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## Article

# An Update on the Link between COVID-19 Vaccination and Mortality

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**Abstract:** This study updates previous research showing that 22 all-cause mortality in 31 European countries increased over time the higher the 21 COVID-19 full vaccination uptake. The update illuminates that a one percentage point increase in 21 full vaccination uptake initially decreased all-cause mortality from Jan to Mar 22 by  $-0.423$  percent (95% CI  $-0.577, -0.270$ ), but the following 14 months, a one percentage point increase in 21 booster vaccination uptake oppositely increased mortality by  $0.366$  percent (95% CI  $0.250, 0.482$ ). The findings indicate that full vaccination initially prevented mortality, but subsequently, booster vaccination, in particular, detrimentally and consistently induced higher mortality. The effects remained robust when controlling for alternative explanations. Studies have argued that heat waves caused mortality in the 22 summer and energy prices caused mortality in the 22-23 winter. However, the update shows that booster vaccination consistently induced higher mortality when neither heat waves nor energy prices were likely explanations.

**Keywords:** COVID-19; full vaccination; booster vaccination; all-cause mortality; excess mortality

## 1. Introduction

This paper updates previous research by Aarstad and Kvitastein [1], showing that all-cause mortality in 31 European countries during the nine first months of 22 increased over time the higher the COVID-19 full vaccination uptake by the end of 21. The following paragraphs address the update's contributions.

First, it adds eight monthly observations of country-level all-cause mortality, extending the study period from Jan 22 to May 23. I.e., it extends the study period from nine to 17 months.

Second, it includes country-level first booster vaccination uptake by the end of 21 as an additional explanatory variable of mortality in the following months. Booster vaccination in this study implies that three doses have been administered, and the motive for including the measure is Uversky, *et al.* [2] asserting that repeated injections can suppress natural antiviral responses in addition to causing autoimmune diseases, including myocarditis and promote cancer growth. In line with these statements, I do not rule out that booster vaccination may affect mortality, which the update investigates.

Third, the update in more detail than the Aarstad and Kvitastein [1] study assesses how the association between vaccination uptake by the end of 21 and subsequent mortality has changed over time. They showed that the overall association increased from Jan 22 to Sep 22, which does not rule out it may have been negative initially, i.e., vaccination having a temporal preventive mortality effect. Subsequently, the association likely turned positive, which implies that vaccination, in the longer run, has had a detrimental harmful mortality effect. To gain more detailed and explicit knowledge about these issues, the update examines the association between vaccination uptake by the end of 21 and mortality for each of the following 17 months of observation. The approach assesses in which months during 22 and 23 the mortality may have decreased or increased from vaccination uptake by the end of 21. It further contributes to other research showing that the preventive effect of COVID-19 vaccination against hospital admission and death waned [3], but to my knowledge, we do not know when the preventive effect may have gone into negative terrain, causing increased mortality.

Fourth, the update particularly emphasizes the association between COVID-19 vaccination uptake and mortality in Apr and May, 22 and 23, plus Sep and Oct in 22. The motive for emphasizing those six months is a recent study arguing that heat waves increased mortality between Jun and Aug in the summer of 22 [4], and a second study arguing that increased energy prices caused increased mortality in the following winter between Nov 22 and Mar 23 [5]. Analyzing Apr and May in 22 and 23, plus Sep and Oct in 22 – during spring and autumn months neither strongly exposed to heat waves nor adverse effects of increased energy prices – partakes to rule out alternative explanations of mortality as suggested by the two studies.

## 2. Materials and Methods

The 31 countries studied are EU member states, plus Norway, Iceland, Switzerland, and Liechtenstein. The dependent variable is monthly country-level excess all-cause mortality compared to average pre-pandemic all-cause mortality in the same months between 16 and 19. Values higher (lower) than zero indicate positive (negative) excess mortality. E.g., a value of 10 (–10) for a given month in a given country indicates ten percent higher (lower) mortality than the pre-pandemic average, i.e., positive (negative) excess mortality. The data were sourced from the Eurostat mortality database [6]. At the time of data collection, the database included complete mortality data from all 31 countries from Jan 22 to May 23, except for Italy, missing data for May 23. In total, the study includes 526 country-level observations of monthly mortality.

Table 1 reports descriptive statistics concerning the dependent variable, excess mortality, for different country-level months (on which I will report in further detail shortly). All the study's statistics are weighted by the countries' population size at the beginning of 20 (intuitively except for minimum and maximum values), as reported in Table 2, sourced from Eurostat [7]. The blue line in Figure 1 reports weighted average excess mortality between Jan 22 and May 23 for all the 31 countries (except for missing data from Italy in May 23). The blue line shows positive excess mortality throughout the study period, except for Feb 23 when it was slightly negative (shortly, I report on the figure in more detail, including the red and green lines).

As independent explanatory variables, the study includes data on country-level full vaccination and first booster vaccination uptake by the end of 21. Full vaccination implies the percentage of the total population in each country having taken the primary course, i.e., two doses, and booster vaccination implies the percentage of the population having taken a third dose. The data were accessed from the European Centre for Disease Prevention and Control [8], except for Switzerland where they were accessed from Our World in Data [9]. Table 2 reports full vaccination and booster vaccination uptake for each country.

To account for alternative explanations, the study includes models controlling for the country-level average 20-21 all-cause mortality relative to the average 16-19 all-cause mortality [10], the life expectancy in 19 [11], the median age in 18 [12], and the purchasing power adjusted per capita GDP in 19 [13] where I include the same number for Liechtenstein as reported for Switzerland. Aarstad and Kvitastein [1] explain the motive for including the two first control variables, but briefly, the concepts account for mortality and the countries' health conditions prior to examining the study's dependent variable. Concerning the two latter control variables, the median age is an additional proxy for a country's health conditions, as people in countries with old populations, *ceteris paribus*, are more prone to sickness and death than in countries with young populations. GDP is an indicator of countries' financial ability to handle challenges related to health issues.

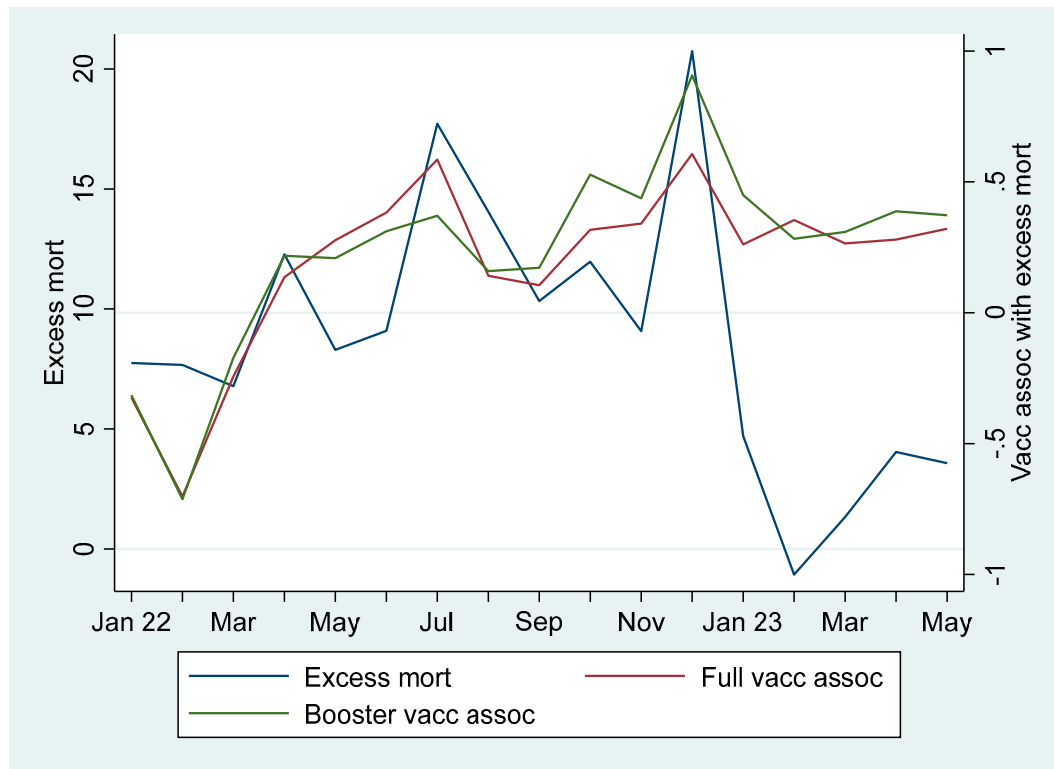
Table 3 reports descriptive statistics concerning the independent- and control variables, and Table 4 reports correlations. Concerning correlations taking high absolute values, potentially causing problems with multicollinearity cf. [14], I shortly address how I deal with the issue.

**Table 1.** Descriptive statistics concerning the dependent variable, excess mortality. Mean values and standard deviations are weighed by countries’ population size.

Months included	Min	Max	Mean	St dev	Monthly obs
Jan22-May23	−27.9	54.4	8.77	8.66	526
Jan22-Mar22	−4.5	54.4	7.40	7.59	93
Apr22-May23	−27.9	38.2	9.06	8.86	433
Apr22, May22, Sep22, Oct22, Apr23, May23	−27.9	36.8	8.52	6.27	185

**Table 2.** Full vaccination uptake, booster vaccination uptake, and countries’ population size. The table is sorted by booster uptake.

Country	Full vacc	Booster vacc	Population
Bulgaria	27.7	4	6951482
Romania	40.8	5.9	19328838
Croatia	53.3	13.2	4058165
Latvia	64.9	13.7	1907675
Poland	55.8	18.4	37958138
Slovakia	48.5	20.2	5457873
Slovenia	55.6	21.5	2095861
Lithuania	65.7	22.6	2794090
Finland	75	23.2	5525292
Estonia	61.3	24.2	1328976
Sweden	69.3	24.3	10327589
Switzerland	67.1	25.2	8606033
Czechia	62.2	27.7	10693939
Netherlands	67.6	27.8	17407585
Norway	73	29	5367580
Spain	74.9	30.2	47332614
Portugal	83.1	30.7	10295909
Liechtenstein	65.4	31.1	38747
Luxembourg	68.2	31.8	626108
Hungary	61	32.4	9769526
Italy	74.9	33.5	59641488
Cyprus	69.4	33.8	888005
Greece	66.3	34.1	10718565
France	73.3	36.1	67320216
Belgium	76.2	37.9	11522440
Germany	72.8	41.4	83166711
Ireland	76.6	43.5	4964440
Malta	82.1	44.5	514564
Austria	70.4	44.7	8901064
Denmark	78.7	46.9	5822763
Iceland	76.4	54.2	364134



**Figure 1.** The blue line shows monthly excess mortality between Jan 22 and May 23. For the same months the red line shows the monthly full vaccination association with excess mortality, and the green line shows the monthly booster vaccination association with excess mortality.

**Table 3.** Descriptive statistics concerning the independent- and control variables. The number of countries included is 31. Mean values and standard deviations are weighed by countries' population size.

Variable	Min	Max	Mean	St dev
Full vacc by the end of 21	27.7	83.1	68.7	10.5
Booster vacc by the end of 21	4	54.2	31.2	9.88
Avg 2020-2021 mort rel to 16-19	101.4	126.1	112.5	5.67
Life expectancy 19	75.1	84.3	81.5	2.51
Median age 18	36.5	47.1	43.3	2.45
Per capita GDP 19	53	254	101.6	24.5

N=31.

**Table 4.** Correlations concerning the independent- and control variables. The number of countries included is 31. Correlations are weighed by countries' population size.

	1	2	3	4	5
Full vacc by the end of 21 (1)	1				
Booster vacc by the end of 21 (2)	0.827***	1			
Avg. 20-21 mort rel to 16-19 (3)	-0.742***	-0.750***	1		
Life expectancy 19 (4)	0.857***	0.629***	-0.620***	1	
Median age 18 (5)	0.316†	0.488**	-0.308†	0.195	1
Per capita GDP 19	0.544**	0.629***	-0.740***	0.515**	0.168

Two-tailed tests of significance. †  $p < 0.10$ ; \*  $p < 0.05$ ; \*\*  $p < 0.01$ ; \*\*\*  $p < 0.001$ . N=31.

### 3. Results

Initially, I carried out explorative analyses, which showed that the independent variables, country-level full- and booster vaccination uptake by the end of 21, respectively, were negatively associated with the dependent variable, excess all-cause mortality, Jan, Feb, and Mar 22. The

following 14 months, i.e., from Apr 22 to May 23, the association turned positive. (Shortly, I will report on and explain these findings in detail.)

Based on the negative associations between the independent and dependent variables from Jan 22 to Mar 22, Models 1-3 (Table 5) conducted multilevel mixed effects random intercept linear regressions for this period. The dependent variable observations of monthly excess mortality were nested at the country level [for further readings, see 15,16]. The regressions were weighted by the countries' population size (cf. Table 2) and report robust standard errors concerning the fixed effects (intercepts and regressors). Model 1 includes full vaccination uptake only as an independent variable. It shows that between Jan 22 and Mar 22 full vaccination uptake was negatively and significantly associated with all-cause mortality. Specifically, a one percentage point increase in country-level full vaccination uptake decreased all-cause mortality by -0.423 percent (95% CI -0.577, -0.270). Model 2 includes booster vaccination uptake only as an independent variable and also shows a significant negative association with the dependent variable, but the effect is weaker than in the previous model. Model 3 includes both the independent variables and indicates that full vaccination uptake had a genuine negative significant association with the dependent variable, while booster vaccination uptake had a non-significant negative association. Having said that, as the correlation between the two independent variables is high, taking a value of 0.827 (cf. Table 4) and reflected in a high variance inflation factor, I do not rule out multicollinearity in Model 3. Therefore, I conclude that both the independent variables had a strong negative association with the dependent variable between Jan 22 and Mar 22, but full vaccination uptake probably had a more dominant and genuine effect on excess mortality than booster vaccination uptake. Thus, full vaccination uptake by the end of 21 seems to have reduced excess mortality the three first months of the following year. In other words, the findings indicate that full vaccination prevented mortality during this period.

Based on the positive associations between the independent and dependent variables from Apr 22 to May 23 in the explorative analyses referred to above (and which I will report on in more detail shortly), Models 4-6 (Table 5) conducted multilevel mixed effects random intercept linear regressions for this period similarly as reported above. Model 4 includes full vaccination uptake only as an independent variable. It shows that between Apr 22 and May 23 full vaccination uptake was positively and significantly associated with all-cause mortality. Model 5 includes booster vaccination uptake only as an independent variable and also shows a significant positive association with the dependent variable, and the effect is stronger than in the previous model. Specifically, a one percentage point increase in country-level booster vaccination uptake increased country-level all-cause mortality by 0.366 percent (95% CI 0.250, 0.482). Model 6 includes both the independent variables and indicates that booster vaccination uptake had a genuine positive significant association with the dependent variable, while full vaccination uptake had a non-significant positive association. Again, multicollinearity may be an issue when including both independent variables in the same model (cf. my discussion above). Therefore, I conclude that both the independent variables had a strong positive association with the dependent variable between Apr 22 and May 23, but booster vaccination uptake probably had a more dominant and genuine effect on excess mortality than full vaccination uptake. Thus, booster vaccination uptake by the end of 21 seems to have increased excess mortality from Apr 22 to May 23. In other words, the findings indicate that booster vaccination had a detrimental effect leading to higher mortality during this period.

**Table 5.** Multilevel mixed-effects random intercept linear regressions with robust standard errors and monthly all-cause mortality as the dependent variable.

	Model 1	Model 2	Model 3	Model 4	Model 5	Model 6
Months included in the analysis	Jan22-Mar22	Jan22-Mar22	Jan22-Mar22	Apr22-May23	Apr22-May23	Apr22-May23
FIXED EFFECTS						
Intercept	36.5*** (5.36)	19.9*** (2.95)	34.4*** (6.30)	-12.5*** (3.94)	-2.37 (1.89)	-6.07 (3.72)
Full vacc by the end of 21	-0.423*** (0.078)		-0.351* (0.138)	0.314*** (0.061)		0.090 (0.090)



	Model 1	Model 2	Model 3	Model 4	Model 5	Model 6
Booster vacc by the end of 21		-0.400*** (0.085)	-0.092 (0.161)		0.366*** (0.059)	0.288* (0.113)
<b>RANDOM EFFECTS</b>						
Residual	29.8 (11.3)	29.8 (11.3)	29.8 (11.3)	61.9 (6.22)	61.9 (6.21)	61.9 (6.21)
Country effect	8.30 (5.76)	12.2 (7.81)	8.05 (5.89)	5.76 (2.34)	3.57 (2.64)	3.29 (2.31)
Wald $\chi^2$	29.3***	22.0***	28.5***	26.2***	38.6***	44.4***
Log pseudo-likelihood	-4.46e9	-4.50e9	-4.45e9	-2.25e10	-2.24e10	-2.24e10
Monthly observations	93	93	93	433	433	433
Variance inflation factor			3.16			3.17

Estimates are weighted by countries' population size, and I report robust standard errors in parentheses concerning the fixed effects. For fixed effects, I also report conservative two-tailed tests of significance. †  $p < 0.10$ ; \*  $p < 0.05$ ; \*\*  $p < 0.01$ ; \*\*\*  $p < 0.001$ .

To account for alternative explanations concerning the negative association between full vaccination uptake and excess mortality between Jan 22 and Mar 22, Table 6 includes the control variables without altering any statistical conclusion. Similarly, Table 7 includes the control variables to account for eventual alternative explanations concerning the positive association between booster vaccination uptake and excess mortality between Apr 22 and May 23, but without altering any statistical conclusion.

**Table 6.** Multilevel mixed-effects random intercept linear regressions with robust standard errors and monthly all-cause mortality as the dependent variable. Months included are Jan 22 to Mar 22.

	Model 1	Model 2	Model 3	Model 4
<b>FIXED EFFECTS</b>				
Intercept	-3.05 (26.6)	-13.7 (40.0)	50.0** (15.9)	37.7*** (5.02)
Full vacc by the end of 2021	-0.310** (0.098)	-0.576*** (0.138)	-0.397*** (0.079)	-0.316*** (0.084)
Avg. 2020-2021 mort rel to 2016-2019	0.282 (0.187)			
Life expectancy 2019		0.745 (0.585)		
Median age 2018			-0.356 (0.380)	
Per capita GDP 2019				-0.085* (0.037)
<b>RANDOM EFFECTS</b>				
Residual	29.8 (11.3)	29.8 (11.3)	29.8 (11.3)	29.8 (11.3)
Country effect	7.19 (6.20)	7.40 (5.46)	7.64 (6.11)	5.39 (4.88)
Wald $\chi^2$	38.9***	35.2***	29.5***	38.9***
Log pseudo-likelihood	-4.44e9	-4.44e9	-4.459	-4.42e9
Variance inflation factor	2.23	3.76	1.11	1.42

Estimates are weighted by country size in population by Jan 1, 20, and I report robust standard errors in parentheses concerning the fixed effects. For fixed effects, I also report conservative two-tailed tests of significance. †  $p < 0.10$ ; \*  $p < 0.05$ ; \*\*  $p < 0.01$ ; \*\*\*  $p < 0.001$ . The number of monthly observations is 93.

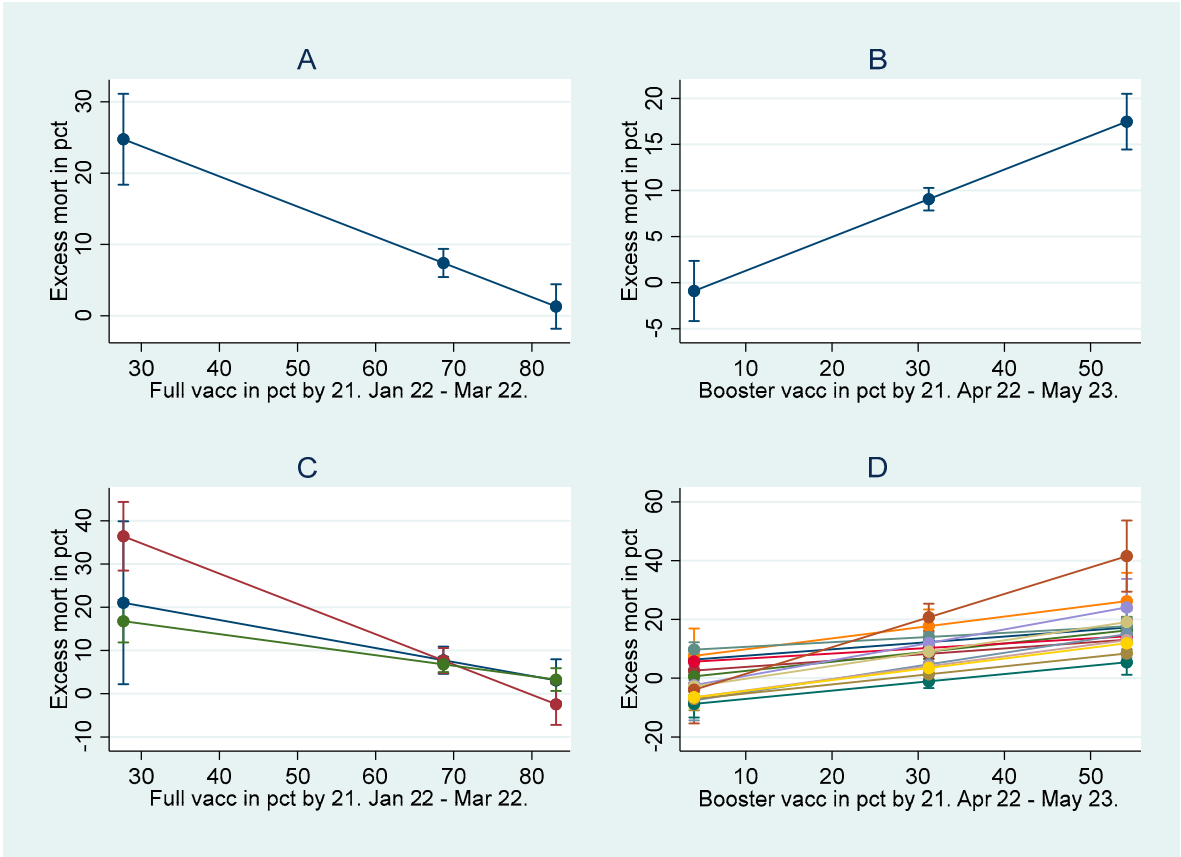
**Table 7.** Multilevel mixed-effects random intercept linear regressions with robust standard errors and monthly all-cause mortality as the dependent variable. Weighted by countries' population size. Months included are Apr 22 to May 23.

	Model 1	Model 2	Model 3	Model 4
<b>FIXED EFFECTS</b>				
Intercept	7.10 (19.5)	-28.6 (25.6)	-12.7 (8.83)	-4.63* (2.28)
Booster vacc by the end of 21	<b>0.334***</b> (0.092)	<b>0.311**</b> (0.094)	<b>0.334***</b> (0.063)	<b>0.300***</b> (0.078)
Avg. 2020-2021 mort rel to 16-19	-0.075 (0.152)			
Life expectancy 19		0.343 (0.339)		
Median age 18			0.262 (0.220)	
Per capita GDP 19				0.043 (0.027)
<b>RANDOM EFFECTS</b>				
Residual	61.9 (6.21)	61.9 (6.22)	61.9 (6.22)	61.9 (6.21)
Country effect	3.49 (2.58)	3.12 (2.04)	3.25 (2.68)	2.94 (2.27)
Wald $\chi^2$	36.1***	39.4***	45.6***	41.9***
Log pseudo-likelihood	-2.24e10	-2.24e10	-2.24e10	-2.24e10
Variance inflation factor	2.30	1.66	1.31	1.66

Estimates are weighted by countries' population size by Jan 1, 20, and I report robust standard errors in parentheses concerning the fixed effects. For fixed effects, I also report conservative two-tailed tests of significance. †  $p < 0.10$ ; \*  $p < 0.05$ ; \*\*  $p < 0.01$ ; \*\*\*  $p < 0.001$ . The number of monthly observations is 433.

Graph A in Figure 2, based on Model 1 in Table 5, shows how the country-level mortality decreased between Jan 22 and Mar 22 as a function of country-level full vaccination uptake. Similarly, Graph B, based on Model 5 in Table 5, shows how the country-level mortality between Apr 22 and May 23 increased as a function of country-level booster vaccination uptake. The graphs report 95% confidential intervals (CIs) for minimum, weighted mean, and maximum vaccination values. Graphs C and D are based on regressions that add interaction terms between month dummies and vaccination uptake. Consistent with Graph A, Graph C shows how the mortality decreased as a function of full vaccination uptake, but in addition, it illustrates the association for each month between Jan 22 and Mar 22. Similarly, and consistent with Graph B, Graph D shows how the mortality increased as a function of booster uptake, but in addition, it illustrates the association for each month between Apr 22 and May 23.





**Figure 2.** Graph A shows country-level variation in excess mortality between Jan 22 and Mar 22 as a function of country-level full vaccination by the end of 21 (based on Model 1, Table 5). Graph B shows country-level variation excess mortality between Apr 22 and May 23 as a function of country-level booster vaccination by the end of 21 (based on Model 5, Table 5). Graphs C and D show monthly country-level variations in excess mortality. Minimum, weighted mean, and maximum values are reported with 95% confidential intervals.

Returning to Figure 1, its red line illustrates the association between full vaccination uptake by the end of 21 and mortality the following 17 months, and its green line similarly illustrates the association between booster vaccination uptake and mortality. The lines show that the associations from Jan 22 to Mar 22 were negative, which aligns with the results I have reported above. However, from Apr 22 the associations turned positive and remained so for the rest of the study period till May 23, which also aligns with the results I have reported above. In addition, the red and green lines show that the associations had increasing trends at least the first half of 22, which aligns with Aarstad and Kvitastein [1] also showing that the association between vaccination uptake and excess mortality increased over time. Roughly between May 22 and Jul 22 the positive association between full vaccination uptake and mortality was stronger than between booster vaccination and mortality, but from Sep 22 the positive association between booster vaccination and mortality was stronger than between full vaccination and mortality, except for Feb 23. The results indicate that booster vaccination over time appears to have a stronger effect on mortality than full vaccination uptake, and they align with findings reported above. Finally, it is worth mentioning that particularly from Apr 22, a change in the *association* between booster vaccination and mortality was strongly correlated with a change in mortality (correlation coefficient is 0.851), which is also possible to observe from the blue and green lines in Figure 1. It indicates that months with increasing mortality mainly occurred in countries with high booster vaccination uptakes.

Table 8 reports statistical details concerning the associations between full- and booster vaccination uptake by the end of 21, respectively, and mortality in the following 17 months. It shows that the monthly associations were statistically significant, i.e.,  $p < 0.05$ , except for a few months where

they either were non-significant (Jan 22 for both vaccination types and Sep 22 for full vaccination) or borderline-significant (Sep 22 for booster vaccination and Jan 23 for full vaccination), i.e.,  $p < 0.10$ .

**Table 8.** The monthly associations between 21 full- and booster vaccination uptake, respectively, and excess mortality. .

Year	Month	Full vacc assoc with mort	Booster vacc assoc with mort
22	Jan	-0.324 (0.214) [-0.761, 0.113]	-0.315 (0.220) [-0.764, 0.135]
	Feb	-0.702*** (0.106) [-0.919, -0.484]	-0.712*** (0.127) [-0.973, -0.452]
	Mar	-0.244*** (0.061) [-0.369, -0.119]	-0.172* (0.069) [-0.314, -0.031]
	Apr	0.136* (0.058) [0.018, 0.254]	0.218** (0.061) [0.094, 0.341]
	May	0.277*** (0.057) [0.159, 0.394]	0.209** (0.067) [0.071, 0.346]
	Jun	0.383*** (0.091) [0.196, 0.569]	0.311** (0.101) [0.104, 0.519]
	Jul	0.585** (0.153) [0.273, 0.897]	0.371* (0.158) [0.047, 0.694]
	Aug	0.141* (0.051) [0.036, 0.246]	0.159** (0.047) [0.063, 0.254]
	Sep	0.105 (0.075) [-0.048, 0.259]	0.172† (0.100) [-0.033, 0.376]
	Oct	0.317* (0.141) [0.030, 0.604]	0.527** (0.160) [0.200, 0.855]
	Nov	0.341** (0.091) [0.154, 0.527]	0.437*** (0.081) [0.272, 0.602]
	Dec	0.607* (0.247) [0.101, 1.11]	0.907*** (0.226) [0.444, 1.369]
23	Jan	0.145† (0.145) [-0.035, 0.557]	0.450** (0.138) [0.168, 0.731]
	Feb	0.354*** (0.065) [0.220, 0.488]	0.282** (0.079) [0.120, 0.445]
	Mar	0.265** (0.074) [0.114, 0.416]	0.309*** (0.059) [0.188, 0.430]
	Apr	0.280*** (0.069) [0.138, 0.422]	0.388*** (0.056) [0.273, 0.502]
	May	0.321*** (0.073) [0.172, 0.470]	0.372*** (0.058) [0.254, 0.491]

The estimates are weighted by the countries' population size. I report robust standard errors in parentheses and 95% CIs in brackets. Also, I report conservative two-tailed tests of significance. †  $p < 0.10$ ; \*  $p < 0.05$ ; \*\*  $p < 0.01$ ; \*\*\*  $p < 0.001$ . N=31, except for May 23 when N=30 (due to missing data from Italy that month). .

Figure 2 shows that the associations between booster vaccination uptake and mortality were positive in Apr, May 22, and 23, and Sep and Oct in 22. Table 8 additionally reveals that the positive associations were statistically significant in all those months except for Sep 22, which was borderline significant. Multilevel mixed effects random intercept linear regression moreover shows an overall positive significant association between booster vaccination uptake and excess mortality when observations for these six months are included as the dependent variable (Table 9). I conclude from the results that booster vaccination uptake consistently induced higher mortality in months when, as suggested by other studies [4,5], neither heat waves nor energy prices were likely explanations.

**Table 9.** Multilevel mixed-effects random intercept linear regressions with robust standard errors and monthly all-cause mortality as the dependent variable. Months included are Apr and May in 22 and 23, plus Sep and Oct in 22.

FIXED EFFECTS	
Intercept	-1.35 (1.96)
Booster vacc by the end of 21	<b>0.316***</b> (0.061)
RANDOM EFFECTS	
Residual	25.6 (2.93)
Country effect	3.84 (2.52)
Wald $\chi^2$	26.6***
Log pseudo-likelihood	-8.39e9

Estimates are weighted the countries' population size by Jan 1, 20, and I report robust standard errors in parentheses concerning the fixed effects. For fixed effects, I also report conservative two-tailed tests of significance. †  $p < 0.10$ ; \*  $p < 0.05$ ; \*\*  $p < 0.01$ ; \*\*\*  $p < 0.001$ . The number of monthly observations is 185.

#### 4. Discussion

This study updates previous research by Aarstad and Kvitastein [1] showing that 22 all-cause mortality in 31 European countries increased over time the higher the 21 COVID-19 full vaccination uptake. The update illuminates that a one percentage point increase in 21 full vaccination uptake initially decreased all-cause mortality from Jan 22 to Mar 22 by  $-0.423$  percent (95% CI  $-0.577, -0.270$ ), but the following 14 months, a one percentage point increase in 21 booster vaccination uptake oppositely increased mortality by  $0.366$  percent (95% CI  $0.250, 0.482$ ). The findings indicate that full vaccination initially prevented mortality, but subsequently, booster vaccination, in particular, detrimentally and consistently induced higher mortality. Studies have argued that heat waves caused mortality in the 22 summer [4] and energy prices caused mortality in the 22-23 winter [5]. However, the update shows that booster vaccination uptake consistently induced higher mortality in months when neither heat waves nor energy prices were likely explanations.

Fortunately, the excess mortality has been reduced from its peak on Dec 22. However, as the weighted average excess mortality in the 31 countries was positive in 20, 21 [1], and 22, one should expect that it would be markedly negative at the beginning of 23, but this is not the case. Instead, the country-level mortality is higher the higher the country-level booster vaccination uptake in particular, and the association remains positive even when the time asymmetry between the determination of the independent and dependent variable has increased. In other words, the more months between the determination of the independent and the dependent variable there is a significant association, the more likely a genuine causal relationship. Moreover, the initial negative association between full vaccination uptake and excess mortality Jan 22 - Mar 22, and later the extended positive association between booster vaccination uptake and excess mortality Apr 22 - May 23, remained statistically significant when controlling for alternative explanations [for a discussion about how time asymmetry and accounting for alternative explanations partake to assess causality, please see Stuart Mill cited in Cook and Campbell, 17, pp. 18-20]. Adding to this, Aarstad and Kvitastein [1] discussed how their study addressed the issue of ecological fallacy [18], e.g., the classical Robinson's paradox [19,20] and the Simpson's paradox [21-24], which also applies to this updated version.

However, despite addressing the issues of causality and ecological fallacy, a limitation of my paper is nonetheless that it researches country-level data only. Therefore, I encourage future research to also investigate person-level data by, for instance, emphasizing different types of vaccination, the timing, and the number of doses given to males and females in different age cohorts as potential carriers of long-term mortality. Similarly, I suggest using person-level chronic morbidity as a dependent variable. Person-level data, too, can have limitations, which the literature has labeled individualistic fallacy [20,24], but additional research addressing COVID-19 vaccination uptakes as potential causes of excess mortality and chronic morbidity – important research questions, in my opinion – at different levels of analysis using different methodologies is warranted, I argue.

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#### References

1. Aarstad, J.; Kvitastein, O.A. Is there a Link between the 2021 COVID-19 Vaccination Uptake in Europe and 2022 Excess All-Cause Mortality? *Preprints* **2023**, doi:10.20944/preprints202302.0350.v1.
2. Uversky, V.N.; Redwan, E.M.; Makis, W.; Rubio-Casillas, A. IgG4 Antibodies Induced by Repeated Vaccination May Generate Immune Tolerance to the SARS-CoV-2 Spike Protein. *Vaccines* **2023**, *11*, doi:10.3390/vaccines11050991.
3. Nordstrom, P.; Ballin, M.; Nordstrom, A. Risk of infection, hospitalisation, and death up to 9 months after a second dose of COVID-19 vaccine: a retrospective, total population cohort study in Sweden. *Lancet* **2022**, *399*, 814-823, doi:10.1016/s0140-6736(22)00089-7.

4. Ballester, J.; Quijal-Zamorano, M.; Méndez Turrubiates, R.F.; Pegenaute, F.; Herrmann, F.R.; Robine, J.M.; Basagaña, X.; Tonne, C.; Antó, J.M.; Achebak, H. Heat-related mortality in Europe during the summer of 2022. *Nature Medicine* **2023**, *29*, 1857-1866, doi:10.1038/s41591-023-02419-z.
5. The Economist. Expensive energy may have killed more Europeans than covid-19 last winter. **2023**, *May 10th*.
6. Eurostat. Excess mortality by month. Available online: [https://ec.europa.eu/eurostat/databrowser/view/DEMO\\_MEXRT\\_\\_custom\\_1210067/bookmark/table?lang=en&bookmarkId=fc27a3a9-082b-461d-830b-a4c7b36caf4f](https://ec.europa.eu/eurostat/databrowser/view/DEMO_MEXRT__custom_1210067/bookmark/table?lang=en&bookmarkId=fc27a3a9-082b-461d-830b-a4c7b36caf4f) (accessed on Jul 15, 2023).
7. Eurostat. Population on Jan 1, 2022. Available online: <https://ec.europa.eu/eurostat/databrowser/view/tps00001/default/table?lang=en> (accessed on Nov 23, 2022).
8. European Centre for Disease Prevention and Control. COVID-19 Vaccine Tracker. Available online: <https://vaccinetracker.ecdc.europa.eu/public/extensions/COVID-19/vaccine-tracker.html#uptake-tab> (accessed on Jul 15, 2023).
9. Our World in Data. Coronavirus (COVID-19) Vaccinations. Available online: <https://ourworldindata.org/covid-vaccinations> (accessed on Jan 15, 2023).
10. Eurostat. Deaths and crude death rate. Available online: <https://ec.europa.eu/eurostat/databrowser/view/tps00029/default/table?lang=en> (accessed on Nov 15, 2022).
11. Eurostat. Life expectancy by age and sex. Available online: [https://ec.europa.eu/eurostat/databrowser/view/demo\\_mlexpec/default/table?lang=en](https://ec.europa.eu/eurostat/databrowser/view/demo_mlexpec/default/table?lang=en) (accessed on Nov 23, 2022).
12. CIA. The World Factbook: Median Age. Available online: <https://www.cia.gov/the-world-factbook/field/median-age/country-comparison/> (accessed on Nov 23, 2022).
13. Eurostat. GDP per capita in PPS. Available online: <https://ec.europa.eu/eurostat/databrowser/view/tec00114/default/table?lang=en> (accessed on Nov 23, 2022).
14. O'Brien, R.M. A caution regarding rules of thumb for variance inflation factors. *Qual. Quant.* **2007**, *41*, 673-690.
15. Raudenbush, S.W.; Bryk, A.S. *Hierarchical Linear Models: Applications and Data Analysis Methods*, 2nd ed.; Sage: Thousand Oaks, CA, 2002.
16. Snijders, T.A.B. Multilevel Analysis. In *International Encyclopedia of Statistical Science*, Lovric, M., Ed.; Springer Berlin Heidelberg: Berlin, Heidelberg, 2011; pp. 879-882.
17. Cook, T.D.; McCleary, R.; McCain, L.J.; Reichardt, C.S.; Fankhauser, G.; Campbell, D.T. *Quasi-experimentation : design & analysis issues for field settings*; Houghton Mifflin Co.: Boston, 1979.
18. Hsieh, J.J. Encyclopedia Britannica. Ecological fallacy. Available online: <https://www.britannica.com/science/ecological-fallacy> (accessed on 26 Jan 2023).
19. Robinson, W.S. Ecological Correlations and the Behavior of Individuals. *Am. Sociol. Rev.* **1950**, *15*, 351-357, doi:10.2307/2087176.
20. Subramanian, S.V.; Jones, K.; Kaddour, A.; Krieger, N. Revisiting Robinson: The perils of individualistic and ecologic fallacy. *International Journal of Epidemiology* **2009**, *38*, 342-360, doi:10.1093/ije/dyn359.
21. Blyth, C.R. On Simpson's Paradox and the Sure-Thing Principle. *J. Am. Stat. Assoc.* **1972**, *67*, 364-366, doi:10.1080/01621459.1972.10482387.
22. Simpson, E.H. The Interpretation of Interaction in Contingency Tables. *Journal of the Royal Statistical Society: Series B (Methodological)* **1951**, *13*, 238-241, doi:<https://doi.org/10.1111/j.2517-6161.1951.tb00088.x>.
23. Rothman, K.J. A pictorial representation of confounding in epidemiologic studies. *Journal of Chronic Diseases* **1975**, *28*, 101-108, doi:[https://doi.org/10.1016/0021-9681\(75\)90066-1](https://doi.org/10.1016/0021-9681(75)90066-1).
24. Loney, T.; Nagelkerke, N.J. The individualistic fallacy, ecological studies and instrumental variables: a causal interpretation. *Emerging Themes in Epidemiology* **2014**, *11*, 18, doi:10.1186/1742-7622-11-18.

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