

Heat, heatwaves and cardiorespiratory hospital admissions in Helsinki, Finland.

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Abstract:

Background: There is a lack of knowledge concerning the effects of ambient heat exposure on morbidity in Northern Europe. Therefore, this study aimed to evaluate the relationships of daily summer-time temperature and heatwaves with cardiorespiratory hospital admissions in the Helsinki metropolitan area, Finland.

Methods: Time-series models adjusted for potential confounders such as air pollution were used to investigate the associations of daily temperature and heatwaves with cause-specific cardiorespiratory hospital admissions, during summer months of 2001-2017. Daily number of hospitalizations was obtained from the national hospital discharge register, weather information from the Finnish Meteorological Institute.

Results: Increased daily temperature was associated with decreased risk of total respiratory hospital admissions and asthma. Heatwave days were associated with 20.5% (95% CI: 6.9, 35.9) increased risk of pneumonia admissions and during long or intense heatwaves also with total respiratory admissions in the oldest age group (≥ 75 years). There were also suggestive positive associations between heatwave days and admissions due to myocardial infarction and cerebrovascular diseases. In contrast, risk of arrhythmia admissions was decreased 20.8% (95% CI: 8.0, 31.8) during heatwaves.

Conclusions: Heatwaves, rather than single hot days, are a health threat affecting the morbidity even in a Northern climate.

Keywords: Heat; Heatwave; Cardiovascular diseases; Respiratory diseases; Hospital admissions; Climate change; ambient temperature; Public health; time series; summer months

Introduction

Global warming is anticipated to increase the frequency and intensity of hot days and heatwaves (1). Exposure to high temperature can lead to many types of adverse physiological changes in the human body (2, 3). Therefore, evaluating the association of heat exposure with human health is of paramount importance in a changing climate.

The positive association between ambient heat exposure and mortality has been well established in countries around the world (4, 5, 6, 7, 8). Results from mortality studies have been the basis for the development of heat-health warning systems and action plans (9). However, there have been far less research on the associations between heat exposure and hospital admissions. This is because the hospital data is typically less easily accessible than mortality data. The majority of the studies on hospital admissions have included either all-cause admissions or the broad diagnosis categories of cardiovascular and respiratory diseases.

Interestingly, studies between heat and hospital admissions have exhibited mixed results (10,11). In addition to hospital admissions due to health conditions directly related to heat stress, fluid or electrolyte imbalance (12, 13, 14), respiratory admissions are commonly found to be associated with heatwaves (15,16,17). Associations have also been detected with hospital admissions due to some less studied disease outcomes (18, 19, 20, 21). Cardiovascular diseases have shown a weak association with hot days in some studies (11, 22), but no such association was found in a meta-analysis by Turner et al (23). Hospital admission studies have focused on extreme heat leaving the need to investigate also the effects of moderate heat (10).

Polar amplification in the northern latitudes is making Northern Europe especially susceptible to the effects of global warming (24). So far, only few studies have been carried out to evaluate the association of high ambient temperatures with hospital admissions in Northern European countries (16, 25, 26, 27). Therefore, this study investigates the association of heat and heatwaves with cardiorespiratory hospital admissions in the Helsinki metropolitan area, Finland. Understanding how morbidity, and not just mortality, is associated with heat exposure in Northern Europe is crucial for formulating adaptation strategies.

Methodology:

Study design and population

Our time series study evaluated associations between daily mean temperature and, daily number of non-elective hospital admissions in the Helsinki metropolitan area, Finland, in June-August 2001-2017. The Helsinki metropolitan area includes the cities of Helsinki, Vantaa, Espoo and Kauniainen, population size of approximately 1.15 million during the period 2001-2017 (28).

Health and environmental data

Health data was obtained from a national registry containing information on daily hospital admissions. The study contained information on daily non-elective hospital admission in June-August 2001-2017 for all cardiovascular diseases (CVD: I00-I99), all respiratory diseases (J00-J99), myocardial infarction (MI: I21-I22), ischemic heart disease (IHD: 20-I25), cerebrovascular diseases

(I60-I61, I63-I64), arrhythmia (I46.0, I46.9, I47-I49), asthma (J45, J46), chronic obstructive pulmonary disease (COPD: J41, J44), and pneumonia (J12-J15, J16.8, J18) in specific age groups (all-ages, 18-64, 65-74 and 75 plus).

Exposures of interest in the study were daily mean temperature and heatwaves.

Daily weather data was provided by the Finnish Meteorological Institute. Data was collected using a fixed weather station at the Helsinki-Vantaa airport. Air pollution data was obtained from the Helsinki Region Environmental Services Authority, and included information on nitrogen dioxide (NO₂), ozone (O₃), inhalable particulate matter (PM₁₀; aerodynamic diameter ≤ 10 μm) and fine particulate matter (PM_{2.5}; aerodynamic diameter ≤ 2.5 μm). For NO₂, PM₁₀ and PM_{2.5}, Kallio urban background station was used. Daily averages were calculated from 1-hour values and at least 18 hourly values had to be available, otherwise daily average was defined as missing. For ozone (O₃), 8-hour moving averages were calculated using hourly data from Luukki measurement site which is an aerial background station. At least, 6 hourly values had to be available, otherwise 8 hour moving average were defined as missing. Second, maximum daily 8-hour moving averages were calculated from 8-hour moving averages and at least 18 values had to be available, otherwise the maximum 8-hour average was defined as missing. Data on daily pollen counts was provided by the Aerobiology Unit, University of Turku.

Statistical analysis

Multivariate Poisson time series regression models were used to evaluate whether daily number of hospital admissions is associated with daily mean temperature. We used daily mean temperature and heatwaves separately as exposure variables in the models. Previous studies have reported that the effect of high temperature on hospital admissions take place only with a short delay. Therefore, we looked at individual daily lags from lag 0 (exposure during the day of admission) to lag 5 (exposure 5 days prior to the day of admission). We used the average of lags 0 and 1 as the main exposure variable.

The effects of heatwaves (extended heat exposure) were investigated by adding an indicator variable for heatwaves in the model. There is no standard definition for a heatwave, but varying combinations of intensity and duration have been used in earlier studies. In the current study, heatwaves were defined using 90th and 95th percentile cutoff points for mean daily temperature in May-August 2001-2017. For heatwaves that lasted nine days or less, the cutoff temperature had to be exceeded in each consecutive day. For heatwaves that lasted ten days or longer, one day with temperature below the cutoff point after the tenth day was allowed, if the cutoff temperature was then again exceeded for at least two consecutive days. At the 90th percentile cutoff point, heatwaves were defined as periods lasting for four or more days, and analyses were also conducted separately for short heatwaves (4-7 days) and long heatwaves (10 or more days). At the 95th cutoff point, length of heatwave was 3 or more days.

Time trends were modelled with a three-way interaction between year, month and day of the week and an indicator for holidays. Linear terms for air pollutants (NO₂, O₃, PM₁₀ and PM_{2.5}),

relative humidity (RH) and barometric pressure (BP) were introduced as potential confounders (selected a priori) in the model. Both lag 0 and the average of lags 1-3 were used for air pollutants, relative humidity and barometric pressure. Moreover, pollen count (a sum of alder, birch, mug wort and grasses) was taken into account using two indicators. The daily count (lag 0) and the sum of count on three previous days (lags 1-3) were categorized into two categories using 100 grains/m³ as a cutoff.

Poisson regression was applied using the glm function in the stats package in R (R Development Core Team, Vienna, Austria)(29). Effect estimates were reported as percentage change with 95% confidence intervals. We performed analysis separately for all ages, 18-64, 65-74 and 75 plus. The shape of association of continuous exposure and covariates were checked with the gam function in the mgcv package using thin plate regression splines. This technique has advantage of not requiring the limitation of choosing the knot locations.

As a sensitivity analysis, we run separate models again for daily average temperature by removing days with low temperature (1st percentile, 9.4 °C) and days with high temperature (99th percentile, 24.5 °C). We also run the models again without air pollutant variables to check the robustness of our results.

Results

Table 1 and Table 2 show descriptive statistics for the outcome and exposure variables used in this study. The majority (46.3%) of all the hospital admissions occurred among persons of 75 years of age and above. The mean daily temperature during the study period was 15.3°C. The average daily mean humidity was 70.1%. Daily mean concentrations of PM_{2.5}, PM₁₀, O₃ and NO₂ were 7.6, 14.1, 80.3 and 17.8 µg/m³, respectively.

Table 1: Daily number of hospital admissions for cardiorespiratory diseases during summer months (June-August) in the Helsinki metropolitan area, Finland 2001-2017.

Disease	All-ages			Age18-64			Age 65-74			Age 75 +		
	Media n	Min .	Max .	Media n	Min .	Max .	Media n	Min .	Max .	Media n	Min .	Max .
Cardiovascular												
All	24	6	45	6	0	19	5	0	19	13	2	31
MI	3	0	11	1	0	5	0	0	6	1	0	9
IHD	5	0	20	1	0	8	1	0	9	2	0	12
Cerebrovascular	4	0	12	1	0	7	1	0	7	2	0	10
Arrhythmia	4	0	14	1	0	5	1	0	7	2	0	10
Respiratory												
All	16	2	37	5	0	20	3	0	12	3	0	12
Asthma	1	0	8	0	0	4	0	0	3	0	0	3
COPD	2	0	9	0	0	4	1	0	6	1	0	7
Pneumonia	5	0	23	1	0	11	1	0	7	2	0	15

Table 2: Daily levels of temperature, relative humidity, barometric pressure and air pollutants during summer months (June-August) in the Helsinki metropolitan area, Finland, 2001-2017.

	Mean	Median	Min.	Max.	SD.	Missing values
Meteorology						
Temperature [°C]	16.8	16.7	6.1	26.6	3.4	0
Relative Humidity [%]	72	73	37	98	12.4	0
Barometric Pressure [hPa]	1012	1012	986	1034	7.1	0
Air pollution [$\mu\text{g}/\text{m}^3$]						
Ozone	76.8	75.9	19.4	156.0	17.6	114
PM _{2.5}	7.6	6.6	0.5	50.2	4.6	32
PM ₁₀	13.2	11.9	1.6	62.9	6.5	24
NO ₂	17.0	16.0	2.9	62.6	7.5	23

Table 3 shows the number of heatwave days and mean daily temperature during heatwave and non-heatwave days.

Table 3: Number of heatwave days and mean temperature during heatwave and non-heatwave days during summer months (June-August) in the Helsinki metropolitan area, Finland 2001-2017

	Number of heatwave days	Average temperature during heatwaves [°C]	Average temperature during control days [°C]
90th percentile			
All heatwaves	113	22.8	16.3
Short heatwaves	62	22.6	16.3
Long heatwaves	51	23.1	16.3
95th percentile			
All Heatwaves	60	23.9	16.6

Table 4 shows the associations of daily temperature with hospital admissions due to cardio-respiratory diseases. No statistically significant increased risk was found for hospital admissions in association with daily mean temperature for any of the disease categories. There was a suggestive positive association for arrhythmia admissions in the 65-74 age group. Respiratory diseases

showed a statistically significant protective association in the all-ages group while asthma showed significant protective association for all-ages and 65-74 years age group. All cardiovascular diseases and COPD showed borderline significant protective associations in the 18-64 and 75 plus years age groups, respectively.

Table 4: Percentage change (95% CI) in daily hospital admissions for cardiorespiratory diseases associated with 1°C increase in daily mean temperature during summer months (June-August), 2000-2017, Helsinki, Finland.

Disease	All-ages	Age 18-64	Age 65-74	Age 75 +
Cardiovascular				
All	-0.49 (-1.14, 0.17)	-1.21 (-2.47, 0.07)	0.02 (-1.38, 1.44)	-0.32 (-1.22, 0.60)
MI	0.14 (-1.70, 2.02)	-0.17 (-3.50, 3.28)	0.36 (-3.52, 4.39)	0.16 (-2.51, 2.90)
IHD	-0.46 (-1.79, 0.88)	-0.73 (-3.22, 1.82)	-0.40 (-3.14, 2.42)	-0.37 (-2.26, 1.57)
Cerebrovascular	-0.70 (-2.22, 0.85)	-0.89 (-3.78, 2.09)	-1.37 (-4.44, 1.81)	-0.19 (-2.38, 2.06)
Arrhythmia	-0.66 (-2.24, 0.95)	-2.44 (-5.58, 0.80)	2.95 (-0.48, 6.50)	-1.52 (-3.69, 0.69)
Respiratory				
All	-1.09 (-1.88, -0.30)	0.53 (-0.90, 1.97)	-0.59 (-2.43, 1.28)	-0.11 (-1.45, 1.25)
Asthma	-3.49 (-6.25, -0.65)	3.25 (-2.11, 8.91)	-8.96 (-16.42, -0.82)	-2.68 (-8.06, 3.01)
COPD	-0.53 (-2.56, 1.55)	1.54 (-2.81, 6.09)	0.97 (-2.60, 4.67)	-2.59 (-5.57, 0.49)
Pneumonia	-0.30 (-1.67, 1.09)	-0.66 (-3.17, 1.92)	-0.33 (-3.47, 2.92)	1.12 (-0.94, 3.22)

Associations between heatwaves and cardiorespiratory hospital admissions varied depending on the type of disease (Table 5). In the analysis where 90th percentile of the temperature distribution was used as a cutoff point, statistically significant positive associations were detected between all heatwaves and pneumonia in all-ages and 18-64 age group and also all respiratory diseases in 18-64 age group. We also found suggestive increased risk for cerebrovascular disease in all-ages and COPD hospitalizations in 18-64 age group. Protective effects were detected for arrhythmia. When short heatwaves were analyzed separately, we found significant positive associations also with COPD admissions in the 18-64 year age group and MI among persons 65-74 years of age.

Long heatwaves (90th percentile cutoff point) and more intense heatwaves (95th percentile) showed significant increased risk for all respiratory hospitalizations in elderly people (75 years of age and above). For hospital admissions due to pneumonia, there was a statistically significant positive association with long heatwaves and a suggestive positive association with intense heatwaves in those aged ≥ 75 .

Sensitivity Analyses

We evaluated the robustness of our results by performing sensitivity analyses. Results remained essentially the same (the results not shown).

Table 5: Percentage change (95% CI) in daily hospital admissions for cardiorespiratory diseases associated with heatwave days in summer months (June-August), 2000-2017, Helsinki, Finland

Heatwaves		All-ages	Age 18-64	Age 65-74	Age 75 +
90 th Percentile cut-off, All heatwaves	Cardiovascular				
	All	-2.14 (-7.71,3.76)	-7.57(-17.73,3.84)	2.01(-10.00,15.62)	-0.57(-8.30,7.81)
	MI	4.77 (-11.03,23.38)	-6.82(-31.64,27.00)	40.42(-0.43,98.04)	-3.55(-23.81,22.09)
	IHD	0.06 (-11.24,12.79)	-3.50(-23.35,21.49)	9.62(-14.50,40.55)	-2.81(-18.03,15.23)
	Cerebrovascular	12.17(-1.93,28.29)	12.12(-14.10,46.35)	17.96(-10.56,55.57)	10.07(-8.85,32.93)
	Arrhythmia	-20.78(-31.81,-7.97)	-34.41(-52.71,-9.03)	-13.13(-35.68,17.31)	-18.44(-33.63,0.24)
	Respiratory				
	All	4.18 (-3.04,11.94)	17.97(3.97,33.86)	-6.11(-20.27,10.57)	10.18(-2.05,23.94)
	Asthma	-4.36 (-27.46,26.08)	17.86(-28.57,94.46)	-33.27(-68.73,42.42)	38.56(-15.15,126.25)
	COPD	-5.03 (-20.95,14.10)	48.05(0.00,119.20)	-9.48(-33.48,23.18)	-20.67(-40.38,5.55)
Pneumonia	20.48(6.85,35.85)	28.94(3.00,61.42)	25.67(-4.24,64.93)	18.25(-0.82,40.98)	
90 th Percentile cutoff, Short heatwaves	Cardiovascular				
	All	-1.32(-8.05,5.91)	-3.64(-16.10,10.68)	3.48(-11.15,20.53)	-1.76(-10.92,8.35)
	MI	17.13(-3.44,42.07)	5.21(-28.12,54.01)	74.86(15.75,164.16)	2.36(-21.99,34.31)
	IHD	8.47(-5.70,24.78)	2.63(-22.04,35.11)	21.29(-9.64,62.81)	5.47(-13.30,28.31)
	Cerebrovascular	12.27(-4.19,31.56)	19.07(-12.40,61.84)	4.48(-25.99,47.49)	12.23(-9.94,39.87)
	Arrhythmia	-26.85(-39.36,-11.76)	-33.82(-55.37,-1.87)	-10.75(-38.04,28.56)	-32.07(-48.07,-11.15)
	Respiratory				
	All	5.24(-3.58,14.86)	27.03(9.49,47.38)	-3.89(-21.44,17.59)	-1.04(-14.72,14.85)
	Asthma	21.58(-10.78,65.68)	47.11(-18,23,164.64)	0.23(-57.34,135.48)	72.41(-1.01,200.29)
	COPD	-6.98(-25.78,16.58)	65.33(3.13,165.06)	-10.06(-38.48,31.50)	-28.29(-49.87,2.59)
Pneumonia	20.31(3.49,39.87)	48.30(13.87,93.15)	33.99(-4.80,88.59)	-0.30(-21.09,25.96)	
90 th Percentile cutoff, Long heatwaves	Cardiovascular				
	All	-3.01(-11.46,6.24)	-13.56(-28.14,3.97)	-1.46(-18.64,19.35)	2.50(-9.60,16.23)
	MI	-12.85(-32.96,13.31)	-18.09(-49.11,31.86)	-6.02(-44.48,59.11)	-12.16(-40.86,30.47)
	IHD	-13.36(-28.82,5.47)	-11.50(-38.68,27.73)	-6.97(-37.40,38.24)	-17.85(-38.41,9.58)
	Cerebrovascular	7.87(-13.15,33.98)	-12.16(-43.75, 37.16)	38.92(-7.90,109.53)	4.53(-23.67,43.14)
	Arrhythmia	-8.34(-26.27,14.57)	-29.16(-57-62,18.40)	-24.17(-52,31,20.57)	10.09(-18.10,47.98)
	Respiratory				
All	-1.01(-11.29,10.46)	-0.85(-18.99,21.35)	-12.54(-31.42,11.54)	22.47(3.14,45.43)	
Asthma	-52.46(-71.80, .19.74)	-22.69(-67.96, 86.48)	-76.19(-93.91,-6.86)	-25.76(-67.86,71.52)	

	COPD Pneumonia	-1.63(-25.35, 29.63) 18.20(-0.94,41.03)	35.11(-26.14,147.14) -0.82(-31.20,42.97)	-7.93(-42.00,46.17) 11.85(-24.42,65.53)	-10.78(-41.60,36.30) 40.65(9.87,80.05)
95 th percentile cut-off, All heatwaves	Cardiovascular All MI IHD Cerebrovascular Arrhythmia Respiratory All Asthma COPD Pneumonia	-2.43(-9.60,5.31) 3.02(-17.04,27.93) 1.83(-13.03,19.24) -3.86(-19.65,15.05) -14.67(-29.62,3.45) 1.77(-7.27,11.71) 11.30(-24.55,64.21) 4.93(-16.87,32.44) 6.80(-8,66,24.88)	-12.15(-24.15,2.23) -13.29(-43.04,32.00) -14.20(-37.31,17.42) -34.11(-54.61,-4.36) -13.72(-41.67,27.61) -0.37(-15.56,17.57) 27.09(-38.56,162.86) 42.26(-15.37,139.15) -5.95(-30.44,27.17)	-1.07(-15.82,16.25) 9.38 (-29.32,69.27) 3.11(-24.99,41.72) 13.26(-20.43,61.20) -18.75(-45.62,21.41) -18.93(-34.51,0.36) -28.62(-72.94,88.25) -25.69(-50.41,11.37) 3.51(-27.49,47.76)	2.22(-8.07,13.67) 8.03(-21.07,47.85) 9.44(-12.54,36.95) 6.23(-17.64,37.00) -13.82(-34.07,12.66) 22.46(5.62,41.98) 68.89(-8.60,212.10) 18.38(-16.41,67.63) 24.16(-0.64,55.15)

Discussion:

This study investigated the associations of daily mean temperature and heatwaves with hospital admissions for cardiorespiratory diseases in summer months (June-August) in the Helsinki metropolitan area, Finland. We found that daily mean temperature was associated with decreased risk of hospitalization for respiratory diseases, asthma, and to lesser extent cardiovascular diseases. In contrast, during heatwaves the risk of hospitalization was increased due to pneumonia, and in some age groups for respiratory diseases in general, COPD and MI. We also found a negative association of heatwaves with arrhythmia hospitalizations.

Our study results showed no increased risk for total cardiovascular admissions in association with daily mean temperature and heatwaves. These findings are in line with a review; in which Turner et al. found no association between summer temperature and cardiovascular morbidity (23). Likewise, previous literature has shown either no or weak association between heat and cardiovascular morbidity (30, 31, 32, 33). One of the potential reasons could be that most of the vulnerable people with acute cardiovascular events might die before reaching to hospital which can explain the low cardiovascular hospital admissions (34).

Our results suggest that heatwaves may yet be associated with specific subtypes of cardiovascular diseases. We found borderline significant association between heatwaves and cerebrovascular admissions in all-ages group. However, there has been no evidence in previous literature for association of heat and heatwaves with cerebrovascular admissions. An overview of the systematic reviews found no association of cerebrovascular disease hospitalizations with heat (35). Our findings also indicated positive association of heatwaves with myocardial infarction admissions in 65-74 year old persons. However, previous studies have found no or a decreased risk of MI admissions during summer months (26, 36, 37).

We also observed a protective association of arrhythmia admissions with heatwaves. A population based study from Canada did not find any association between heat and arrhythmia admissions (38). There are too few studies to draw conclusions about the association between heat and arrhythmia hospital admissions.

We found a positive association of heatwaves with hospital admissions due to respiratory diseases in general, pneumonia and COPD. These results are consistent with the findings from previous studies on respiratory morbidity (15, 39, 40, 41). The possible explanation for a stronger association between heatwaves and respiratory admissions than between heatwaves and cardiovascular diseases is that respiratory diseases are not as acutely fatal as cardiovascular diseases. This means that more people are able to reach hospitals (42). In extreme heat, human body maintains safe body temperature through thermoregulation which is associated with increase in skin blood flow, cardiac output and pulmonary ventilation. Though body heat is lost by sweating, hyperthermia can cause hyperventilation leading to thermal hyperpnea (increase in respiratory rate and tidal volume). These responses from thermoregulatory mechanism can lead to an increase in risk for respiratory and COPD hospitalizations (43, 44). Andreson et al. (2013)

argued that, in addition to the thermoregulatory response, acute respiratory effects may also be caused by the direct effect of breathing hot air (45).

In our study long heatwaves and more intense heatwaves (95th percentile cutoff point) were associated with risk of total respiratory and pneumonia hospitalizations only in the oldest age category. These findings are in line with other studies which reported elderly people as the most vulnerable group for respiratory morbidity effects in extreme and long heatwaves (46, 47, 48). However, with different definitions of heatwaves we also found associations between heatwaves and respiratory health in the age group 18-64.

The risk of morbidity association with heatwaves may vary with length and intensity. A study by Levy et al. showed that length of heatwaves increase the risk of emergency department arrivals (49). Likewise, a study in Australia also found increase in infant's hospitalizations with duration of heatwave (50). The effect of heatwaves on morbidity may vary across different regions and still needs more investigation.

Finland, being a North European country, has cold summer temperatures. This is most likely the reason that we found no positive associations or even some protective associations in connection with daily temperature. However, our results further suggested that heatwaves pose a public health concern regarding morbidity (as indicated by hospital admissions).

One of the strengths of our study is that it was conducted in a Northern climate where only few studies have taken place. We used long time series data and had in practice a hundred percent coverage of hospital admissions. Thus reliable evidence on population level associations could be provided. Also, this study evaluated the associations of temperature with cause-specific diagnosis and in age groups rather than for broader disease categories and total population. The potential confounding effect of air pollution was controlled using all major indicators of air pollution. However, this study has some limitations as well. We were not able to include persons younger than 18 years because the limited number of cases in the age group would have led to imprecise effect estimates. Also, the study only included cardiovascular and respiratory hospital admissions. Admissions due to other causes could also be affected by heat exposure.

Conclusion:

In conclusion, we found no associations and even protective associations between daily mean temperature and cardiorespiratory hospital admissions. However, our results suggest that heatwaves are a health threat and affect morbidity even in the northern climate. Moreover, not only the elderly are at risk but preventive programs should also take other age groups into account.

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Author contributions:

Conceptualization, Hasan Bin Sohail, Virpi Kollanus and Timo Lanki; Data curation, Hasan Bin Sohail, Pekka Tiittanen, Virpi Kollanus and Timo Lanki; Formal analysis, Hasan Bin Sohail, Pekka Tiittanen and Timo Lanki; Funding acquisition, Timo Lanki; Investigation, Hasan Bin Sohail, Pekka Tiittanen, Virpi Kollanus and Timo Lanki; Methodology, Hasan Bin Sohail, Pekka Tiittanen and Timo Lanki; Project administration, Timo Lanki; Resources, Timo Lanki; Supervision, Alexandra Schneider and Timo Lanki; Validation, Pekka Tiittanen; Writing – original draft, Hasan Bin Sohail; Writing – review & editing, Hasan Bin Sohail, Pekka Tiittanen, Virpi Kollanus, Alexandra Schneider and Timo Lanki.

Conflict of Interest:

The authors declare no conflict of interest.

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