 0 days) 79.9% of infections could be prevented, compared to 41.8% and 4.9% for 3 and 7 days delay respectively.

Effect on total number of infections

Eight of the 15 model-based studies assessed the effect of CTAs on reducing the total number of infections. (8-11, 13, 17, 18, 20) Two studies indicated that the success of CTAs

in reducing the total number of infections could only be ensured with a high adoption rate of that app. (8, 13) Another study showed that with a high CTA adoption rate of 75%, there would be no more new infections occurring within three months after implementation. (11) It was found that adequate hygiene and social distancing measures are needed to enable CTAs to reduce the total number of infections. (8, 9, 17, 18) Especially in areas where there is low compliance to social distancing, a sufficiently high adoption rate of a CTA is essential to maintain control of an outbreak. (9)

The height of the peak number of new infections can, according to one study, be reduced by half with a 50% adoption rate of a CTA (18), whereas another study showed that this could be achieved with an adoption rate as low as 20%. (20) Another study demonstrated that at 27% CTA adoption rate, a quarter of all new infections can be prevented. (17) However, according to another study that used a similar adoption rate, the number of infections would stabilize, but the epidemic would be maintained by core groups in densely populated areas. (18) There may be a period of time of more than two months between implementation of interventions (such as CTAs) and the effect of that implementation on the total number of SARS-CoV-2 infections. (13)

Effect on number of hospitalizations

None of the 15 model-based studies assessed the effect of CTAs on the number of hospitalizations due to SARS-CoV-2 infection, possibly because the number of hospitalizations is expected to be proportional to the number of infections, only with a time-delay. A German study did look into the effect of a CTA on the number of days that intensive care unit (ICU) capacity was exceeded. (9) They found in their simulations that – based on the German population, and assuming an ICU capacity of 24.000 beds – a CTA adoption rate of 20% would prevent exceedance of ICU capacity at any point in time. In contrast, if no contact tracing (either manual or digital) would be used, ICU capacity would be exceeded on a quarter of days.

Effect on mortality rate

Three of the 15 model-based studies assessed the effect of CTAs on mortality rate. (9, 18, 20) One study demonstrated that a high adoption rate (80%) of a CTA would result in an 85% reduction in mortality rate, over a period of 500 days (9). Another study found that a low CTA adoption rate (25%) is associated with a 10% decrease in mortality rate, an average adoption rate (50%) with 25% decrease, and a high adoption rate (75%) with 40-

60% decrease. (18) A third study showed that at 40% adoption rate, during the peak of an outbreak, a reduction in number of deaths by 97% could be achieved. (20)

Discussion

Empirical evidence regarding the effectiveness of using CTAs for detection of SARS-CoV-2 is still limited. Currently, no randomized studies have been performed, and only two observational comparative studies were identified in this systematic review. Although some benefits of using CTAs for detection of SARS-CoV-2 were observed, both studies were deemed to be of low methodological quality. However, the results of these studies were in accordance with the 15 included, higher quality, model-based studies assessing effectiveness of CTAs. These studies showed that CTAs can be effective and a valuable addition to manual contact tracing. CTA use resulted in a lower R, lower total number of infections, and lower mortality rate. These reductions were already observed at relatively low adoption rates (e.g. 20%), though higher adoption rates of CTAs resulted in greater reductions. Shortening delays between CTA notification and diagnostic testing may increase its effectiveness.

Strengths & Limitations

This rapid systematic review assesses key features, quality, and main clinical and epidemiological outcomes of a set of studies, both empirical and model-based, on effectiveness of CTAs for SARS-CoV-2. To our knowledge, no such systematic review has been published, assessing these specific properties. Methodological quality of empirical studies was assessed using standardized tools. No such tool was available in literature for model-based studies, and as such a set of key features used in other systematic reviews on this topic was used. This set was validated by experts in mathematical modelling.

To fully appreciate the findings from this systematic review, some considerations should be taken into account. First, the studies found through the literature search may not be a comprehensive set. Studies on SARS-CoV-2 are published at a rapid, almost daily, basis in various online repositories. Although we cannot ensure that all studies on the effectiveness of CTAs for SARS-CoV-2 have been identified, we believe that the set of included studies that we have identified is a representative sample.

Furthermore, effectiveness of CTAs for SARS-CoV-2 described in model-based studies is complex. Numerous input variables used in the models interact with one another, and consequently affect effectiveness of, for example, adoption rate of CTAs on clinical or epidemiological outcomes. Summarizing these findings into a general effectiveness is difficult, and will always suffer from simplification of a system of complex interactions. Though we feel that providing some (conditional) findings from these studies will help provide some general insight in the impact CTAs can have on clinical and epidemiological outcomes for SARS-CoV-2.

Conclusion & implications for further research

Current evidence on the effectiveness of CTAs for SARS-CoV-2 is predominantly based on modelling studies, which indicate that there is potential in beneficially affecting key clinical and epidemiological outcomes. High quality empirical evidence, either from experimental or methodologically sound observational studies, is needed in order to be able to draw more robust conclusions regarding effectiveness of CTAs for SARS-CoV-2.

Contributorship statement

Conception and design: KJ, KGMM, LH

Planning: KJ, MCJB, JAAGD, MSO, RWMV, RS, KGMM, MEEK, LH

Acquisition of data: RS

Conduct: KJ, MCJB, JAAGD, MSO, RWMV, RS, KGMM, MEEK, LH

Analysis: KJ, MCJB, KGMM, MEEK, LH

interpretation of data: KJ, MCJB, KGMM, MEEK, LH

Reporting: KJ, MCJB, JAAGD, MSO, RWMV, RS, KGMM, MEEK, LH

References

- 1. Anglemyer A, Moore TH, Parker L, Chambers T, Grady A, Chiu K, et al. Digital contact tracing technologies in epidemics: a rapid review. Cochrane Database Syst Rev. 2020;8:CD013699.
- 2. Davalbhakta S, Advani S, Kumar S, Agarwal V, Bhoyar S, Fedirko E, et al. A Systematic Review of Smartphone Applications Available for Corona Virus Disease 2019 (COVID19) and the Assessment of their Quality Using the Mobile Application Rating Scale (MARS). J Med Syst. 2020;44(9):164.
- 3. Nussbaumer-Streit B, Mayr V, Dobrescu AI, Chapman A, Persad E, Klerings I, et al. Quarantine alone or in combination with other public health measures to control COVID-19: a rapid review. Cochrane Database Syst Rev. 2020;4:CD013574.
- 4. Braithwaite I, Callender T, Bullock M, Aldridge RW. Automated and partly automated contact tracing: a systematic review to inform the control of COVID-19. Lancet Digit Health. 2020;2(11):e607-e21.
- 5. CASP Checklist [Internet]. Critical appraisal skills programme. 2018 [cited 15-10-2020]. Available from: https://casp-uk.net/wp-content/uploads/2018/01/CASP-Cohort-Study-Checklist 2018.pdf.
- 6. EPOC Resources for review authors [Internet]. Cochrane Effective Practice and Organisation of Care (EPOC). 2017 [cited 15-10-2020]. Available from: https://epoc.cochrane.org/resources/epoc-resources-review-authors.
- 7. Bulchandani VB, Shivam S, Moudgalya S, Sondhi SL. Digital Herd Immunity and COVID-19. medRxiv. 2020:2020.04.15.20066720.
- 8. Cencetti G, Santin G, Longa A, Pigani E, Barrat A, Cattuto C, et al. Digital Proximity Tracing in the COVID-19 Pandemic on Empirical Contact Networks. medRxiv. 2020:2020.05.29.20115915.
- 9. Grimm V, Mengel F, Schmidt M. Extensions of the SEIR Model for the Analysis of Tailored Social Distancing and Tracing Approaches to Cope with COVID-19. medRxiv. 2020:2020.04.24.20078113.
- 10. Guttal V, Krishna S, Siddharthan R. Risk assessment via layered mobile contact tracing for epidemiological intervention. medRxiv. 2020:2020.04.26.20080648.
- 11. Shamil MS, Farheen F, Ibtehaz N, Khan IM, Rahman MS. An Agent Based Modeling of COVID-19: Validation, Analysis, and Recommendations. medRxiv. 2020:2020.07.05.20146977.
- 12. Pollmann TR, Pollmann J, Wiesinger C, Haack C, Shtembari L, Turcati A, et al. The impact of digital contact tracing on the SARS-CoV-2 pandemic a comprehensive modelling study. medRxiv. 2020:2020.09.13.20192682.
- 13. Scott N, Palmer A, Delport D, Abeysuriya R, Stuart RM, Kerr CC, et al. Modelling the impact of relaxing COVID-19 control measures during a period of low viral transmission. Med J Aust. 2021;214(2):79-83.
- 14. Kucharski AJ, Klepac P, Conlan AJK, Kissler SM, Tang ML, Fry H, et al. Effectiveness of isolation, testing, contact tracing, and physical distancing on reducing transmission of SARS-CoV-2 in different settings: a mathematical modelling study. Lancet Infect Dis. 2020;20(10):1151-60.
- 15. Kretzschmar ME, Rozhnova G, Bootsma MCJ, van Boven M, van de Wijgert J, Bonten MJM. Impact of delays on effectiveness of contact tracing strategies for COVID-19: a modelling study. Lancet Public Health. 2020;5(8):e452-e9.
- 16. Ferretti L, Wymant C, Kendall M, Zhao L, Nurtay A, Abeler-Dörner L, et al. Quantifying SARS-CoV-2 transmission suggests epidemic control with digital contact tracing. Science. 2020;368(6491):eabb6936.
- 17. Currie DJ, Peng CQ, Lyle DM, Jameson BA, Frommer MS. Stemming the flow: how much can the Australian smartphone app help to control COVID-19? Public Health Res Pract. 2020;30(2).
- 18. Ferrari A, Santus E, Cirillo D, Ponce-de-Leon M, Marino N, Ferretti MT, et al. Simulating SARS-CoV-2 epidemics by region-specific variables and modeling contact tracing app containment. NPJ Digit Med. 2021;4(1):9.

- 19. Bradshaw WJ, Alley EC, Huggins JH, Lloyd AL, Esvelt KM. Bidirectional contact tracing could dramatically improve COVID-19 control. Nat Commun. 2021;12(1):232.
- 20. Nuzzo A, Tan CO, Raskar R, DeSimone DC, Kapa S, Gupta R. Universal Shelter-in-Place Versus Advanced Automated Contact Tracing and Targeted Isolation: A Case for 21st-Century Technologies for SARS-CoV-2 and Future Pandemics. Mayo Clin Proc. 2020;95(9):1898-905.
- 21. Kurita J, Sugawara T, Ohkusa Y. Estimated effectiveness of school closure and voluntary event cancellation as COVID-19 countermeasures in Japan. J Infect Chemother. 2021;27(1):62-4.
- 22. Endo A, Leclerc QJ, Knight GM, Medley GF, Atkins KE, Funk S, et al. Implication of backward contact tracing in the presence of overdispersed transmission in COVID-19 outbreak. medRxiv. 2020:2020.08.01.20166595.
- 23. Chen CM, Jyan HW, Chien SC, Jen HH, Hsu CY, Lee PC, et al. Containing COVID-19 Among 627,386 Persons in Contact With the Diamond Princess Cruise Ship Passengers Who Disembarked in Taiwan: Big Data Analytics. J Med Internet Res. 2020;22(5):e19540.
- 24. Kendall M, Milsom L, Abeler-Dorner L, Wymant C, Ferretti L, Briers M, et al. Epidemiological changes on the Isle of Wight after the launch of the NHS Test and Trace programme: a preliminary analysis. Lancet Digit Health. 2020;2(12):e658-e66.



Table 1. Descriptive characteristics of included studies

Characteristics of empirical epidemiological and model-based studies looking at effectiveness of contact- and tracing apps for SARS-CoV-2. N/R = 1 not reported, R = 1 reproduction number, $R_0 = 1$ baseline reproduction number

3	Study	Country	Study	Sample size /	Time	Population	Specific	Intervention	Comparison	Outcome(s)	Main findings
,		(of first	type	# of	horizon	· opanacion	setting(s)				
0		author)	.,,,,	simulations			55tm.B(5)				
11	Bradshaw	Germany	Modelling	500 or 1000	52 weeks	General	-	Contact tracing	- Manual	- R	- Bidirectional tracing will enable more
12	2020			simulations	or 10,000	population		app (Bluetooth)	contact	- Outbreak	effective control of COVID-19
13	(peer				cases			with quarantine	tracing	control	- Switching from forward to bidirectional
14	reviewed)								- Current		tracing can reduce R by 0.3 if the tracing
15						/			practice		time window is sufficiently wide
16											,
17											- High adoption of bidirectional manual and digital contact tracing is 3x more
18											effective at outbreak control compared to
19											current practice
20						4					•
21	Bulchandani	USA	Modelling	4000	N/R	Susceptible	- / 🔘	Contact tracing	-	- R	- Outbreak control is possible regardless of
22	2020			simulations		population		app (not		- Outbreak	proportion of asymptomatic transmission
24	(preprint)					(i.e. no	4	specified) with		control	- Outbreak control requires a contact
25						immunity)		quarantine			tracing app adoption of 75%-95%
26	Cencetti	Italy	Madalling	20	FO days	General	Linivarsity	Contact tracing		- R	- Reduction of R and outbreak control is
27	2020	Italy	Modelling	simulations	50 days		- University	Contact tracing app (Bluetooth)	_	- N - Outbreak	dependent on contact tracing efficiency,
28	(preprint)			Silliulations		population	Campus - High school	with quarantine		control	isolation efficiency, and R_0
29	(ргерине)						- Workplace	with quarantine		Control	,
30											- Outbreak control can be achieved
31											through tracing and isolation, provided
32											that hygiene and social distancing
33											measures limit R ₀ to 1.5
34											- Outbreak control not feasible if contact
35											tracing app adoption is insufficient or if R ₀
36											is >2
37	Chen 2020	Taiwan	Empirical	3000	40 days	General	-	Public Warning	Current	- Respiratory	- Contact tracing and SMS feedback
88	(peer		·	individuals	<i>'</i>	population		System SMS	practice	syndrome	resulted in less cases of respiratory
39	reviewed)					(Taiwan)		(GPS) with		- Pneumonia	syndrome (16.87 vs 19.23 per 1000) and
10 11								quarantine &			
· I -											

· _											
•								symptom			pneumonia (2.36 vs 3.81 per 1000)
								monitoring			compared to the general population
											- Resource requirements for manual
'											contact tracing could be reduced by using
'											contract tracing apps combined with big
;											data analytics
)											uata allalytics
0	Currie 2020	Australia	Modelling	Not reported	12	General	-	COVIDSafe	No contact	- Outbreak	- Outbreak control by a contact tracing
1	(peer				months	population		contact tracing	tracing app	control	app can be achieved when adoption is
	reviewed)					(Australia)		app (Bluetooth)		- Cumulative	sufficient, and is combined with testing
3								with quarantine		incidence	and social distancing
4					4					SARS-CoV-2	- Cumulative incidence of SARS-CoV-2 can
5						/					within 8 months (depending social
6											distancing and testing intensity) be
7											reduced to:
8											
9											o 13-24% at an app adoption of 27%
0						1					o 17-35% at an app adoption of 40%
11											o 36-59% at an app adoption of 61%
2											o 47-76% at an app adoption of 80%
3 –	Ferrari 2020	Italy	Modelling	5500	50 days	General	-	Contact tracing	-	- R	- Reduction of R below 1.0 can be
4	(peer			simulations	300 days	population		app (not		- Outbreak	achieved when contact tracing apps have
	reviewed)			(per scenario)	400 days	(Italy)		specified) with		control	sufficient adoption, efficacy of case
6								quarantine &		- Cumulative	identification, and compliance to
7								symptom		incidence	quarantine
8								monitoring	UA	SARS-CoV-2	- Outbreak control can be achieved using
9										(symptomatic	contact tracing apps combined with
0)	voluntary self-quarantine and efficient
1										- Mortality	case isolation, depending population
3											density and transportation
4											
5											- Outbreak control was achieved with 75%
6											app adoption rate
7											- Cumulative incidence can be suppressed
8											with 25% app adoption rate, but
- 1											outbreaks will be sustained by districts
9										l	-
9											with high population density

2											
3 4 5 6 7											 Mortality was reduced by: 10% at 25% app adoption rate 25% at 50% app adoption rate 40-60% at 75% app adoption rate
8 9 10 11 12 13 14 15	Ferretti 2020 (peer reviewed)	China	Modelling	40 simulations (pairs)	12 days 20 days	General population (China)	- Home - Train - Work	Contact tracing app (Bluetooth) with quarantine	Manual contact tracing	R	- Manual contact tracing is not able to stop outbreaks due to delays (~ 3 days), whereas contact tracing apps are able to prevent outbreaks - Reduction of R below 1.0 is feasible using instantaneous (red. without delays) contact tracing apps
16 17 18 19 20 21 22 23 24 25 26	Grimm 2020 (preprint)	Germany	Modelling	N/R	500 days	General population (Germany)	- High risk of severe course of infection - Low risk of severe course of infection	Contact tracing app (not specified) with quarantine	- No intervention - Uniform social distancing - Group specific social distancing	- Cumulative incidence SARS-CoV-2 - # of days ICU capacity exceeded - Mortality	- ICU capacity and mortality can be kept low by using contact tracing apps combined with tailored social distancing and personal protection measures - ICU capacity was not exceeded at any point with a contact tracing app adoption of 20% or more - Mortality was reduced by 85% when a high (80%) adoption rate of the contact tracing app was achieved
27 28 29 30 31 32 33 34 35 36	Guttal 2020 (preprint)	N/R	Modelling	N/R	150-200 days	General population	-	Contact tracing app (Bluetooth) with quarantine	0/1/	Cumulative incidence SARS-CoV-2	- Peak cumulative incidence can be flattened significantly even when a small fraction of cases are identified using contact tracing apps, tested and isolated - Peak cumulative incidence can strongly be reduced even if contact tracing app testing is only performed in the most probable individuals (p > 0.8)

2											
3 4 5 6 7 8 9	Kendall 2020 (peer reviewed)	United Kingdom	Empirical Modelling	Population- size Isle of Wight Population- size UK (except Wales)	<2 months	General population (Isle of Wight and UK (except Wales))	- Close	NHS contact tracing app (version 1) (Bluetooth) with social distancing	- Social	- R - Cumulative incidence SARS-CoV-2	- Reduction of R from 1.3 to 0.5 was achieved after implementation of a contact tracing app - Cumulative incidence of SARS-CoV-2 reduced by 87% in 2-3 weeks after implementation of a contact tracing app - Contact tracing apps, with short delays
11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31	2020 (peer reviewed)	ds		simulations	04	population	contacts - Casual contacts	app (Bluetooth) with quarantine	distancing without contact		and high coverage for testing and tracing, could substantially reduce the R, alleviating more stringent control measures - Reduction of the R from 1.2 with social distancing alone to 0.8 (95% CI 0.7–1.0) by adding a contact tracing app with an adoption of 80% - Reduction of the R through contact tracing apps is more effective compared to manual contact tracing, with respectively 17.6% and 2.5% reduction of R compared to no contact tracing - Reduction in transmission rate (reflective of R) depends on tracing delay 0 79.9% with 0-day testing delay 0 41.8% with 3-day testing delay 0 4.9% with 7-day testing delay
32 33 34 35 36 37 38 39 40 41 42	Kucharski 2020 (peer reviewed)	United Kingdom	Modelling	25,000 simulations	N/R	General population (UK)	- Household - Work - School - Other	Contact tracing app (Bluetooth) with quarantine	-	- R - Outbreak control	- Combining contact tracing app with quarantine and reduce transmission more than mass testing or self-isolation alone - Reduction in transmission rate (reflective of R) was 47% when contact tracing app was used at 53% adoption rate - Maintaining an R < 1.0 requires a combination of self-isolation, contact tracing, and physical distancing

_											
8 4 5 6											- Outbreak control in a scenario where incidence is high requires a considerable number of individuals to be quarantined after contact tracing
7 8 9 10 11 12 13	Kurita 2020 (peer reviewed)	Japan	Modelling	N/R	5 months	General population (Japan)	-	COCOA contact tracing app (Bluetooth) with quarantine	-	R	- Reduction of R < 1.3 using a contact tracing app is not feasible if there are no voluntary restrictions - Reduction of R < 1.0 is feasible if contact tracing app adoption is 10% combined with 15% compliance for voluntary restrictions against going out
115 116 117 118 119 220 221 222 223 224 225 226 227 228	Nuzzo 2020 (peer reviewed)	USA	Modelling	N/R	400 days 150 days	Susceptible individuals	10	Contact tracing app (GPS,WiFi, and/or Bluetooth) with quarantine	Shelter in place	- Cumulative incidence SARS-CoV-2 - Mortality	- Contact tracing apps can mitigate infection spread similar to universal shelter-in-place, but with considerably fewer individuals isolated - Cumulative peak incidence can be reduced by 49% at 20% app adoption rate - Cumulative peak incidence can be reduced by 90% at 50% app adoption rate (similar to 40% compliance to shelter in place) - Mortality can be reduced by 23% at 20% app adoption rate
29 30 31 32 33 34 35 36 37 38 39 40	Pollmann 2020 (preprint)	Germany	Modelling	100 simulations	500 days	General population	-	Contact tracing app (Bluetooth) with quarantine		- R - Outbreak control - Cumulative incidence SARS-CoV-2	- Recursive tracing by contact tracing apps is more efficient than 1-step-tracing - Contact tracing apps alone cannot bring R below 1.0, unless 100% adoption is approached, and app notifications are strictly followed by quarantining and testing - Reducing an R _o of >3.0, in which 40% are asymptomatic SARS-CoV-2 carriers, below 1.0, can only be achieved by a contact tracing app if combined with other
17											

2											
3 4											interventions such as social distancing and/or random testing
5 6 7 8											- Reducing R significantly requires a contact tracing app adoption rate of at least 60%
9 10 11											- Cumulative incidence is reduced at any percentage of contact tracing app adoption
12 13 14 15	Scott 2020 (peer reviewed)	Australia	Modelling	N/R	3.5 months	Susceptible population (Victoria, Australia)	Various*	COVIDSafe contact tracing app (Bluetooth) with quarantine	-	Cumulative incidence SARS-CoV-2	- Impact of policy changes on cumulative incidence can take >2 months to become apparent
16 17 18 19) <u></u>	The questions			- Opening pubs/bars was identified as the greatest risk for increasing incidence of SARS-CoV-2. This could be mitigated by either of these measures:
20 21 22 23 24 25							(6)	with quarantine			 30% app adoption rate is achieved Transmission within venues was reduced by >40% through physical distancing policies Manual contact tracing was used that enabled >60% of contacts to be
26 27 28 29 30									0/1/	<u>,</u>	traced - Cumulative incidence is unlikely to be significantly impacted when app adoption rates are low-moderate
31 32 33 34 35 36 37 38 39 40	Shamil 2020 (preprint)	Banglades h	Modelling	N/R	60 days (Ford County) 120 days (New York city)	Susceptible population	- Healthcare workers - Students - Service holders - Unemployed people	Contact tracing app (not specified) with quarantine	- Lockdown - Extra personal protection	- R - Cumulative incidence SARS-CoV-2	- Reduction of R below 1.0 can be achieved within 3 weeks at 60% app adoption rate - Cumulative incidence approach zero within 3 months when 75% app adoption rate is achieved - Cumulative incidence is reduced by 3.5% when using a contact tracing app compared to not using one
41											compared to flot using one

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2					
3 [- Cumulative incidence is reduced by 4.6%
4					after 90 days when either:
5					
5					 All doctors, nurses, healthcare
7					workers and 50% of service holders
3					are using a contact tracing app for 2
9					days
10					o 75% of the population are using a
11					contact tracing app for 2 days

beer telien only

^{12 *} Household, school, work, community, church, professional sports, community sports, beaches, entertainment, cafés / restaurants, pubs / bars, public transport, national parks, public parks, large events, child care, social networks, and aged care

Table 2. Properties of model-based studies

Model-specific characteristics of model-based studies looking at effectiveness of contact- and tracing apps for SARS-CoV-2. Hyphens (-) indicate a continuous range between numbers, semicolons indicate separate distinct values. R = Reproductionnumber, N/A = not applicable, N/R = not reported, ODE = ordinary differential equations, PDE = partial differential equations, HH = household

	Model-related	Model-related properties		Contact- and tracing app related properties			Disease-related properties				Modifyable properties		
Study	Modeltype	Input parameter properties	Tracing direction	# of sequential generations	Adoption rate app	R	Incubation time	Infectious period	Probability of disease transmission	Delay symptom onset and testing	Delay testing and feedback app	Quarantine effectiveness	
Bradshaw 2020 (peer reviewed)	Branching- process model	Distributions	Bidirectional	Infinite generations	53;80%	2.5	5.5 days	Fitted to curve, value not specified	Fitted to curve, value not specified	1 days	0 days	90%	
Bulchandani 2020 (preprint)	Branching- process model	Based on exponential distributions	Bidirectional	3-infinite generations	0-100%	3.0	N/A ¹	N/A	N/R	N/A ²	0 days	100%	
Cencetti 2020 (preprint)	Continuous weighted temporal network	Distributions	Forward	1 generation	60;80;100%	1.2;1.5;2.0	Fitted to curve, value not specified	Fitted to curve, value not specified	Fitted to curve, value not specified	2 days	0 days	0-100%	
Currie 2020 (peer reviewed)	ODE compartmental model	Based on exponential distributions	Forward	1 generation	0;27;40;61;80%	2.5	2.0 days	11 days	N/R	3 days	N/R	90%	
Ferrari 2020 (peer reviewed)	ODE compartmental model	Based on exponential distributions	Forward	1 generation	0;25;50;75%	1.5	5.1 days	10 days	10%	2 days	N/R	90%	
Ferretti 2020 (peer reviewed)	PDE compartmental model	Distributions	Forward	1 generation	0-100%	2.0	5.5 days	12 days	Fitted to curve, value not specified	1.6 days	0 days	0-100%	
Grimm 2020 (preprint)	ODE compartmental model	Based on exponential distributions	Forward	1 generation	20-80%	2.2;3.0	5.0 days	10;12.5;14;20 days	N/R	N/R	N/R	100%³	

		Model-related properties		Contact- and tracing app related properties			Disease-related properties				Modifyable properties		
	Study	Modeltype	Input parameter properties	Tracing direction	# of sequential generations	Adoption rate app	R	Incubation time	Infectious period	Probability of disease transmission	Delay symptom onset and testing	Delay testing and feedback app	Quarantine effectiveness
0 1 2	Guttal 2020 (preprint)	Individual- based network model	Based on exponential distributions	Bidirectional	>1 generation	100%	3.0;4.0	N/A	20 days	0.2%	N/R	N/R	100%
3 4 5	Kretzschmar 2020 (peer reviewed)	Branching- process model	Distributions	Forward	1 generation	20;40;60;80;100%	2.5	6.4 days	10 days	2-12%	0 days	0 days	0;20;40; 60;80;100%
6 7 8	Kucharski 2020 (peer reviewed)	Individual- based network model	Distributions	Forward	1 generation	53%	2.6	5.0 days	5 days	20% within HH 6% outside HH 50% less for asymptomatic	0 days	0 days	90%
0 1	Kurita, 2020 (peer reviewed)	ODE compartmental model	Based on exponential distributions	N/R	1 generation	0;10;20;30;40;50; 60;70;80;90;100%	1.5	6.6 days	N/R	N/R	2 days	0 days	N/R
2 3 4 5	Nuzzo, 2020 (peer reviewed)	ODE compartmental model	Based on exponential distributions	N/A ⁴	N/A ⁴	0;10;20;30;40;50; 60;70;80;90%	3.02	5.1 days	N/R	Fitted to curve, value not specified	N/R	N/R	100%
6 7 8 9	Pollmann 2020 (preprint)	ODE compartmental model	Based on exponential distributions & distributions	Bidirectional	>1 generation	60;75;90;100%	2.0-3.0- 4.0	4.0;7.4 days	10 days	7% ⁵	0;2;4;6 days	N/R	100%
1 2 3	Scott 2020 (peer reviewed)	Agent-based model	Distributions	Forward	1 generation	0-50%	Fitted to curve, value not specified	4.6 days	8-14 days	Fitted to curve, value not specified	1 day	1 day	0% in HH 80-100% in other settings
4 5 6 7	Shamil 2020 (preprint)	Agent-based model	Distributions	Forward	1 generation	60;75%	Fitted to curve, value not specified	6.0 days	10 days	N/R	0 days	0 days	100%

¹ Fraction of infections before symptoms is relevant

² Isolation based on positive notification, not a positive test

³ Changing app coverage coveres imperfect isolation

⁴ No true tracing, fixed proportion cases will self-isolate

⁵ Time-dependent, maximum value reported in table

Table 3. Critical appraisal of empirical studies

Table - Critical appraisal empirical epidemiological studies looking at effectiveness of contact- and tracing apps for SARS-CoV-2

Study	Confounding?	Selection bias: participants?	Selection bias: missing data?	Information bias: intervention misclassification / non- compliance?	Information bias: Misclassification of the outcome?	Other concerns?	Overall risk of bias
Chen 2020 (peer reviewed)	Yes*	No	Unclear	No	Unclear	None	High
Kendall 2020 (peer reviewed)	Yes	No	Unclear	No	No	Competing interests and funding not reported	High
Only adjusted fo	or age			erien			

^{*} Only adjusted for age

Table 4. Critical appraisal of model-based studies

Table - Critical appraisal model based studies looking at effectiveness of contact- and tracing apps for SARS-CoV-2

Study	Were empirical distributions used for a varying infectiousness since time of infection?	Were various different scenarios evaluated for important model assumptions and parameter values?	Were models reported transparently? (i.e. no black box)	Other concerns?	Overall study validity
Bradshaw 2020 (peer reviewed)	Yes	Yes	Yes	External funding ¹	High
Bulchandani 2020 (preprint)	No	Yes	Yes	Competing interests & funding not reported	High
Cencetti 2020 (preprint)	Yes	Yes	Yes	No	High
Currie 2020 (peer reviewed)	Yes	Yes	Yes	No	High
Ferrari 2020 (peer reviewed)	No	Yes	Yes	Competing interests ²	High
Ferretti 2020 (peer reviewed)	Yes	Yes	Yes	No	High
Grimm 2020 (preprint)	No	Yes	Yes	No	High
Guttal 2020 (preprint)	Yes	Yes	Yes	Competing interests and funding not reported	High

Kretzschmar 2020 (peer reviewed)	Yes	Yes	Yes	No	High
Kucharski 2020 (peer reviewed)	Yes	Yes	Yes	Funding ³ , though no influence of funder on study results	High
Kurita, 2020 (peer reviewed)	No	No*	Unclear	Type of model used unclear	Low
Nuzzo, 2020 (peer reviewed)	No	No*	Yes	Potential competing interests ⁴	Low
Pollmann 2020 (preprint)	Yes	Yes	Yes	Competing interests and funding not reported	High
Scott 2020 (peer reviewed)	Yes	Yes	Yes	Funding ⁵	High
Shamil 2020 (preprint)	No	Yes	Unclear	No	Low

^{*} Scenarios were limited only to variation in rate of adoption of the contact- and tracing app and voluntary quarantine

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² E.S. works for Bayer, is collaborating to COVID Safe Paths app, by MIT, and advising LEMONADE tracing app, by Nuland. A.S.C. works for Roche Pharma. M.T.F is consultant for Ely Lilly.

³ Wellcome Trust, UK Engineering and Physical Sciences Research Council, European Commission, Royal Society, Medical Research Council.

⁴ Dr Raskar is the founder of a non-profit to facilitate digital contact tracing. The other authors report no potential competing interests.

⁵ Funding by the Burnet Institute



Supplementary file 1. Search strategy

Search strategy

On October 28th 2020 the comprehensive set of studies included in the COAP database (available on https://ispmbern.github.io/covid-19/living-review/collectingdata.html) was loaded in Endnote X9.

The dataset consisted of 82,401 references related to research on COVID-19. The following search was performed within this dataset:

(contact OR tracing OR track OR tracking OR warn OR warning) AND (smartphone OR app OR smartwatch OR device OR mobile OR smart phone OR bluetooth OR wearable OR iphone OR cell phone)

Background COAP database

The COAP database is a repository provided by Bern University, in which studies related to COVID-19 are incorporated. (available on https://ispmbern.github.io/covid-19/living-review/collectingdata.html)

Studies included in this repository are extracted on a daily basis from EMBASE (OVID), MEDLINE (PubMed), BioRxiv, and MedRxiv. References that are not yet available in the repository are added based on the date of publication provided by the aforementioned databases. The date on which the reference is added to the COAP database is included under the heading 'strategydate'.

Search strategies used for the COAP database are updated on a regular basis. An overview of these updates can be found below.

Initial search: 01.01.2020

MEDLINE

("Wuhan coronavirus" [Supplementary Concept] OR "COVID-19" OR "2019 ncov"[tiab] OR (("novel coronavirus"[tiab] OR "new coronavirus"[tiab]) AND (wuhan[tiab] OR 2019[tiab])) OR 2019-nCoV[All Fields] OR (wuhan[tiab] AND coronavirus[tiab])))))

EMBASE

ncov OR (wuhan AND corona) OR COVID

BioRxiv/MedRxiv

ncov or corona or wuhan or COVID

Update #1: 26.03.2020

MEDLINE

("Wuhan coronavirus" [Supplementary Concept] OR "COVID-19" OR SARS-CoV-2 OR "2019 ncov"[tiab] OR (("novel coronavirus"[tiab] OR "new coronavirus"[tiab]) AND (wuhan[tiab] OR 2019[tiab])) OR 2019-nCoV[All Fields] OR (wuhan[tiab] AND coronavirus[tiab]))

EMBASE

(nCoV or 2019-nCoV or ((new or novel or wuhan) adj3 coronavirus) or covid19 or covid-19 or SARS-CoV-2).mp.

BioRxiv/MedRxiv

ncov or corona or wuhan or COVID or SARS-CoV-2 With the kind support of the <u>Public Health & Primary Care Library PHC</u>, and following guidance of the <u>Medical Library Association</u>

Update #2: 01.04.2020

From 01.04.2020, we retrieve the currate BioRxiv/MedRxiv dataset Link

Update #3: 29.04.2020

MEDLINE

("coronavirus"[MH] OR "coronavirus infections"[MH] OR "coronavirus"[TW] OR "corona virus"[TW] OR "HCoV"[TW] OR "nCov"[TW] OR "covid"[TW] OR "covid19"[TW] OR "Severe Acute Respiratory Syndrome Coronavirus 2"[TW] OR "SARS-CoV2"[TW] OR "SARS-CoV2"[TW] OR "SARS Coronavirus 2"[TW] OR "MERS-CoV"[TW]) AND (2019/1/1:3000[PDAT])

Update #4: 01.05.2020

EMBASE

(SARS coronavirus/ or middle east respiratory syndrome/ or severe acute respiratory syndrome/ or (coronavirus* or corona virus* or HCoV* or ncov* or covid or covid19 or sars-cov* or sars-covo* or Sars-coronavirus* or Severe Acute Respiratory Syndrome Coronavirus*).mp.) and 20191201:20301231.(dc).

Update #5: 30.10.2020

EMBASE

(exp SARS-related coronavirus/ or severe acute respiratory syndrome/ or coronavirus disease 2019/ or (coronavir* or corona virus* or HCoV* or ncov* or 2019 cov or covid or covid19 or sars-cov* or sars-covo* or sars-coronavirus* or Severe Acute Respiratory Syndrome Coronavirus* or nCoV).mp.) and 20191101:20301231.(dc).

MEDLINE

("severe acute respiratory syndrome coronavirus 2"[Supplementary Concept] OR "COVID-19" [Supplementary Concept] OR "coronavirus" OR "corona virus" OR "HCoV" OR "nCoV" OR "2019 CoV" OR "covid" OR "covid19" OR "Severe Acute Respiratory Syndrome Coronavirus 2" OR "SARS-CoV2" OR "SARS-CoV 2" OR "SARS Coronavirus 2") AND (2019/11/01:3000/12/31[PDAT])

Supplementary file 2. Method for critical appraisal of empirical studies

Method used for critical appraisal of empirical epidemiologic studies

Confounding

Have the authors identified all important confounding factors? Yes / No / Unclear

Were the identified confounding factors <u>adjusted</u> for in the design and/or analysis? Yes / No

/ Unclear

- Model-based adjustment of confounders
- Stratification
- Matching
- No adjustment required (randomization)

Selection bias

Was patient exposure / intervention status at inclusion likely to result in bias? Yes / No /

Unclear

- Non-randomized study
- Randomized study with issues regarding allocation concealment or non-random sequencing
- Stringent exclusion criteria

Was missing data or loss to follow-up during the study likely to result in bias? Yes / No /

Unclear

- Missingness likely not completely at random (i.e. not MCAR or % of missingness different between groups)
- No methods described for handling missingness (i.e. imputation)
- Other methods explored to prevent missingness (i.e. cross checking data sources)

Informationbias

Was measurement of exposure / administration of the intervention likely to result in bias? Yes

/ No / Unclear

- Blinding
- Standardization
- Objective
- Non-compliance
- Breaking protocol

Was measurement of <u>outcome</u> likely to result in bias? Yes / No / Unclear

- Blinding
- Standardization
- Objective (note: if this is the case item should be scored 'No')

Other concerns? FREE TEXT

Items to consider (but not limited to)

- Reporting bias
- Conflict of interest

Supplementary file 3. Method for critical appraisal of model-based studies

Method used for critical appraisal of model based studies

Were empirical distributions used for a varying infectiousness since time of infection?

Yes / No / Unclear

Keywords indicating distributions were used

- Weibull
- Log-normal
- Exponential distribution

Were various different scenarios evaluated for important model assumptions and parameter values? Yes / No / Unclear

Keywords indicating uncertainty was taken into account

- Sensitivity analysis
- Scenario analysis

Were models reported transparently? (i.e. no black box) Yes / No / Unclear

Key elements indicating that model can be reproduced

- (differential) Equation specified
- Behavior of agents specified
- Graphic representation of model
- All variables and distributions specified

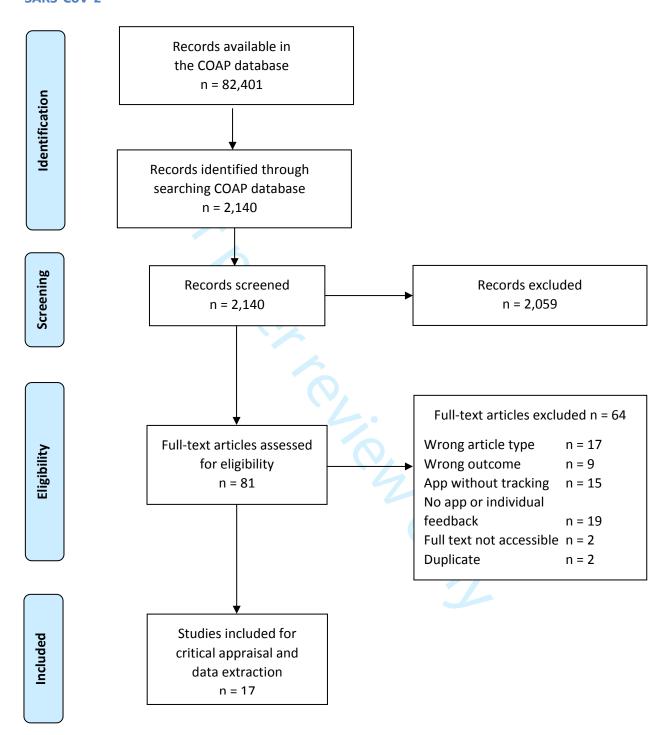
Other concerns? FREE TEXT

Items to consider (but not limited to)

- Reporting bias
- Conflict of interest
- Illogical properties of the model not captured by the criteria above

Supplementary file 4. Flowchart study selection

Flowchart regarding selection of studies looking at effectiveness of contact- and tracing apps for SARS-CoV-2



Supplementary file 5. Excluded studies

Studies not meeting inclusion criteria after full text screening, and excluded from analyses (n=64)

Reference Reason for exclusion Aleta 2020 No app or individual feedback Aleta 2020 No app or individual feedback Ayres 2020 Wrong outcome Bian 2020 Wrong article type Bianconi 2020 Full text not accessible Braithwaite 2020 Braithwaite 2020 Duplicate	
Aleta 2020 No app or individual feedback Ayres 2020 Wrong outcome Bian 2020 Wrong article type Bianconi 2020 Full text not accessible Braithwaite 2020 Wrong article type	
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Bianconi 2020 Full text not accessible Braithwaite 2020 Wrong article type	
Braithwaite 2020 Wrong article type	
Duplicate	
Braun 2020 Full text not accessible	
Brooks-Pollock 2020 No app or individual feedback	
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Drake 2020 Wrong article type	
Drew 2020 App without tracking	
Fateh-Moghadam 2020 App without tracking	
Fenton 2020 Wrong outcome	
Firth 2020 No app or individual feedback	
Gozzi 2020 App without tracking	
Grantz 2020 Wrong outcome	
Güemes 2020 App without tracking	
Haller 2020 Wrong article type	
Huang 2020 Wrong outcome	
Hussein 2020 No app or individual feedback	
Jian 2020 Wrong outcome	
Kassaye 2020 App without tracking	
Kendall 2020 Duplicate	
Khataee 2020 Wrong article type	
Kogan 2020 Wrong outcome	
Kretzschmar 2020 Duplicate	
Lambert 2020 Wrong article type	
Leith 2020 Wrong article type	
Liu 2020 No app or individual feedback	
Maghdid 2020 Wrong article type	
Marín-García 2020 Wrong article type	
Menni 2020 App without tracking	
Menni 2020 App without tracking	
Milenkovic 2020 No app or individual feedback	
Mishra 2020 App without tracking	
Morley 2020 No app or individual feedback	

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PRISMA 2009 Checklist

Section/topic	#	Checklist item	Reported on page #		
TITLE	ITLE				
Title	1	Identify the report as a systematic review, meta-analysis, or both.	1		
ABSTRACT					
Structured summary	2	Provide a structured summary including, as applicable: background; objectives; data sources; study eligibility criteria, participants, and interventions; study appraisal and synthesis methods; results; limitations; conclusions and implications of key findings; systematic review registration number.	3		
INTRODUCTION	•				
Rationale	3	Describe the rationale for the review in the context of what is already known.	5		
Objectives	4	Provide an explicit statement of questions being addressed with reference to participants, interventions, comparisons, outcomes, and study design (PICOS).	5-6		
METHODS					
Protocol and registration	5	Indicate if a review protocol exists, if and where it can be accessed (e.g., Web address), and, if available, provide registration information including registration number. Given the rapid nature of this systematic review, no protocol was registered beforehand	-		
Eligibility criteria	6	Specify study characteristics (e.g., PICOS, length of follow-up) and report characteristics (e.g., years considered, language, publication status) used as criteria for eligibility, giving rationale.	6		
Information sources	7	Describe all information sources (e.g., databases with dates of coverage, contact with study authors to identify additional studies) in the search and date last searched.	6, 29		
Search	8	Present full electronic search strategy for at least one database, including any limits used, such that it could be	7		
		repeated.	Suppl. File page 1-3		
Study selection	9	State the process for selecting studies (i.e., screening, eligibility, included in systematic review, and, if applicable, included in the meta-analysis).	7		
Data collection process	10	Describe method of data extraction from reports (e.g., piloted forms, independently, in duplicate) and any processes for obtaining and confirming data from investigators.	7-8		
Data items	11	List and define all variables for which data were sought (e.g., PICOS, funding sources) and any assumptions and simplifications made.	7-9		
Risk of bias in individual studies	12	Describe methods used for assessing risk of bias of individual studies (including specification of whether this was done at the study or outcome level), and how this information is to be used in any data synthesis.	7 Suppl. File		
		For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml	page 7, 8		



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PRISMA 2009 Checklist

Summary measures	13	State the principal summary measures (e.g., risk ratio, difference in means).	7-8
Synthesis of results		Describe the methods of handling data and combining results of studies, if done, including measures of consistency (e.g., l^2) for each meta-analysis.	8

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Page 1 of 2					
Section/topic	#	Checklist item	Reported on page #		
Risk of bias across studies	15	Specify any assessment of risk of bias that may affect the cumulative evidence (e.g., publication bias, selective reporting within studies).	-		
Additional analyses	16	Describe methods of additional analyses (e.g., sensitivity or subgroup analyses, meta-regression), if done, indicating which were pre-specified.	-		
RESULTS	•	·			
Study selection	17	Give numbers of studies screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally with a flow diagram.	8 Suppl. File page 4-6		
Study characteristics	18	For each study, present characteristics for which data were extracted (e.g., study size, PICOS, follow-up period) and provide the citations.	8, 9, 17-25		
Risk of bias within studies	19	Present data on risk of bias of each study and, if available, any outcome level assessment (see item 12).	9, 26-28		
Results of individual studies	20	For all outcomes considered (benefits or harms), present, for each study: (a) simple summary data for each intervention group (b) effect estimates and confidence intervals, ideally with a forest plot.	10-13		
Synthesis of results	21	Present results of each meta-analysis done, including confidence intervals and measures of consistency. Not applicable, as only qualitative assessment was possible	-		
Risk of bias across studies	22	Present results of any assessment of risk of bias across studies (see Item 15).	-		
Additional analysis	23	Give results of additional analyses, if done (e.g., sensitivity or subgroup analyses, meta-regression [see Item 16]).	-		
DISCUSSION					
Summary of evidence	24	Summarize the main findings including the strength of evidence for each main outcome; consider their relevance to key groups (e.g., healthcare providers, users, and policy makers).	13		
Limitations	25	Discuss limitations at study and outcome level (e.g., risk of bias), and at review-level (e.g., incomplete retrieval of identified research, reporting bias).	13, 14		
Conclusions	26	Provide a general interpretation of the results in the context of other evidence, and implications for future research.	14		
FUNDING	1				



PRISMA 2009 Checklist

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4 5	Funding	27	Describe sources of funding for the systematic review and other support (e.g., supply of data); role of funders for the systematic review.	2
6 =		<u> </u>		<u>'</u>
8	From: Moher D, Liberati A, Tetzlaff doi:10.1371/journal.pmed1000097	J, Altm	an DG, The PRISMA Group (2009). Preferred Reporting Items for Systematic Reviews and Meta-Analyses: The PRISMA Statement. PLoS Med	6(7): e100009
9			For more information, visit: www.prisma-statement.org .	
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