

Original Investigation | Global Health Household Transmission of SARS-CoV-2 A Systematic Review and Meta-analysis

Zachary J. Madewell, PhD; Yang Yang, PhD; Ira M. Longini Jr, PhD; M. Elizabeth Halloran, MD, DSc; Natalie E. Dean, PhD

Abstract

IMPORTANCE Crowded indoor environments, such as households, are high-risk settings for the transmission of severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2).

OBJECTIVES To examine evidence for household transmission of SARS-CoV-2, disaggregated by several covariates, and to compare it with other coronaviruses.

DATA SOURCE PubMed, searched through October 19, 2020. Search terms included *SARS-CoV-2* or *COVID-19* with secondary attack rate, household, close contacts, contact transmission, contact attack rate, or family transmission.

STUDY SELECTION All articles with original data for estimating household secondary attack rate were included. Case reports focusing on individual households and studies of close contacts that did not report secondary attack rates for household members were excluded.

DATA EXTRACTION AND SYNTHESIS Meta-analyses were done using a restricted maximumlikelihood estimator model to yield a point estimate and 95% CI for secondary attack rate for each subgroup analyzed, with a random effect for each study. To make comparisons across exposure types, study was treated as a random effect, and exposure type was a fixed moderator. The Preferred Reporting Items for Systematic Reviews and Meta-analyses (PRISMA) reporting guideline was followed.

MAIN OUTCOMES AND MEASURES Secondary attack rate for SARS-CoV-2, disaggregated by covariates (ie, household or family contact, index case symptom status, adult or child contacts, contact sex, relationship to index case, adult or child index cases, index case sex, number of contacts in household) and for other coronaviruses.

RESULTS A total of 54 relevant studies with 77 758 participants reporting household secondary transmission were identified. Estimated household secondary attack rate was 16.6% (95% CI, 14.0%-19.3%), higher than secondary attack rates for SARS-CoV (7.5%; 95% CI, 4.8%-10.7%) and MERS-CoV (4.7%; 95% CI, 0.9%-10.7%). Household secondary attack rates were increased from symptomatic index cases (18.0%; 95% CI, 14.2%-22.1%) than from asymptomatic index cases (0.7%; 95% CI, 0%-4.9%), to adult contacts (28.3%; 95% CI, 20.2%-37.1%) than to child contacts (16.8%; 95% CI, 12.3%-21.7%), to spouses (37.8%; 95% CI, 25.8%-50.5%) than to other family contacts (17.8%; 95% CI, 11.7%-24.8%), and in households with 1 contact (41.5%; 95% CI, 31.7%-51.7%) than in households with 3 or more contacts (22.8%; 95% CI, 13.6%-33.5%).

Key Points

Question What is the household secondary attack rate for severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2)?

Findings In this meta-analysis of 54 studies with 77 758 participants, the estimated overall household secondary attack rate was 16.6%, higher than observed secondary attack rates for SARS-CoV and Middle East respiratory syndrome coronavirus. Controlling for differences across studies, secondary attack rates were higher in households from symptomatic index cases than asymptomatic index cases, to adult contacts than to child contacts, to spouses than to other family contacts, and in households with 1 contact than households with 3 or more contacts.

Meaning These findings suggest that households are and will continue to be important venues for transmission, even in areas where community transmission is reduced.

Supplemental content

Author affiliations and article information are listed at the end of this article.

(continued)

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Abstract (continued)

CONCLUSIONS AND RELEVANCE The findings of this study suggest that given that individuals with suspected or confirmed infections are being referred to isolate at home, households will continue to be a significant venue for transmission of SARS-CoV-2.

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Introduction

The coronavirus disease 2019 (COVID-19) pandemic is caused by severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2), which is spread via direct or indirect contact with infected people via infected respiratory droplets or saliva, fomites, or aerosols.^{1,2} Crowded indoor environments with sustained close contact and conversations, such as households, are a particularly high-risk setting.³

The World Health Organization China Joint Mission reported human-to-human transmission in China largely occurred within families, accounting for 78% to 85% of clusters in Guangdong and Sichuan provinces.⁴ Stay-at-home orders reduced human mobility by 35% to 63% in the United States,⁵ 63% in the United Kingdom,⁶ and 54% in Wuhan,⁷ relative to normal conditions, which concomitantly increased time at home. Modeling studies demonstrated that household transmission had a greater relative contribution to the basic reproductive number after social distancing (30%-55%) than before social distancing (5%-35%).⁸ While current US Centers for Disease Control and Prevention recommendations are to maintain 6 feet of distance from a sick household member, this may be difficult to achieve in practice and not be fully effective.⁹

The household secondary attack rate characterizes virus transmissibility. Studies can collect detailed data on type, timing, and duration of contacts and identify risk factors associated with infectiousness of index cases and susceptibility of contacts. Our objective was to estimate the secondary attack rate of SARS-CoV-2 in households and determine factors that modify this parameter. We also estimated the proportion of households with index cases that had any secondary transmission. Furthermore, we compared the SARS-CoV-2 household secondary attack rate with that of other severe viruses and with that to close contacts for studies that reported the secondary attack rate for both close and household contacts.

Methods

Definitions

We estimated the transmissibility of SARS-CoV-2 within the household or family by the empirical secondary attack rate by dividing the number of new infections among contacts by the total number of contacts. Household contacts include anyone living in the same residence as the index case. Family contacts include the family members of index cases, including individuals who live outside the index case's household. Close contact definitions varied by study and included physical proximity to an index case, exceeding a minimum contact time, and/or not wearing effective protection around index cases before the index case was tested.

Search Strategy

Following Preferred Reporting Items for Systematic Reviews and Meta-analyses (PRISMA) reporting guideline, we searched PubMed using terms including *SARS-CoV-2* or *COVID-19* with *secondary attack rate*, *household*, *close contacts*, *contact transmission*, *contact attack rate*, or *family transmission* (eTable 1 in the Supplement) with no restrictions on language, study design, time, or place of publication. The last search was conducted October 19, 2020.

Eligibility Criteria

Eligibility criteria are described in eAppendix 1 in the Supplement. All articles with original data for estimating household secondary attack rate were included. Case reports focusing on individual households and studies of close contacts that did not report secondary attack rates for household members were excluded.

Data Extraction

One of us (Z.J.M.) extracted data from each study. Details appear in eAppendix 2 in the Supplement.

Evaluation of Study Quality and Risk of Bias

To assess the methodological quality and risk of bias of included studies of SARS-CoV-2, we used the same modified version of the Newcastle-Ottawa quality assessment scale for observational studies used by Fung et al.^{10,11} Studies received as many as 9 points based on participant selection (4 points), study comparability (1 point), and outcome of interest (4 points). Studies were classified as having high (\leq 3 points), moderate (4-6 points), and low (\geq 7 points) risk of bias. One of us (Z.J.M.) evaluated the study quality and assigned the quality grades.

Statistical Analysis

Meta-analyses were done using a restricted maximum-likelihood estimator model to yield Freeman-Tukey double arcsine-transformed point estimates and 95% CI for secondary attack rate for each subgroup analyzed, with a random effect for each study.¹² For comparisons across covariates (ie, household or family, index case symptom status, adult or child contacts, contact sex, relationship to index case, adult or child index cases, index case sex, number of household contacts, study location, universal or symptomatic testing, dates of study) and comparisons with close contacts and other viruses, study was treated as a random effect, and the covariate was a fixed moderator. Variables had to have been collected in at least 3 studies to be included in meta-analyses. The Cochran Q test and l^2 statistic are reported as measures of heterogeneity. l^2 values of 25%, 50%, and 75% indicated low, moderate, and high heterogeneity, respectively.¹³ Stastistical significance was set at a 2-tailed $\alpha = .05$. All analyses were done in R version 4.0.2 using the package metafor (R Project for Statistical Computing).^{14,15}

When at least 10 studies were available, we used funnel plots, Begg correlation, and Egger test to evaluate publication bias, with significance set at P < .10.^{16,17} If we detected publication bias, we used the Duval and Tweedie trim-and-fill approach for adjustment.¹⁸

Results

We identified 54 relevant published studies that reported household secondary transmission, with 77 758 participants (eTable 1 in the Supplement).¹⁹⁻⁷² A total of 16 of 54 studies (29.6%) were at high risk of bias, 27 (50.0%) were moderate, and 11 (20.4%) were low (eTable 2 in the Supplement). Lower quality was attributed to studies with 1 or fewer test per contact (35 studies [64.8%]), small sample sizes (31 [57.4%]), and secondary attack rate not disaggregated by covariates (28 [51.9%]).

A description of index case identification period and methods and symptom status is provided in eTable 3 in the Supplement. Most studies did not describe how co-primary index cases were handled or whether secondary infections could have been acquired from outside the household, both of which can inflate the empirical secondary attack rate. Testing and monitoring strategies varied between studies, often reflecting variations in local testing guidelines implemented as part of contact tracing (eTable 4 and eAppendix 3 in the Supplement).

Figure 1 summarizes secondary attack rates for 44

studies^{19-26,28-30,32-36,38-45,47-57,59,61-63,65-67,69,70} of household contacts and 10 of family contacts.^{26,31,37,45,58,60,65,68,71,72} Estimated mean secondary attack rate for household contacts was 16.4% (95% CI, 13.4%-19.6%) and family contacts was 17.4% (95% CI, 12.7%-22.5%). One study⁴⁰

Figure 1. Secondary Attack Rates (SAR) of Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2) for Household Contacts and Family Contacts

Source	Location	Participants, No.	Participants with SARS-CoV infection, No.	-2 SAR (95% CI)		Weight, %
Household contacts						
Boscolo-Rizzo et al, ²⁴ 2020	Treviso Province, Italy	121	54	0.45 (0.36-0.54)	-	1.93
Patel et al, ⁵³ 2020	London, UK	185	79	0.43 (0.36-0.50)	_	2.17
Rosenberg et al, 55 2020	New York, US	343	131	0.38 (0.33-0.43)		2.43
Dattner et al, 29 2020	Bnei Brak, Israel	2824	981	0.35 (0.33-0.37)	-#-	2.//
Lopez Bernal et al,47 2020	UK	4/2	161	0.34 (0.30-0.38)		2.54
Wu et al, 03 2020	Zhunai, China	148	48	0.32 (0.25-0.40)		2.08
	Atlanta UC	155	4/	0.30 (0.23-0.38)		2.12
Louis et al. 44 2020	Attallita, US	100	51	0.29 (0.21-0.36)		1.92
Dawson at al. 30 2020	Wisconsin, US	100	16	0.26 (0.21-0.34)		2.25
Wang et al 63 2020	Rojijing Chipa	225	77	0.23 (0.13-0.30)		2.01
Han 35 2020	South Koroa	1/	3	0.23 (0.13-0.28)		2.47
Röhmer et al 23 2020	Bayaria Cormany	24	5	0.21 (0.03-0.47)		0.00
Bao at al 21 2020	Cheonan South Korea	24	37	0.18 (0.13-0.24)		0.37
Vin et al 67 2020	Oinadao Muncinal China	106	19	0.18 (0.11-0.24)		2.52
Wu et al 66 2020	Hangzhou, China	280	50	0.18 (0.11-0.20)		2.01
Hu et al 36 2020	Hunan China	200	491	0.18 (0.16-0.19)	+	2.45
ling et al ³⁹ 2020	Guangzhou China	542	93	0 17 (0 14-0 20)		2.70
l vngse et al ⁴⁹ 2020	Denmark	2226	371	0 17 (0 15-0 18)		2.02
Doung-ngern et al 32 2020	Thailand	230	38	0 17 (0 12-0 22)		2.77
Li et al ⁴⁵ 2020	Wuhan China	392	64	0.16 (0.13-0.20)		2.55
Zhang et al ⁷⁰ 2020	China	62	10	0.16 (0.08-0.26)		1 70
Wang et al 62 2020	Reijing China	714	111	0.16 (0.13-0.18)		2.67
Park et al ⁵² 2020	Seoul South Korea	225	34	0.15 (0.11-0.20)		2.07
Fateh-Moghadam et al ³⁴ 2020	Trento Italy	3546	500	0.14 (0.13-0.15)	+	2.35
Islam and Noman ³⁸ 2020	Chattogram Bangladesh	46	6	0 13 (0 05-0 25)		1 55
Park et al ⁵¹ 2020	South Korea	10 592	1248	0 12 (0 11-0 12)		2.81
Phirivasart et al ⁵⁴ 2020	Pattani Province Thailand	106	12.10	0 11 (0 06-0 18)		2.01
Bi et al ²² 2020	Shenzhen China	686	77	0 11 (0 09-0 14)		2.68
Arnedo-Pena et al ²⁰ 2020	Castellon Spain	745	83	0 11 (0 09-0 14)		2.69
Adamik et al ¹⁹ 2020	Poland	32 023	3553	0 11 (0 11-0 11)		2.83
Malheiro et al ⁵⁰ 2020	Fastern Porto Portugal	780	83	0 11 (0 09-0 13)		2.02
Chaw et al ²⁶ 2020	Brunei	264	28	0 11 (0 07-0 15)		2.7 0
Burke. ²⁵ 2020	US	19	2	0.11 (0.00-0.29)		1.00
Luo et al. ⁴⁸ 2020	Guangzhou, China	1015	105	0.10 (0.09-0.12)		2.73
Laxminarayan et al, ⁴² 2020	Tamil Nadu and Andhra Pradesh, India	4065	380	0.09 (0.08-0.10)	•	2.80
Shah et al, ⁵⁶ 2020	Gujarat, India	386	34	0.09 (0.06-0.12)	- - -	2.61
Son et al, ⁵⁷ 2020	Busan, South Korea	196	16	0.08 (0.05-0.12)	_ _	2.44
Korea CDC, ⁴¹ 2020	South Korea	119	9	0.08 (0.03-0.13)	_ 	2.26
Cheng et al, ²⁸ 2020	Taiwan	151	10	0.07 (0.03-0.11)		2.38
Yung et al, ⁶⁹ 2020	Singapore	200	13	0.06 (0.03-0.10)		2.48
Lee et al, ⁴³ 2020	Busan, South Korea	23	1	0.04 (0.00-0.18)		1.42
Draper et al, ³³ 2020	Northern Territory, Australia	51	2	0.04 (0.00-0.11)		1.98
Kim et al, ⁴⁰ 2020	South Korea	208	1	0.00 (0.00-0.02)	-	2.72
Subgroup estimate				0.164 (0.134-0.196)	\diamond	100
Family contacts	Zhajiang Province, China	508	180	032(020025)		11.05
van der Hoek et al 60 2020	Nothorlands	17/	103	0.32 (0.20-0.33)		11.UD Ω ΕΛ
Wang of al 64 2020	Wuhan China	12	+/ 10	0.27 (0.21-0.34)		0.04
Dong et al 31 2020	Tianiin China	259	53	0.20 (0.12-0.37)		4.42 0.70
Hup et al 37 2020	Zhajiana Provinca China	835	151	0.18 (0.16-0.20)		9.70 11 66
Chen et al 27 2020	Ningho China	272	49	0.18 (0.10-0.21)		10.00
Liu et al 46 2020	Guanadona Provinco China	2/2	330	0.10(0.14-0.23)		10.00
7hang et al 71 2020		2771 Q2	12	0.14(0.12-0.13) 0.13(0.07-0.21)		7 52
Virat al 68 2020	Wuhan China	1396	143	0.10(0.00-0.21)	-	7.JJ 17.17
7huang et al 72 2020	Guanadona Provinco China	3697	276	0.07(0.03-0.12)		12.17
Znually et al, 2020	Guanguong Frovince, Chilld	2021	270	0.07 (0.07=0.00)	-	12.31
Subgroup estimate				0.174 (0.127-0.225)		100
Combined estimate ^a				0.166 (0.140-0.193)	◆	
				(0 0.25 0.5 SAR (95% CI)	0.75

Point sizes are an inverse function of the precision of the estimates, and bars correspond to 95% Cls. CDC indicates Centers for Disease Control and Prevention.

^a Weights for the combined estimate are available in eTable 8 in the Supplement.

restricted index cases to children (age <18 years), resulting in a substantially lower secondary attack rate of 0.5%. Excluding this outlier, the combined secondary attack rate for household and family contacts was 17.1% (95%, 14.6%-19.7%). Secondary attack rates for household and family contacts were more than 3 times higher than for close contacts (4.8%; 95% CI, 3.4%-6.5%; P < .001) (eFigure 2 in the Supplement). Significant heterogeneity was found among studies of household $(l^2 = 96.9\%; P < .001)$, family $(l^2 = 93.0\%; P < .001)$, and close $(l^2 = 97.0\%; P < .001)$ contacts. No significant publication bias was observed for studies of household, family, or close contacts (eFigure 3 in the Supplement). Secondary attack rates were not significantly different when restricting to 38 studies^{19,20,22,23,26-31,34-40,42,44-51,54-57,60,62,63,65,67-69,72} with low or moderate risk of bias (15.6%; 95%, 12.8%-18.5%) (eFigure 4 in the Supplement). There were no significant differences in secondary attack rates between 21 studies in China^{22,27,31,36,37,39,45,46,48,58,61-68,70-72} and 33 studies from other countries^{19-21,23-26,28-30,32-35,38,40-44,47,49-57,59,60,69} (eFigure 5 in the Supplement), 18 studies that tested symptomatic contacts^{19-21,24,25,28,29,33,34,41,47,50,53,56,58,59,61,64} and 33 studies that reported testing all contacts^{22,23,26,27,30,31,35-40,42-46,48,49,51,52,54,55,57,60,63,65-67,69-72} (eFigure 6 in the Supplement), and 16 early studies^{22,23,25,31,37,39,45,58,61,63-66,68,71,72} (January-February) and 20 later studies^{19,24,26,29,30,32-35,38,42,44,50,53-56,59,60,69} (March-July) (eFigure 7 in the Supplement).

To study the transmissibility of asymptomatic SARS-CoV-2 index cases, eFigure 8 in the Supplement summarizes 27 studies^{19-21,23-26,30,32-34,44,45,47,50,52-54,56,59-61,63,64,68,69,72} reporting household secondary attack rates from symptomatic index cases and 4 studies^{26,43,44,52} from asymptomatic or presymptomatic index cases. Estimated mean household secondary attack rate from symptomatic index cases (18.0%; 95% CI, 14.2%-22.1%) was significantly higher than from asymptomatic or presymptomatic index cases (0.7%; 95% CI, 0%-4.9%; *P* < .001), although there were few studies in the latter group. These findings are consistent with other household studies^{28,70} reporting asymptomatic index cases as having limited role in household transmission.

There is evidence for clustering of SARS-CoV-2 infections within households, with some households having many secondary infections while many others have none.⁷³⁻⁷⁵ For example, 1 study⁵⁵ reported that 26 of 103 (25.2%) households had all members test positive. This is consistent with observation of overdispersion in the number of secondary cases per index case across a range of settings.³ While most studies reported only the average number of secondary infections per index case, some also reported transmission by household.^{44,55,56,63,65,69} **Figure 2** summarizes the proportion of households with any secondary transmission. Using an empirical analysis based on secondary attack rates and mean number of contacts per household, we found the proportion of

Figure 2. Mean Number of Contacts per Household, Secondary Attack Rate (SAR) of Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2), and Proportion of Households Reporting Any Secondary Transmission From Index Cases

		Mean	Conta	octs	House	holds	Proportion of households with any secondary transmission		Weight
Source	Location	contacts	Total	Infected	Total	Infected	(95% CI)		%
Wu et al, ⁶⁵ 2020	Zhuhai, China	4.229	148	48	35	22	0.63 (0.46-0.78)	_	12.78
Rosenberg et al, ⁵⁵ 2020	New York, US	3.330	343	131	103	63	0.61 (0.52-0.70)	A	14.62
Lewis et al, ⁴⁴ 2020	Utah and Wisconsin, US	3.241	188	52	58	32	0.55 (0.42-0.68)	<u>_</u>	13.78
Wang et al, ⁶³ 2020	Beijing, China	2.702	335	77	124	41	0.33 (0.25-0.42)	_ _	14.84
Shah et al, ⁵⁶ 2020	Gujarat, India	5.216	386	34	74	16	0.22 (0.13-0.32)	— — •	14.43
Yung et al, ⁶⁹ 2020	Singapore	1.493	200	13	134	7	0.05 (0.02-0.10)		15.36
Draper et al, ³³ 2020	Northern Territory, Australia	1.821	51	2	28	1	0.04 (0.00-0.15)		14.19
Model estimate							0.317 (0.134-0.534)		100
								0 0.25 0.5 0.75 Proportion of households with any secondary transmission (95% CI)	1

The expected proportion of households with any secondary transmission (represented by the triangles) was calculated as proportion with at least 1 secondary infection in a household = $1 - (1 - SAR)^n$, where *n* is the mean number of contacts for that study

(eTable 5 in the Supplement). Point sizes are an inverse function of the precision of the estimates, and bars correspond to 95% CIs.

households with any secondary transmission was lower than expected in a setting with no clustering (eg, most transmission is not characterized by a minority of infected individuals) (eTable 5 in the Supplement). Ideally, future studies will assess this formally by fitting a β binomial to quantify overdispersion in the full data.

A number of studies examined factors associated with susceptibility of household contacts to infection (eTable 6 in the Supplement). Age was the most examined covariate, with most studies^{20,29,36-39,45,46,48,49,55,63,65,68} reporting lower secondary transmission of SARS-CoV-2 to child contacts than adult contacts. In 5 studies,^{20,36,39,48,49} individuals older than 60 years were most susceptible to SARS-CoV-2 infection. Contact age was not associated with susceptibility in 9 studies,^{26,28,32,44,47,58,66,67,70} although these were typically less powered to detect a difference. **Figure 3** summarizes 15 studies^{22,26,29,37,39,42,44,45,47,49,55,59,60,63,65} reporting separate secondary attack rates to children and adult contacts. The estimated mean household secondary attack rate was significantly higher to adult contacts (28,3%; 95% CI, 20,2%-37,1%) than to child contacts (16,8%; 95% CI, 12,3%-21,7%; *P* < .001). Significant heterogeneity was found among studies of adult (*l*² = 96,8%; *P* < .001) and child contacts (*l*² = 78,9%; *P* < .001). Begg (*P* = .03) and Egger (*P* = .03) tests were statistically significant for studies of adult but not child contacts (eFigure 9 in the Supplement). One study of adults⁶³ had a high secondary attack rate in the forest plot. Excluding this study improved the funnel plot symmetry and resulted in a secondary attack rate to adult contacts of 26,3% (95% CI, 19,3%-33,2%).

The second most examined factor was sex of exposed contacts, which was not associated with susceptibility for most studies^{20,22,26,32,36,39,44,45,47,49,58,65-67,70} except 3.^{38,46,68} eFigure 10 in the Supplement summarizes results from 11 studies^{20,39,42,44,45,47,49,58,65,67,69} reporting household secondary attack rates by contact sex. Estimated mean household secondary attack rate to female contacts (20.7%; 95% CI, 15.0%-26.9%) was not significantly different than to male contacts (17.7%; 95% CI, 12.4%-23.8%). Significant heterogeneity was found among studies of female contacts ($l^2 = 87.4\%$; P < .001) and male contacts ($l^2 = 87.7\%$; P < .001). Moderate asymmetry was observed in the funnel plots, which was significant for studies of female contacts from Egger test (P = .07) but not male contacts (eFigure 11 in the Supplement). However, imputation of an adjusted effect size using the trim-and-fill method did not significantly change the secondary attack rate to female contacts (19.7%; 95% CI, 13.9%-25.6%).

Spouse relationship to index case was associated with secondary infection in 4 studies^{26,45,46,58} of 6 in which this was examined.^{65,67} Infection risk was highest for spouses, followed by nonspouse family members and other relatives, which were all higher than other contacts.⁴⁶ **Figure 4** summarizes results from 7 studies^{26,44,46,58,65,67} reporting household secondary attack rates by relationship. Estimated mean household secondary attack rate to spouses (37.8%; 95% CI, 25.8%-50.5%) was significantly higher than to other contacts (17.8%; 95% CI, 11.7%-24.8%). Significant heterogeneity was found among studies of spouses ($I^2 = 78.6\%$; P < .001) and other relationships ($I^2 = 83.5\%$; P < .001).

Several studies examined factors associated with infectiousness of index cases. Older index case age was associated with increased secondary infections in 3 studies^{20,47,67} of 9 in which this was examined.^{22,36,39,44,63,65} eFigure 12 in the Supplement summarizes results from 3 studies^{42,44,51} reporting household secondary attack rates by index case age. Estimated mean household secondary attack rate from adults (15.2%; 95% CI, 6.2%-27.4%) was not significantly different than that from children (7.9%; 95% CI, 1.7%-16.8%). Index case sex was associated with transmission in 3 studies^{42,44,67} of 9 in which this was examined.^{20,36,45,47,63,65} eFigure 13 in the Supplement summarizes results from 7 studies^{20,42,44,45,65,67,69} reporting household secondary attack rates by index case sex. Estimated mean household secondary attack rate from female contacts (16.6%; 95% CI, 11.2%-22.8%) was not significantly different than from male contacts (16.4%; 95% CI, 9.0%-25.5%).

Critically severe index case symptoms was associated with higher infectiousness in 6 studies^{20,38,46-48,67} of 9 in which this was examined.^{44,63,70} Index case cough was associated with

infectivity in 2 studies ^{20,65} of 8 in which this was examined ^{45-48,63,67} (eAppendix 4 in the Supplement).

Contact frequency with the index case was associated with higher odds of infection, specifically at least 5 contacts during 2 days before the index case was confirmed,⁷⁰ at least 4 contacts and 1 to 3 contacts,⁶³ or frequent contact within 1 meter.^{22,67,68} Smaller households were associated with transmission in 4 studies^{20,39,47,49} of 7 in which this was examined.^{55,63,65} **Figure 5** summarizes results from 6 studies^{20,47,49,55,61,65} reporting household secondary attack rates by number of

Figure 3. Secondary Attack Rates (SAR) of Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2) for Adult (≥18 Years) and Child (<18 Years) Household and Family Contacts

			Participants								
Source	Location	Participants, No.	SARS-CoV-2 infection, No.	SAR (95% CI)							Weight, %
Wang et al, ⁶³ 2020	Beijing, China				1						
Adults		92	64	0.70 (0.60-0.79)					-		- 6.07
Children		36	13	0.36 (0.21-0.53)				•	-		2.81
Rosenberg et al, ⁵⁵ 2020	New York, US	102		0.40(0.41.0.50)							6.57
Adults		182	88	0.48 (0.41-0.56)					_		6.57
		150	42	0.27 (0.20-0.34)							0.05
Dattner et al, 29 2020	Bhei Brak, Israel	1448	637	0 44 (0 41-0 47)				-			7 21
Children		1376	344	0.25 (0.23-0.27)			-	_			9.92
Lopez Bernal et al 47 2020		1570	544	0.25 (0.25 0.27)			_				5.52
Adults	UK	297	119	0.40 (0.35-0.46)							6.85
Children		175	42	0.24 (0.18-0.31)			_				7.01
Wu et al. ⁶⁵ 2020	Zhuhai. China										
Adults	,	112	43	0.38 (0.30-0.48)							6.20
Children		31	5	0.16 (0.05-0.31)							3.05
Teherani et al, ⁵⁹ 2020	Atlanta, US										
Adults		64	20	0.31 (0.20-0.43)							5.64
Children		44	11	0.25 (0.13-0.39)			-				3.48
Lewis et al, ⁴⁴ 2020	Utah and										
Adults	Wisconsin, US	120	33	0.28 (0.20-0.36)				_			6.34
Children		68	19	0.28 (0.18-0.39)							4.46
van der Hoek et al, 60 2020a	Netherlands	67	22	0.24 (0.22, 0.46)			_				F 66
Childron		107	23	0.34 (0.23-0.46)		_					5.00
	Zhaijang Dravinca	107	24	0.22 (0.13-0.31)							J.04
Adults	China	510	108	0 21 (0 18-0 25)			_				7 08
Children		325	43	0.13 (0.10-0.17)			-				8.69
ling et al. ³⁹ 2020	Guangzhou China										
Adults	Guangenou, enna	412	85	0.21 (0.17-0.25)							7.03
Children		125	8	0.06 (0.03-0.11)	-						7.51
Lyngse et al, ⁴⁹ 2020	Denmark										
Adults		1367	257	0.19 (0.17-0.21)			-				7.22
Children		859	114	0.13 (0.11-0.16)							9.76
Li et al, ⁴⁵ 2020	Wuhan, China										
Adults		292	60	0.21 (0.16-0.25)		-					6.92
Children		100	4	0.04 (0.01-0.09)	-	<u> </u>					7.51
Bi et al, ²² 2020	Shenzhen, China	462	61	0.12 (0.10.0.10)							7.00
Adults		462	61	0.13 (0.10-0.16)							7.09
	Durrani	103	10	0.10 (0.06-0.15)							7.66
Adults	Brunei	179	16	0.09(0.05-0.14)							6.86
Children		85	12	0.14 (0.07-0.22)							5 71
Laxminarayan et al, ⁴² 2020	Tamil Nadu and	55									5.7 ±
Adults	Andhra Pradesh, India	2671	245	0.09 (0.08-0.10)							7.28
Children		941	85	0.09 (0.07-0.11)		-					9.93
Adults estimate				0.283 (0.202-0.371)			\sim	•			100
Children estimate				0.168 (0.123-0.217)		<					100
					0		0.25	0.5	j	0.	75
							SAR	(95% CI)			

Point sizes are an inverse function of the precision of the estimates and bars correspond to 95% Cls.

^a Study of family contacts.

contacts in the household. Estimated mean household secondary attack rate for households with 1 contact (41.5%; 95% Cl, 31.7%-51.7%) was significantly higher than households with at least 3 contacts (22.8%; 95% Cl, 13.6%-33.5%; P < .001) but not different than households with 2 contacts (38.6%; 95% Cl, 17.9%-61.6%). There was significant heterogeneity in secondary attack rates between studies with 1 contact ($I^2 = 52.9\%$; P = .049), 2 contacts ($I^2 = 93.6\%$; P < .001), or 3 or more contacts ($I^2 = 91.6\%$; P < .001). Information was not available on household crowding (eg, number of people per room).

eFigure 14 in the Supplement summarizes 7 studies⁷⁶⁻⁸² reporting household secondary attack rates for SARS-CoV, and 7 studies⁸³⁻⁸⁹ for Middle East respiratory syndrome coronavirus (MERS-CoV). Estimated mean household secondary attack rate was 7.5% (95% CI, 4.8%-10.7%) for SARS-CoV and 4.7% (95% CI, 0.9%-10.7%) for MERS-CoV (eTable 7 in the Supplement), both lower than the household secondary attack rate of 16.6% for SARS-CoV-2 in this study (*P* < .001). The SARS-CoV-2 secondary attack rate was also higher than secondary attack rates reported for HCoV-NL63 (0-12.6%), HCoV-OC43 (10.6-13.2%), HCoV-229E (7.2-14.9%), and HCoV-HKU1 (8.6%).⁹⁰⁻⁹² Household secondary attack rates for SARS-CoV-2 were within the mid-range of household secondary attack rates reported for influenza, which ranged from 1% to 38% based on polymerase chain reaction-confirmed infection.⁹³

Discussion

We synthesized the available evidence on household studies of SARS-CoV-2. The combined household and family secondary attack rate was 16.6% (95% CI, 14.0%-19.3%), although with significant heterogeneity between studies. This point estimate is higher than previously observed

Figure 4. Secondary Attack Rates (SAR) of Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2) for Household and Family Contacts by Relationship to Index Case

Source	Location	Participants, No.	with SARS-CoV-2 infection, No.	SAR (95% CI)				Weigh %
Wu et al, ⁶⁵ 2020	Zhuhai, China							
Spouse		23	12	0.52 (0.32-0.72)				10.44
Other		120	36	0.30 (0.22-0.39)		<mark>-</mark>		11.32
Sun et al, ⁵⁸ 2020 ^a	Zhejiang Province, China							
Spouse		119	76	0.64 (0.55-0.72)				17.83
Other		479	113	0.24 (0.20-0.28)				16.41
ewis et al, ⁴⁴ 2020.	Utah and Wisconsin, US							
Spouse		33	11	0.33 (0.18-0.50)				12.65
Other		155	41	0.26 (0.20-0.34)				12.62
(in et al, ⁶⁷ 2020	Qingdao Municipal, China							
Spouse		16	4	0.25 (0.06-0.50)				9.28
Other		90	15	0.17 (0.10-0.25)				10.94
Li et al, ⁴⁵ 2020	Wuhan, China							
Spouse		90	25	0.28 (0.19-0.38)				17.18
Other		202	35	0.17 (0.12-0.23)	—			14.25
iu et al, ⁴⁶ 2020.	Guangdong Province, China							
Spouse		563	131	0.23 (0.20-0.27)				20.53
Other		1878	199	0.11 (0.09-0.12)	-			18.46
Chaw et al, ²⁶ 2020	Brunei							
Spouse		31	13	0.42 (0.25-0.60)				12.09
Other		233	15	0.06 (0.04-0.10)				16.00
Spouse estimate				0.378 (0.258-0.505)				100
Other estimate				0.178 (0.117-0.248)				100
					0	0.25	0.5	0.75
						SAR (95	5% CI)	

Point sizes are an inverse function of the precision of the estimates and bars correspond to 95% CIs.

^a Study of family contacts.

secondary attack rates for SARS-CoV and MERS-CoV. Households are favorable environments for transmission. They are what are known as 3Cs environments, as they are closed spaces, where family members may crowd and be in close contact with conversation.⁹⁴ There may be reduced use of personal protective equipment relative to other settings.

That secondary attack rates were not significantly different between household and family contacts may indicate that most family contacts are in the same household as index cases. Household and family contacts are at higher risk than other types of close contacts, and risks are not equal within households. Spouses were at higher risk than other family contacts, which may explain why the secondary attack rate was higher in households with 1 vs 3 or greater contacts. Spouse relationship to the index case was also a significant risk factor observed in studies of SARS-CoV and H1N1.^{82,95} This may reflect intimacy, sleeping in the same room, or longer or more direct exposure to index cases. Further investigation is required to determine whether sexual contact is a transmission route. Although not directly assessed, household crowding (eg, number of people per room) may be more important for SARS-CoV-2 transmission than the total number of people per household, as has been demonstrated for influenza.⁹⁶⁻⁹⁸

The finding that secondary attack rates were higher to adult contacts than to child contacts is consistent with empirical and modeling studies.^{99,100} Lower infection rates in children may be

Figure 5. Secondary Attack Rates (SAR) of Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2) by the Number of People Living in the Same Household as the Index Case

Source	Location	Participants, No.	Participants with SARS-CoV-2 infection, No.	SAR (95% CI)								Weight %
Rosenberg et al, ⁵⁵ 2020	New York, US											
1 Contact		31	13	0.42 (0.25-0.60)		_						12.20
2 Contacts		30	18	0.60 (0.42-0.77)								15.53
≥3 Contacts		282	100	0.35 (0.30-0.41)				_				17.35
Lopez Bernal et al, ⁴⁷ 2020	UK											
1 Contact		77	38	0.49 (0.38-0.61)			-					20.20
2 Contacts		106	43	0.41 (0.31-0.50)								18.49
≥3 Contacts		289	80	0.28 (0.23-0.33)		-	-					17.54
Wu et al. ⁶⁵ 2020	Zhuhai. China											
1 Contact	,	5	2	0.40 (0.02-0.86)								2.74
2 Contacts		14	8	0.57 (0.30-0.82)								12.32
≥3 Contacts		124	38	0.31 (0.23-0.39)			-	_				15.12
Wang et al. ⁶¹ 2020	Wuhan. China											
1 Contact	,	27	15	0.56 (0.36-0.74)			_					11.05
2 Contacts		21	15	0.71 (0.50-0.89)				-		-		14.50
≥3 Contacts		56	17	0.30 (0.19-0.43)			-					11.75
Lyngse et al. ⁴⁹ 2020	Denmark											
1 Contact		368	103	0.28 (0.24-0.33)		-	-					31.92
2 Contacts		432	64	0.15 (0.12-0.18)	-	-						19.72
≥3 Contacts		1426	204	0.14 (0.13-0.16)								19.42
Arnedo-Pena et al, ²⁰ 2020	Castellon, Spain											
1 Contact	,	92	40	0.43 (0.33-0.54)				-	_			21.89
2 Contacts		173	16	0.09 (0.05-0.14)								19.43
≥3 Contacts		397	27	0.07 (0.05-0.10)								18.82
1 Contact estimate				0.415 (0.317-0.517)					-			100
2 Contacts estimate				0.386 (0.179-0.616)								100
≥3 Contacts estimate				0 228 (0 136-0 335)								100

Point sizes are an inverse function of the precision of the estimates, and bars correspond to 95% Cls.

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SAR (95% CI)

attributed to asymptomatic or mild disease, reduced susceptibility from cross-immunity from other coronaviruses, ¹⁰¹ and low case ascertainment, ¹⁰² but the difference persisted in studies in which all contacts were tested regardless of symptoms. Higher transmission rates to adults may be influenced by spousal transmission. Given the increased risk to spousal contacts, future studies might compare child contacts and nonspouse adult contacts to ascertain whether this difference persists. Limited data suggest children have not played a substantive role in household transmission of SARS-CoV-2.^{40,103-105} However, a study in South Korea of 10 592 household contacts noted relatively high transmission from index cases who were aged 10 to 19 years.⁵¹ Although children seem to be at reduced risk for symptomatic disease, it is still unclear whether they shed virus similarly to adults.¹⁰⁶

We did not find associations between household contact or index case sex and secondary transmission. The World Health Organization reports roughly even distribution of SARS-CoV-2 infections between women and men worldwide, with higher mortality in men.¹⁰⁷

We found significantly higher secondary attack rates from symptomatic index cases than asymptomatic or presymptomatic index cases, although less data were available on the latter. The lack of substantial transmission from observed asymptomatic index cases is notable. However, presymptomatic transmission does occur, with some studies reporting the timing of peak infectiousness at approximately the period of symptom onset.^{108,109} In countries where infected individuals were isolated outside the home, this could further alter the timing of secondary infections by limiting contacts after illness onset.¹¹⁰

Household secondary attack rates were higher for SARS-CoV-2 than SARS-CoV and MERS-CoV, which may be attributed to structural differences in spike proteins,¹¹¹ higher basic reproductive rates,¹¹² and higher viral loads in the nose and throat at the time of symptom onset.¹¹³ Symptoms associated with MERS-CoV and SARS-CoV often require hospitalization, which increases nosocomial transmission, whereas less severe symptoms of SARS-CoV-2 facilitate community transmission.¹¹³ Similarly, presymptomatic transmission was not observed for MERS-CoV or SARS-CoV.^{114,115}

Limitations

Our study had several limitations. The most notable is the large amount of unexplained heterogeneity across studies. This is likely attributable to variability in study definitions of index cases and household contacts, frequency and type of testing, sociodemographic factors, household characteristics (eg, density, air ventilation), and local policies (eg, centralized isolation). Rates of community transmission also varied across locations. Given that studies cannot always rule out infections from outside of the home (eg, nonhousehold contacts), household transmission may be overestimated. For this reason, we excluded studies that used antibody tests to diagnose household contacts. Furthermore, many analyses ignored tertiary transmission within the household, classifying all subsequent cases as secondary to the index case. Eighteen

studies^{19-21,24,25,28,29,33,34,41,47,50,53,56,58,59,61,64} involved testing only symptomatic household contacts, which would miss asymptomatic or subclinical infections, although secondary attack rate estimates were similar across studies testing all vs only symptomatic contacts.

Important questions remain regarding household spread of SARS-CoV-2. Chief among them is the infectiousness of children to their household contacts and the infectiousness of asymptomatic, mildly ill, and severely ill index cases. This study did not provide additional elucidation of factors influencing intergenerational spread. People unable to work at home may have greater risk of SARS-CoV-2 exposure, which may increase transmission risk to other household members. There may be overdispersion in the number of secondary infections per index case, which could be caused by variations in viral shedding, household ventilation, or other factors.

Conclusions

The findings of this study suggest that households are and will continue to be important venues for transmission, even where community transmission is reduced. Prevention strategies, such as

increased mask-wearing at home, improved ventilation, voluntary isolation at external facilities, and targeted antiviral prophylaxis, should be further explored.

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Corresponding Author: Zachary J. Madewell, Department of Biostatistics, University of Florida, PO Box 117450, Gainesville, FL 32611 (zmadewell@ufl.edu).

Author Affiliations: Department of Biostatistics, University of Florida, Gainesville (Madewell, Yang, Longini, Dean); Fred Hutchinson Cancer Research Center, Seattle, Washington (Halloran); Department of Biostatistics, University of Washington, Seattle (Halloran).

Author Contributions: Drs Madewell and Dean had full access to all of the data in the study and take responsibility for the integrity of the data and the accuracy of the data analysis.

Concept and design: Madewell, Longini, Dean.

Acquisition, analysis, or interpretation of data: All authors.

Drafting of the manuscript: Madewell, Longini, Dean.

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SUPPLEMENT.

eFigure 1. PRISMA Flow Diagram for Review of Household Secondary Attack of SARS-CoV-2, MERS-CoV, SARS-CoV, and Other Coronaviruses

eFigure 2. Secondary Attack Rates of SARS-CoV-2 for Studies of Close Contacts

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Coronavirus 2 (SARS-CoV-2) for Household Contacts and Family Contacts

eAppendix 1. Eligibility Criteria

eAppendix 2. Data Extraction

eAppendix 3. Additional Description of Studies

eAppendix 4. Additional Description of Risk Factors

eReferences.