Infection Risk Prediction in Healthcare Settings: Lessons from COVID-19 Contact Tracing

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ABSTRACT

Objective: Contact tracing aids epidemic control by enabling early detection and isolation without overburdening healthcare systems despite potential challenges. This study aimed to evaluate the practical application of contact and risk assessment-based models in predicting SARS-CoV-2 infection following exposure among healthcare workers in a large tertiary public university hospital in Türkiye.

Material and Methods: The study was designed as a retrospective cohort study, including contact tracing data from 3389 exposed healthcare workers from March 23, 2020, to October 22, 2021. Contact-based (mask use, contact duration and distance) and exposure risk-assessment-based (low, medium, high-risk) models with and without having symptoms were generated using logistic regression. SARS-CoV-2 infection was defined as having a positive SARS-CoV-2 RT-PCR test result. Adjustments were made to the models for demographic and occupational variables, previous infection, and vaccination. Model parameters were compared.

Results: Of 3389 exposed healthcare workers, 2451 underwent RT-PCR testing. Among those tested, RT-PCR positivity was 5.9% (144/2451). Lack of personal protective equipment use (odds ratio [OR]=1.64, 95% confidence interval [CI]=1.03-2.66) and \geq 15 minutes of contact duration (1.89, 1.21-3.09) were significantly associated with RT-PCR positivity. In the risk-assessment model, being a high-risk contact increased the odds of RT-PCR positivity (OR=2.76, 95% CI=1.61-5.03). Adding the presence of symptoms to contact-based and risk assessment models improved model parameters (Akaike information criterion [AIC]: from 1086.1 to 1083.1; Tjur's R²: from 0.016 to 0.019, respectively).

Conclusion: The inclusion of being symptomatic improved the contact-based and risk assessment-based models. Institutions should be encouraged to incorporate symptom inquiries into risk assessment protocols in response to newly emerging respiratory virus epidemics. Institutions lacking the capacity for extensive contact tracing are recommended, at minimum, to track symptomatic exposed workers for epidemic control.

Keywords: COVID-19, health personnel, contact tracing, risk assessment, healthcare worker

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Received: January 05, 2024 Accepted: February 19, 2024 Published: March 08, 2024

Suggested citation:

Keskin S, Emecen AN, Ergör A. Infection risk prediction in healthcare settings: Lessons from COVID-19 contact tracing. Infect Dis Clin Microbiol. 2024:1:44-54.

DOI: 10.36519/idcm.2024.315



INTRODUCTION

ontact tracing has been recognized as a critical process in controlling infectious disease J epidemics. Its objectives include early identification of potentially infectious cases and prevention of the emergence of new clusters (1). In the context of the severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) infection, in-hospital contact-tracing efforts ensure the well-being of frontline healthcare workers (HCWs) to maintain essential healthcare services (2). For this reason, many hospitals from different nations implemented infection risk assessments for HCWs according to international/national guidelines, institutional needs, and resource capacities (3-6). Yet, the key measures to evaluate infection risk were based on exposure: use of masks, contact distance and duration (7, 8).

Large-scale hospital-based studies found household exposure (3, 9), contact with a patient (3), being symptomatic (5), and inadequate use of personal protective equipment (PPE) (9) as the significant factors that contributed to testing positive for SARS-CoV-2 after exposure. In addition, three studies in Türkiye found that high-risk exposures (<1-meter distance, >15-minute duration, no mask) had a higher risk of infection (10-12). The work environment and contact type also affected exposure and SARS-CoV-2 infection risk (12); for example, the index case was more likely to be another coworker (11, 13).

Early on in the SARS-CoV-2 pandemic, international organizations proposed infection risk assessment algorithms for HCWs. The World Health Organization (WHO) recommended infection risk assessment using only the exposure risk. The WHO classified exposure risk as low and high based on contact characteristics (i.e., use of masks, contact distance, and duration). However, the complexity of in-hospital contact scenarios and not including symptom assessment in the algorithms resulted in risk categorization and management limitations. Moreover, contact tracing and exposure risk assessment were time-consuming and resource-intensive processes requiring trained personnel to ensure standardized evaluation (14, 15).

Given these challenges, a need for an accurate and easy-to-use contact tracing strategy for respiratory

virus infections that require a rapid, global response in the future remains. This study aimed to evaluate models derived from real-world hospital-based contact tracing data to predict SARS-CoV-2 infection in a simplified manner.

MATERIALS AND METHODS

In this retrospective cohort study, we examined all SARS-CoV-2 infection and contact tracing data collected between March 23, 2020, and October 22, 2021, from HCWs at the Dokuz Eylül University Research and Application Hospital in İzmir, Türkiye. The hospital had a workforce of nearly 4000 employees, and during the pandemic, provided both outpatient and inpatient care for coronavirus disease 2019 (COVID-19) patients.

Dokuz Eylül University Non-Interventional Research Ethics Committee approved the study with the number 2022/17-15.

Risk Assessment Procedure

Upon identifying a confirmed COVID-19 case among HCWs, our institution implemented rigorous contact tracing and risk assessment procedures. If the primary case was an HCW, a list of potentially exposed coworkers was compiled for contact tracing, and the contact tracing period started two days before symptom onset for symptomatic cases. For asymptomatic cases, contact tracing started two days before the first positive SARS-CoV-2 RT-PCR (RT-PCR) test result. All coworkers potentially exposed

HIGHLIGHTS

- Infection risk prediction is enhanced by including symptomatic status among healthcare workers.
- There was no significant relationship between contact distance and increased infection risk.
- Personal protective equipment use reduced RT-PCR positivity in healthcare settings.
- Contact duration over 15 minutes increased transmission risk significantly.
- Focusing on protecting vulnerable healthcare workers was found to be essential instead of a zero-infection policy.

	Primary case	line of second s	>1-m contact distance		≤1-m contact distance	
Algorithm		Use of masks	Low intensity	High intensity	Low intensity	High intensity
		CP(+) / E(+)	No risk	Low	Low	Low
la stituti se sl	HCW	CP(+) / E(-)	Low	Low	Low	Medium
Institutional		CP(-) / E(+)	Low	Medium	Medium	High
		CP(-) / E(-)	Medium	Medium	High	High
МоН	Patient	Pt(+) / E(+)	No Risk	No Risk	No Risk	No Risk
		Pt(+) / E(-)	Low	Medium	Low	Medium
		Pt(-) / E(+)	Low	Medium	Low	Medium
		Pt(-) / E(-)	Low	High	Low	High

Table 1. Levels of exposure risk for HCWs: algorithms from institutional and Ministry of Health sources based on the primary case, use of masks, contact distance and intensity.

MoH: Ministry of Health, HCW: Healthcare workers, CP: COVID-19 HCW, E: Exposed HCW, Pt: COVID-19 patient. (+): using a mask; (-): not using a mask.

High Intensity: For HCW-to-HCW contacts, activities done face-to-face such as consuming meals, drinking tea or coffee, smoking, conversing face-to-face, coughing, sneezing, or any action sustained for a duration exceeding 15 minutes. For patient-to-HCW contacts, conducting respiratory procedures such as non-invasive ventilation, specimen collection, aspiration, nebulizer administration, cardiopulmonary resuscitation (CPR), oropharyngeal examination, ophthalmological assessments, endoscopy, bronchoscopy, and similar procedures.

Low Intensity: For HCW-to-HCW contacts, activities done face-to-face (such as consuming meals, drinking tea or coffee, smoking, conversing face-to-face, coughing, sneezing), including brief periods of rest lasting less than 15 minutes, sitting without conversing, and exchanging documents or objects while adhering to hand hygiene protocols. For patient-to-HCW contacts, activities are not included in the high-intensity category.

to an infected case during these periods were identified. Trained public health residents conducted telephone interviews with the exposed HCWs under the supervision of senior occupational health and epidemiology professionals. Symptoms associated with the illness were evaluated as fever, cough, respiratory symptoms, myalgia and malaise, loss of taste and smell, and diarrhea.

We utilized exposure risk assessment using two distinct algorithms: one for patient-to-HCW contacts and the other for HCW-to-HCW contacts. Patient-to-HCW contacts were assessed following the guidelines of the Turkish Ministry of Health (16). This national algorithm included the patient's mask use, the HCW's mask use, and PPE suitability for healthcare procedures. Since no national guidelines were available for evaluating contacts among HCWs, our institution developed an algorithm for HCW-to-HCW interactions, which was aligned with international guidelines and expert opinions within our institution. The risk assessment algorithm for HCW-to-HCW contacts included contact duration, distance, mask use, and the intensity of interactions, such as handshaking or having meals or coffee together (13). Both algorithms are summarized in Table 1.

Following the assessment, exposure risk levels were classified as low, medium, or high. Low or medium-risk HCWs were permitted to continue working while being monitored for symptoms for 14 days from the last contact with a SARS-CoV-2 infected person. Medium-risk HCWs underwent an RT-PCR test on the seventh day of the first contact. Highrisk HCWs were subject to work restrictions and home quarantined until RT-PCR testing was conducted on the seventh day. After the assessment, informative text messages detailing work restrictions, quarantine procedures, and scheduled RT-PCR testing dates were sent to the HCWs. Routine RT-PCR testing was recommended for medium to **Table 2.** Characteristics of SARS-CoV-2 exposed HCWs(n=3389).

	n (%)			
Gender (female)	1989 (58.7)			
Age, mean±SD	35.2±9.0			
Profession				
Physician	843 (24.9)			
Nurse	1033 (30.5)			
Allied healthcare personnel	140 (4.1)			
Auxiliary staff	1106 (32.6)			
Support staff	267 (7.9)			
Previously infected	100 (3.0)			
Two-dose vaccination	380 (11.2)			
Case type				
Colleague	2292 (67.6)			
Patient	1097 (32.4)			
Contact distance				
≥1 meter	1038 (30.6)			
<1 meter	2351 (69.4)			
Contact duration				
<15 minutes	1329 (39.2)			
≥15 minutes	2060 (60.8)			
PPE use	1929 (56.9)			
Use of surgical masks	1695 (50)			
Use of respiratory masks	234 (6.9)			
Use of goggles or face shields	85 (2.5)			
Having symptoms	824 (24.3)			
Risk category				
Low	1185 (35.0)			
Medium	1173 (34.6)			
High	1031 (30.4)			
SARS-CoV-2 RT-PCR tested	2451 (72.3)			
SARS-CoV-2 RT-PCR positive*	144 (5.9)			

SD: Standard deviation, PPE: Personal protective equipment. *Test positivity calculated based on those tested.

high-risk HCWs but not for low-risk HCWs. Regardless of risk category, HCWs experiencing any symptoms were instructed to immediately report to the occupational health outpatient clinic for the sameday RT-PCR testing.

Study Variables

Professions were categorized as physicians (medical doctors, dentists, pharmacists), nurses, allied healthcare personnel (laboratory technicians, dieticians, social workers, physiotherapists, anesthetic technicians), auxiliary staff (administrative staff, secretaries, office workers), and support staff (cleaning staff, porters, repair workers, garden maintenance staff, kitchen workers, security staff). Symptoms identified in the risk assessment phone interview were acknowledged as symptom presence. HCWs vaccinated with at least two doses of inactivated or mRNA COVID-19 vaccines 14 days before the exposure was classified as two-dose vaccinated. A positive RT-PCR result at least 14 days before the first exposure date was defined as a previous infection. The use of respiratory or surgical masks constituted PPE use. We followed the exposed workers until 14 days after the last contact with a SARS-CoV-2-infected person and assessed their RT-PCR test results within that timeframe. Individuals with all negative results were considered PCR-negative, and those with at least one positive result were regarded as PCR-positive, regardless of any negative results.

Statistical Analysis

Categorical variables were presented as numbers with percentages (n, %) and were analyzed using the chi-squared test. According to the normal distribution assessment, the Student's t-test or the Mann-Whitney U test was conducted to compare continuous variables. Factors associated with reporting a positive PCR test were evaluated using logistic regression among the contacted HCWs who had RT-PCR testing. The measure of risk was the odds ratio (OR) with 95% confidence interval (CI).

To investigate the practical use of contact and risk assessment-based models incorporating symptoms and other variables, we built four different models with the "enter" method. Model 1 and Model 2 were contact-based models. Model 1 involved an assessment of the globally accepted measures: use of PPE, contact duration, and distance. Model 2 incorporated Model 1 variables plus the presence of model utilizing the risk categories (low, medium, symptoms. For Model 3, we built a risk assessment and high risk) as summarized in Table 1. In Model 4,

Table 3. Comparison of the HCWs who tested RT-PCR negative and positive within 14 days after the last exposure (n=2451).

	PCR negative n=2307 n (%)	PCR positive n=144 n (%)	p*	OR (95% CI)
Gender				
Female	1369 (59.3)	87 (60.4)	0.07	Ref.
Male	938 (40.7)	57 (39.6)	- 0.87 -	0.96 (0.68-1.35)
Age, mean±SD	34.9±8.8	33.9±9.2	0.17	0.99 (0.97-1.01)
Profession				
Physician	569 (24.7)	28 (19.4)		Ref.
Nurse	697 (30.2)	41 (28.5)		1.19 (0.73-1.98)
Allied healthcare personnel	98 (4.2)	5 (3.5)	0.22	1.06 (0.35-2.61)
Auxiliary staff	747 (32.4)	60 (41.7)		1.63 (1.03-2.62)
Support staff	196 (8.5)	10 (6.9)		1.05 (0.47-2.14)
Previously infected				
Yes	65 (2.82)	3 (2.08)		Ref.
No	2242 (97.2)	141 (97.9)	0.80	1.30 (0.47-5.54)
Two doses vaccinated	l			
Yes	261 (11.3)	13 (9.03)	0.40	Ref.
No	2046 (88.7)	131 (91.0)	- 0.48 -	1.27 (0.74-2.40)
Case type				
Colleague	1703 (73.8)	117 (81.2)		Ref.
Patient	604 (26.2)	27 (18.8)	0.06	0.65 (0.42-0.99)
Contact distance				
≥1 meter	581 (25.2)	25 (17.4)	0.04	Ref.
<1 meter	1726 (74.8)	119 (82.6)	- 0.04 -	1.59 (1.04-2.53)
Contact duration	· · · · · · · · · · · · · · · · · · ·			
<15 minutes	688 (29.8)	23 (16.0)	0.001	Ref.
≥15 minutes	1619 (70.2)	121 (84.0)	- 0.001 -	2.22 (1.44-3.59)
PPE use	· · · · · · · · · · · · · · · · · · ·			
Yes	1046 (45.3)	42 (29.2)	0.001	Ref.
No	1261 (54.7)	102 (70.8)	<0.001	2.01 (1.40-2.94)

Yes	919 (39.8)	35 (24.3)	-0.001	Ref.
No	1388 (60.2)	109 (75.7)	- <0.001 -	2.06 (1.41-3.08)
Use of respiratory masks				
Yes	127 (5.50)	7 (4.86)	0.00	Ref.
No	2180 (94.5)	137 (95.1)	0.89 -	1.12 (0.55-2.70)
Use of goggles or face shields				
Yes	25 (1.08)	3 (2.08)	0.22	Ref.
No	2282 (98.9)	141 (97.9)	0.23	0.49 (0.17-2.17)
Having symptoms				
No	1585 (68.7)	79 (54.9)	0.001	Ref.
Yes	722 (31.3)	65 (45.1)	0.001	1.81 (1.28-2.54)
Risk assessment category				
Low	396 (17.2)	16 (11.1)		Ref.
Medium	995 (43.1)	31 (21.5)	<0.001	0.77 (0.42-1.46)
High	916 (39.7)	97 (67.4)		2.60 (1.55-4.64)

Continue to Table 3

*Chi-square or Student-t test.

OR: Odds ratio, PPE: Personal protective equipment.

we included Model 3 variables plus the presence of symptoms. All models were adjusted to account for demographic variables (age, gender), occupational variables (profession, case type), and COVID-19 history (previous infection, vaccination). To compare the discriminative power and fits of the models, we calculated the AUC (area under the curve), Tjur's R², and Akaike's information criterion (AIC). To evaluate the diagnostic accuracy, we computed the diagnostic odds ratio (DOR) for various thresholds of the models, ranging from 1% to 15%. The DOR was calculated as (Sensitivity / [1- Sensitivity]) / ([1-Specificity] / Specificity). The DOR remains unaffected by the prevalence of the disease (17). It quantifies the odds of positive test results in individuals with RT-PCR positivity compared to the odds of positive test results in those without RT-PCR positivity.

As a sensitivity analysis, we repeated all models with inverse probability weighting (IPW) to address potential selection bias stemming from unequal probabilities of RT-PCR testing among risk categories. The variables (age, having symptoms, risk category, and case type) used for IPW were profession and the statistically significant variables in the comparison of RT-PCR-tested and not-tested workers (Supplementary Table 1). Statistical analyses were conducted using R version 4.2.3, with double-sided *p*-values of less than 0.05 considered significant.

RESULTS

A total of 3389 contacts that occurred from March 23, 2020, to October 22, 2021, were included in the study. The characteristics of exposed HCWs are presented in Table 2. The majority of contacts occurred among colleagues (67.6%). The mean age of the exposed HCWs was 35.2±9.0, and the majority were women (58.7%). Of the contacts, 69.4% occurred within 1 meter of distance, and 60.8% lasted 15 minutes or more. PPE was used in 56.9% of the contacts. Exposure risk was reported as low in 35.0% of the contacts, followed by medium (34.6%) and high risk (30.4%). Of the contacts, 24.3% were experiencing symptoms at the time of risk assessment. Among the 2451 (72.3%) exposed HCWs who had SARS-CoV-2 PCR testing, 144 (5.9%) had a positive RT-PCR test result. Clinically, all cases were asymptomatic or with mild illness with no intensive care unit (ICU) admission or death.

	Contact-based models		Risk assessment-based models		
	Model 1	Model 2	Model 3	Model 4	
	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)	
No PPE use	1.64 (1.03-2.66)	1.61 (1.01-2.62)	-	-	
<1 meter contact distance	1.40 (0.89-2.28)	1.32 (0.84-2.16)	-	-	
≥ 15-minute contact duration	1.89 (1.21-3.09)	1.83 (1.17-3.00)	-	-	
Having symptoms	-	1.50 (1.05-2.14)	-	1.47 (1.03-2.09)	
Risk category					
Low	-	-	Ref.	Ref.	
Medium	-	-	0.77 (0.42-1.47)	0.77 (0.42-1.47)	
High	-	-	2.76 (1.61-5.03)	2.62 (1.53-4.80)	
Model parameters*				·	
AUC, 95% CI	0.65 (0.60-0.69)	0.65 (0.61-0.70)	0.67 (0.63-0.72)	0.67 (0.63-0.72)	
Accuracy, %	50.2	64.0	60.0	55.7	
Sensitivity, %	75.7	60.4	70.1	74.3	
Specificity, %	48.6	64.4	58.8	54.5	
PPV, %	8.4	9.6	9.6	9.3	
NPV, %	97.0	96.3	96.9	97.1	
AIC	1086.1	1083.1	1069.4	1066.9	
Tjur's R ²	0.016	0.019	0.022	0.025	

Table 4. Logistic regression results using contact-based and risk assessment-based models predicting SARS-CoV-2 RT-PCR positivity within 14-days after the last exposure.

OR: Odds ratio, CI: Confidence interval, PPE: Personal protective equipment, AUC: Area under the curve, PPV: Positive predictive value, NPV: Negative predictive value, AIC: Akaike information criterion.

Model 1 included PPE use, contact distance, duration, and adjustment variables.

Model 2 included PPE use, contact distance, duration, having symptoms, and adjustment variables.

Model 3 included the risk category and adjustment variables.

Model 4 included the risk category, having symptoms, and adjustment variables.

Adjustment variables: Demographic variables (age, gender), occupational variables (profession, case

type) and COVID-19 history (previous infection, vaccination).

*Thresholds are set to maximize Youden's index for each model.

A comparison of HCWs who tested positive and negative for PCR after exposure is shown in Table 3. Auxiliary staff showed an increased risk of testing positive compared to physicians, with an OR of 1.63 (95% CI=1.03-2.62). Previously infected (p=0.80) and fully vaccinated individuals (p=0.48) demonstrated no significant difference in PCR positivity within 14 days post-exposure. Contact of < 1-meter (OR=1.59, 95% CI=1.04-2.53), >15 minutes duration (OR=2.22, 95% CI=1.44-3.59), no use of PPE (OR=2.01, 95% CI=1.40-2.94), and having symptoms (OR=1.81, 95% CI=1.28-2.54) were associated with a significant increase in the odds of PCR positivity.

Table 4 presents logistic regression results for the PCR positivity within 14 days after the last exposure. In the contact-based Model 1, no PPE use (OR=1.64, 95% CI=1.03-2.66) and \geq 15 minutes of contact duration (OR=1.89, 95% CI=1.21-3.09) were significantly associated with PCR positivity. In Mod-

Threshold	Contact-based models		Risk assessment-based models		
Thi conord	Model 1	Model 2	Model 3	Model 4	
1%	-	-	-	-	
2%	5.80	2.09	1.86	2.49	
3%	1.93	2.46	2.55	2.81	
4%	2.45	2.81	3.00	3.47	
5%	2.71	2.45	3.33	3.04	
6%	2.29	2.58	2.89	2.99	
7%	2.22	2.53	2.61	2.62	
8%	2.38	2.53	2.65	2.52	
9%	2.55	2.60	2.79	2.93	
10%	2.87	2.71	3.01	3.01	
11%	2.73	3.25	2.99	3.48	
12%	4.39	3.49	3.69	3.96	
13%	4.02	4.28	5.29	3.67	
14%	3.76	6.08	4.11	5.41	
15%	4.03	6.65	6.49	9.46	

Table 5. The diagnostic odds ratios of the contact-based or risk assessment-based model thresholds range from 1% to 15%.

Model 1 included PPE use, contact distance, duration, and adjustment variables. Model 2 included PPE use, contact distance, duration, having symptoms, and adjustment variables.

Model 3 included the risk category and adjustment variables.

Model 4 included the risk category, having symptoms and adjustment variables. Adjustment variables were: Demographic variables (age, gender), occupational variables (profession, case type) and COVID-19 history (previous infection, vaccination).

el 2, symptoms were added to Model 1 variables, which improved model parameters (AIC from 1086.1 to 1083.1; Tjur's R² from 0.016 to 0.019, respectively). According to the risk assessment-based Model 3, being evaluated as a high-risk contact increased the odds of PCR positivity (OR=2.76, 95% CI=1.61-5.03). Model 4, which combined risk assessment and the presence of symptoms, had the best fit compared to other models with an AIC of 1066.9 and Tjur's R² of 0.025. IPW models revealed similar results, as summarized in <u>Supplementary Table 2</u>.

Table 5 shows the DOR calculated for four models, with varied thresholds ranging from 1% to 15%. At the minimum threshold (2%), Model 1 had the high-

est DOR of 5.80, and at the maximum threshold (15%), Model 4 had the highest DOR of 9.46.

DISCUSSION

This study evaluated factors associated with developing RT-PCR positivity among exposed HCWs by analyzing one of the largest single-center contact tracing and exposure risk data from the COVID-19 pandemic. Our findings indicate that exposure risk assessment variables (i.e., use of PPE, contact distance and duration) significantly predict the development of infection, and whether the contact has symptoms or not improves this prediction.

Rigorous contact tracing can limit disease spread but may not be feasible once a substantial surge of new cases is reported daily (18). Managing a large number of in-hospital contacts requires significant human resources, timely communication, meticulous record-keeping, data management, and coordination efforts. As the full quarantine of exposed workers may be unrealistic, international organizations suggested exposure risk algorithms. Building upon the institutionally modified algorithms, our study found that a lack of PPE use and ≥15 minutes of contact duration increased the risk of RT-PCR positivity in contact-based models, but no association was found for contact distance.

Our findings are consistent with at least four meta-analyses (19-22), which also concluded that masks effectively prevent infection in HCWs. The data used in the studies was inherently based on one infector-infectee pair's contact details. When there was widespread community transmission, the true protective effect of masking could be limited as prolonged mask use is difficult to adhere to, causing discomfort and breathing difficulties (23, 24). For this reason, a more strategic use of masks could be considered, making it stricter in high-risk areas such as those dealing with immunosuppressed patients or ICUs while adopting a more relaxed approach in other areas.

Contrary to a meta-analysis conducted during the early stages of the pandemic (25), this study found that contact distance had no significant effect on infection. Theoretically, social distancing reduces the exposure of virus-containing droplets/aerosols from one person to another as it limits the frequency of people sharing the same space. However, accurately assessing the contact distance proves challenging during the risk assessment as it may fluctuate throughout the contact period. Most existing literature focuses on country-level measures and outcomes, leaving a gap to explore person-to-person transmission dynamics (26-28).

The duration of contact is essential for risk assessment (9-12). We found that \geq 15 minutes of contact increased the transmission risk. Integrating systematic pauses or breaks in prolonged healthcare activities could serve as a strategic intervention to limit direct contact and reduce transmission risk among HCWs. Controversially, for the workplace contacts of HCWs, Murugesan et al. did not identify a significant relationship between contact duration and symptom onset after exposure (29). Limiting contact duration during patient care is not always feasible in hospital settings, particularly in emergency care and surgeries. In such cases, controlling other measures, such as using masks and ventilation, should be critical to minimize transmission from a definite COVID-19 case.

Our risk assessment procedure indicated a connection between the presence of symptoms and transmission, which aligns with the findings of Mark et al. (5). Despite its significance, symptom status is not directly a part of risk assessment procedures (7, 8, 16). Incorporating symptom-based approaches into risk assessment models can pose challenges. Lack of confirming the probable COVID-19 case due to the not-perfect sensitivity of RT-PCR may lead to operational confusion of exposed HCW management. Additionally, the subjective nature of symptom reporting introduces prolonged sick leaves.

Similar to the findings of Durmaz et al. and Kalem et al., this study identified a 4.2% PCR positivity (144/3389) among contacts, with reported symptoms at 24.3% during interviews, notably higher (10, 11). During the pandemic, protective measures like prolonged mask-wearing may have led to symptoms (headaches, fatigue) that could be mistaken for COVID-19. Healthcare workers may have meticulously examined these symptoms, which could blend with genuine COVID-19 indicators. Additionally, PCR test sensitivity and sampling issues might have obstructed complete diagnostic accuracy. Despite our healthcare staff not being restricted by testing limitations, early national policies that curtailed PCR testing for asymptomatic cases may have impacted symptom reports. Furthermore, the presence of a group in the study that did not undergo testing could have also influenced the PCR positivity rate.

One of the important practical applications of the study lies in its ability to empower decision-makers with infection probabilities and discrimination power of the models based on real-world data. Through a user-friendly web-embedded platform, users can input relevant variables, enabling the models to generate real-time predictions for newly emerged respiratory virus infections that could require a response on a global scale. By utilizing this, the institutions can efficiently save time and personnel resources. We suggest lower probability thresholds and models with higher DORs for departments where infection-prone patients are admitted, such as hematology and oncology units, to minimize the HCW to patient transmission.

This study had several limitations. Initially, it was subject to recall bias, as contact details were based on personal statements. Subsequently, ascertainment bias could have arisen from unequal PCR testing probabilities among contact risk categories. We attempted to address selection bias by rebuilding inverse probability-weighted models. Furthermore, the single-center nature of the study may limit its generalizability to other hospital settings and other countries. To establish the utility of the models, external validation using datasets from different institutions is necessary. Due to the limited availability of variant analysis kits in our institution, comprehensive data on variants were lacking, potentially causing overlooked differences attributed to various variants. Lastly, this study may not have fully differentiated between workplace infections and community-based acquisitions. Additionally, it may not always be clear whether a contact was already positive prior to the interaction, independent of the exposure event. However, the policy within our institution allows for rapid PCR screening of symptomatic employees, which suggests that such cases are likely rare.

In conclusion, in this tertiary hospital study, the inclusion of being symptomatic made an improvement on the contact-based and risk assessment-based models. Based on these data, institutions with sufficient resources are encouraged to incorporate symptom inquiries into their risk assessment protocols in response to newly emerged respiratory virus infections that may seriously affect public health. Meanwhile, at minimum, institutions lacking the capacity for extensive contact tracing are recommended to track symptomatic exposed HCWs for outbreak control. Aiming for a zero-infection policy is extremely challenging and even unrealistic in the long term; hospitals should focus their mitigation efforts on evidence-based strategies, consequently enhancing both patient and HCW safety.

Ethical Approval: The Non-Interventional Research Ethics Committee of Dokuz Eylül University approved the study on May 11, 2022 with the decision number 2022/17-15.

Informed Consent: N.A.

Peer-review: Externally peer-reviewed

Author Contributions: Concept – S.K., A.N.E., A.E.; Design – S.K., A.E., A.N.E.; Supervision – S.K., A.E.; Funding – A.E.; Data Collection and/or Processing – S.K., A.N.E., A.E.; Analysis and/or Interpretation

– S.K., A.N.E.; Literature Review – S.K., A.N.E.; Writer – S.K., A.N.E., A.E.; Critical Reviews – S.K., A.N.E., A.E.

Conflict of Interest: The authors declare no conflict of interest.

Financial Disclosure: The authors declared that this study has received no financial support.

Acknowledgement: We would like to express our sincere gratitude to Dalya Güriş for her valuable suggestions and guidance.

REFERENCES

- 1 Coronavirus disease (COVID-19): Contact tracing [Internet]. Geneva: World Health Organization (WHO). [cited June 14, 2023]. Available from: https://www.who.int/news-room/questions-and-answers/item/coronavirus-disease-covid-19-contact-tracing
- 2 Breeher L, Boon A, Hainy C, Murad MH, Wittich C, Swift M. A Framework for sustainable contact tracing and exposure investigation for large health systems. Mayo Clin Proc. 2020;95(7):1432-44. [CrossRef]
- 3 Bailie CR, Leung VK, Orr E, Singleton E, Kelly C, Buising KL, et al. Performance of hospital-based contact tracing for COVID-19 during Australia's second wave. Infect Dis Health. 2022;27(1):15-22. [CrossRef]
- 4 Goniewicz K, Khorram-Manesh A, Hertelendy AJ, Goniewicz M, Naylor K, Burkle FM. Current response and management decisions of the European Union to the COVID-19 outbreak: A Review. Sustainability. 2020;12(9):3838. [CrossRef]
- 5 Mark E, Saidel-Odes L, Borer A, Nativ R, Sterer B, Yarmitsky S, Dreiher J. The appropriateness of the decision to quarantine healthcare workers exposed to a severe acute respiratory coronavirus virus 2 (SARS-CoV-2)-positive coworker based on national guidelines. Infect Control Hosp Epidemiol. 2023;44(6):920-5. [CrossRef]
- 6 Spiegel JM, Zungu M, Yassi A, Lockhart K, Wilson KS, Okpani AI, et al. Protecting healthcare workers during a pandemic: what can a WHO collaborating centre research partnership contribute? Rev Panam Salud Publica. 2023;47:e33. [CrossRef]

- 7 Interim guidance for managing healthcare personnel with SARS-CoV-2 infection or exposure to SARS-CoV-2 [Internet]. Atlanta: Centers for Disease Control and Prevention (CDC). [cited May 12, 2023]. Available from: <u>https://www.cdc.gov/coronavirus/2019-ncov/hcp/guidance-risk-assesment-hcp.html</u>
- 8 Interim guidance, risk assessment and management of healthcare workers in the context of COVID-19 [Internet]. Geneva: World Health Organization Western Pacific Region. [cited May 12, 2023]. Available from: <u>https://apps.who.int/iris/bitstream/ handle/10665/334366/WPR-DSE-2020-031-eng.pdf</u>
- 9 Sertcelik A, Cakir B, Metan G. Evaluation of risk factors for developing COVID-19 in healthcare professionals working at two university hospitals in Turkey. Work. 2023;74(3):799-809. [CrossRef]
- 10 Durmaz S, Kucuk EF, Simsek S, Durgun K, Karakas EB, Durusoy R. Outcomes of COVID-19 contact tracing in hospital healthcare workers: A retrospective cohort study. J Pak Med Assoc. 2022;72(4):707-13. [CrossRef]
- 11 Kaya Kalem A, Kayaaslan B, Eser F, Hasanoglu İ, Ayhan M, Coskun B, et al. Investigation of the relation between risk assessment of exposure and nosocomial SARS-CoV-2 transmission in healthcare workers: a prospective single-centre study. BMJ Open. 2022;12(1):e056858. [CrossRef]
- 12 Şahan C, Yıldırım D, Özyürek D, Çoban T, Kara ŞA, Öntaş E, et al. Sources of infection and risk of COVID-19 for healthcare workers at a tertiary hospital. Erciyes Med J. 2022;44(5):501-7. [CrossRef]

- 13 Emecen AN, Keskin S, Boncukcu Eren E, Yildirim Ustuner B, Celik SG, Suner AF, et al. Impact of social contacts on SARS-CoV-2 exposure among healthcare workers. Occup Med (Lond). 2022;72(1):10-6. [CrossRef]
- 14 Ho HJ, Zhang ZX, Huang Z, Aung AH, Lim WY, Chow A. Use of a real-time locating system for contact tracing of health care workers during the COVID-19 pandemic at an infectious disease center in Singapore: Validation study. J Med Internet Res. 2020;22(5):e19437. [CrossRef]
- 15 Hong P, Herigon JC, Uptegraft C, Samuel B, Brown DL, Bickel J, et al. Use of clinical data to augment healthcare worker contact tracing during the COVID-19 pandemic. J Am Med Inform Assoc. 2021;29(1):142-8. [CrossRef]
- 16 COVID-19 (SARS-CoV-2 Enfeksiyonu) temaslı takibi, salgın yönetimi, evde hasta izlemi ve filyasyon, 2021 [Internet]. Ankara: T.C. Sağlık Bakanlığı Halk Sağlığı Genel Müdürlüğü. [cited May 12, 2023]. Available from: <u>https://covid19.saglik.gov.</u> tr/Eklenti/41623/0/covid-19rehberitemaslitakibievdehastaizlemivefilyasyon-021021pdf.pdf
- 17 Glas AS, Lijmer JG, Prins MH, Bonsel GJ, Bossuyt PM. The diagnostic odds ratio: a single indicator of test performance. J Clin Epidemiol. 2003;56(11):1129-35. [CrossRef]
- 18 Operational considerations for adapting a contact tracing program to respond to the COVID-19 pandemic in non-U.S. settings, 2021 [Internet]. Atlanta: National Center for Immunization and Respiratory Diseases (U.S.). Division of Viral Diseases. [cited July 26, 2023]. Available from: <u>https://stacks.cdc.gov/view/cdc/110057</u>
- 19 Chen Y, Wang Y, Quan N, Yang J, Wu Y. Associations between wearing masks and respiratory viral infections: A meta-analysis and systematic review. Front Public Health. 2022;10:874693. [CrossRef]
- 20 Hajmohammadi M, Saki Malehi A, Maraghi E. Effectiveness of using face masks and personal protective equipment to reducing the spread of COVID-19: A systematic review and meta-analysis of case-control studies. Adv Biomed Res. 2023;12:36. [CrossRef]
- 21 Soleman SR, Lyu Z, Okada T, Sassa MH, Fujii Y, Mahmoud MAM, et al. Efficacy of personal protective equipment to prevent environmental infection of COVID-19 among healthcare workers: a systematic review. Environ Health Prev Med. 2023;28:1. [CrossRef]

- **22** Tian C, Lovrics O, Vaisman A, Chin KJ, Tomlinson G, Lee Y, et al. Risk factors and protective measures for healthcare worker infection during highly infectious viral respiratory epidemics: A systematic review and meta-analysis. Infect Control Hosp Epidemiol. 2022;43(5):639-50. [CrossRef]
- 23 Purushothaman PK, Priyangha E, Vaidhyswaran R. Effects of prolonged use of facemask on healthcare workers in tertiary care hospital during COVID-19 pandemic. Indian J Otolaryngol Head Neck Surg. 2021;73(1):59-65. [CrossRef]
- **24** Gyapong F, Debrah E, Oforiwaa M, Isawumi A, Mosi L. Challenges and adverse effects of wearing face masks in the COVID-19 era. Challenges. 2022;13(2):67. [CrossRef]
- 25 Chu DK, Akl EA, Duda S, Solo K, Yaacoub S, Schünemann HJ; COVID-19 Systematic Urgent Review Group Effort (SURGE) study authors. Physical distancing, face masks, and eye protection to prevent person-to-person transmission of SARS-CoV-2 and COVID-19: a systematic review and meta-analysis. Lancet. 2020;395(10242):1973-87. [CrossRef]
- 26 Islam N, Sharp SJ, Chowell G, Shabnam S, Kawachi I, Lacey B, Massaro JM, et al. Physical distancing interventions and incidence of coronavirus disease 2019: natural experiment in 149 countries. BMJ. 2020;370:m2743. [CrossRef]
- 27 Kwon S, Joshi AD, Lo CH, Drew DA, Nguyen LH, Guo CG, et al. Association of social distancing and face mask use with risk of COVID-19. Nat Commun. 2021;12(1):3737. [CrossRef]
- 28 Wellenius GA, Vispute S, Espinosa V, Fabrikant A, Tsai TC, Hennessy J, et al. Impacts of social distancing policies on mobility and COVID-19 case growth in the US. Nat Commun. 2021;12(1):3118. [CrossRef]
- **29** Murugesan M, Venkatesan P, Ramasamy J, Samuel P, Karthik R, Rose W, et al. Transmission dynamics of COVID-19 and utility of contact tracing in risk assessment of health-care worker exposure during COVID-19 pandemic. Indian J Community Med. 2022;47(1):82-6. [CrossRef]