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Original Research

Vaccination against SARS-CoV-2 protects from morbidity, mortality and sequelae from COVID19 in patients with cancer



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KEYWORDS

COVID-19; SARS-CoV-2; Vaccines; Prevention; Cancer; Clinical efficacy **Abstract** *Background:* Although SARS-CoV-2 vaccines immunogenicity in patients with cancer has been investigated, whether they can significantly improve the severity of COVID-19 in this specific population is undefined.

Methods: Capitalizing on OnCovid (NCT04393974) registry data we reported COVID-19 mortality and proxies of COVID-19 morbidity, including post-COVID-19 outcomes, according to the vaccination status of the included patients.

Results: 2090 eligible patients diagnosed with COVID-19 between 02/2020 and 11/2021 were included, of whom 1930 (92.3%) unvaccinated, 91 (4.4%) fully vaccinated and 69 (3.3%) partially vaccinated. With the exception of a higher prevalence of patients from the UK (p = 0.0003) and receiving systemic anticancer therapy at COVID-19 diagnosis (p = 0.0082) among fully vaccinated patients, no demographics/oncological features were associated with vaccination status. The 14-days case fatality rate (CFR) (5.5% vs 20.7%, p=0.0004) and the 28-days CFR (13.2% vs 27.4%, p=0.0028) demonstrated a significant improvement for fully vaccinated patients in comparison with unvaccinated patients. The receipt of prior full vaccination was also associated with reduced symptomatic COVID-19 (79.1% vs 88.5%, p = 0.0070), need of COVID-19 oriented therapy (34.9% vs 63.2%, p = 0.0070)p < 0.0001), complications from COVID-19 (28.6% vs 39.4%, p = 0.0379), hospitalizations due to COVID-19 (42.2% vs 52.5%, p = 0.0007) and oxygen therapy requirement (35.7% vs 52%, p = 0.0036). Following Inverse Probability Treatment Weighting (IPTW) procedure no statistically significant difference according to the vaccination status was confirmed; however, all COVID-19 related outcomes were concordantly in favour of full vaccination. Among the 1228 (58.8%) patients who underwent a formal reassessment at participating centres after COVID-19 resolution, fully vaccinated patients experienced less sequelae than unvaccinated patients (6.7% vs 17.2%, p = 0.0320).

Conclusions: This analysis provides initial evidence in support of the beneficial effect of SARS-CoV-2 vaccines against morbidity and mortality from COVID-19 in patients with cancer. © 2022 Elsevier Ltd. All rights reserved.

1. Introduction

Patients with cancer are intrinsically more vulnerable to morbidity and mortality from Coronavirus Disease 2019 (COVID-19) [1,2]. All levels of cancer care including screening, diagnosis and treatment have been heavily impacted by the Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2) pandemic [3,4]. In addition to the threat imposed by acute morbidity and mortality from COVID-19 in cancer patients, recent evidence highlights that oncological continuity of care can be further disrupted by long-term consequences of COVID-19, which affect approximately 15% of patients with cancer who recover from the acute phase [5].

Widespread vaccination against SARS-Cov-2 represents a highly important public health measure to reduce the severity and lethality of SARS-Cov-2.

The immunogenicity and safety profile of SARS-Cov-2 vaccines have been investigated across several types of cancers and irrespective of exposure to recent anticancer therapy [6–8]. Evidence of seroconversion following SARS-Cov-2 vaccination confirmed an antibody response in >90% of patients with solid tumours [9,10], which is comparable to the general population [11]. However, several reports highlight that a proportion of patients with cancer, such as those with haematological malignancies undergoing anti-CD20 treatments, elicit a diminished immune response to vaccines with seroconversion rates of <60% [9,10].

Evidence on the efficacy of SARS-Cov-2 vaccines from randomized clinical trials is limited to patients with stable oncological disease and off immunosuppressive anti-cancer therapy at the time of vaccination [12]. Therefore, unresolved questions exist around whether efficacy of SARS-CoV-2 vaccinaes is independent from anti-cancer treatments and whether vaccinal immunity protects from long-term consequences of COVID-19.

With these premises in mind, we performed a dedicated update of the OnCovid registry to provide initial evidence regarding the magnitude of clinical benefit of SARS-CoV-2 vaccines in influencing outcomes of COVID-19 in a large real-world oncological population.

2. Study design and outcomes

OnCovid (NCT04393974) is a European registry study that collects data from consecutive patients with solid/haematologic malignancy diagnosed with COVID-19.

By the data lock of 22/12/2021, the registry included 3237 patients diagnosed with COVID-19 between 27/02/2020 and 30/11/2021. Patients with unknown vaccination statuses were excluded. Similarly, to maintain consecutive accrual, we excluded centres that did not actively enrol subjects during the March—December accrual timeframe. A list of participating centres with eligible patients for this analysis is provided in Supplementary Table 1.

The primary objective of this study was to describe COVID-19 mortality in patients with cancer according to SARS-CoV-2 vaccination status. As secondary objectives, we estimated the impact of COVID-19 vaccines on COVID-19 symptoms and morbidity. In addition, we evaluated whether receipt of SARS-CoV-2 vaccination was associated with the occurrence of COVID-19 sequelae among patients who underwent a clinical reassessment at the participating centres.

Patients were categorized as fully vaccinated at the time of COVID-19 diagnosis if they had received two doses of the BNT162b2, mRNA-1273 and ChAdOx1-S vaccines or in case of infection diagnosed at least 28 days after a single dose of the Ad.26.COV2.S vaccine [13]. Patients who received at least one vaccination, without meeting the above-mentioned criteria, were considered partially vaccinated.

Acknowledging the competing influence of the underlying malignancy in determining clinical outcomes, we elected the all-cause 14-days CFR as the clinical end points of interest, in an attempt of differentiating early (COVID-19 related) from late (cancer-related) mortality as already done in with our registry [14]. Considering the limited number of 14-days events recorded among fully vaccinated patients, which prevented the planned Inverse Probability of Treatment Weighting (IPTW) procedure, we also evaluated the 28-days CFR.

First, we reported the distribution of key demographics and oncological characteristics consistently associated with clinical outcomes in the study population [5,14–18] across the vaccination categories, subsequently, we analyzed COVID-19 related outcomes according to the vaccination status, with a formal comparison between fully vaccinated and unvaccinated patients.

We then reported the prevalence and distribution of COVID-19 associated symptoms, and as proxies of COVID-19 morbidity, we analyzed other COVID-19 related outcomes reproducibly described in the registry [5,14–17], including the need of COVID-19 oriented therapy, the incidence of COVID-19 complications, the hospitalization rate and the need of oxygen therapy.

Although recognizing that the unbalanced sample size of the vaccination subgroups did not allow a powered and formal weighted comparison, we performed an exploratory IPTW procedure including key baseline demographics and oncological characteristics, to provide a preliminary adjusted estimation of the CFR and COVID-19 related outcomes.

Considering the evidence of a mild decrease over time of the antibody response to SARS-CoV-2 vaccination [19,20], we also provided a descriptive analysis of COVID-19 outcome according to vaccination timing, including only patients with an available date of vaccination. For this purpose, fully vaccinated patients were considered those who had received two doses of the

BNT162b2, mRNA-1273 and ChAdOx1-S vaccines at least 7 days before the infection and those who received one dose of the Ad.26.COV2.S vaccine at least 28 days before the breakthrough infection [13,21–23]. Patients who received at least one vaccination, without meeting the above-mentioned criteria, were considered partially vaccinated, while those patients diagnosed with COVID-19 more than 6-months following the complete vaccination were considered separately.

Lastly, in order to describe the potential role of SARS-CoV-2 vaccines in reducing the occurrence of COVID-19 sequelae in patients with cancer, we focused on those patients who underwent a formal clinical assessment at the participating centres after COVID-19 recovery as previously done [5], and reported the incidence of COVID-19 sequelae according to their vaccination status. Sequelae were assessed by treating physicians as per local practice, and categorized according to the system/organ involved into: respiratory symptoms (including dyspnoea and chronic cough), residual fatigue, weight loss, neuro-cognitive sequelae (including cognitive, visual impairment, ano/dysosmia – age/dysgeusia, headache, confusion, lethargy) and others (including other organs dysfunctions, residual fever, muscle cramps, arthralgia, skin conditions etc). For the purpose of the analysis, COVID-19 sequelae were further clustered as: respiratory (either alone or combined with other complications) and post-COVID-19 fatigue.

A detailed description of the study methodology and statistical analysis is provided in Supplementary eMethods.

3. Results

3.1. Vaccination and patients' characteristics

At the time of database lock, the registry included 3237 patients. A total of 129 patients were excluded due to the unconfirmed date of COVID-19 diagnosis and missing mortality outcome, another 133 patients were excluded because of unknown vaccination status. A further group of 885 patients previously entered from centres that did not enrol patients for the present update was further excluded to maintain consecutive accrual and minimize selection bias (Fig. 1).

In total, 2090 eligible patients (67.2%) were included in this analysis, including 1930 (92.3%) unvaccinated, 91 (4.4%) fully vaccinated and 69 (3.3%) partially vaccinated patients. Among fully vaccinated patients 54 (59.3%) received the BNT162b2 vaccine, 18 (19.8%) received the mRNA-1273 vaccine, 16 (17.6%) received the ChAdOx1-S vaccine and 3 (3.3%) received the Ad.26.COV2.S vaccine. Among partially vaccinated patients 32 (46.4%) received the BNT162b2 vaccine, 27 (39.1%) received the mRNA-1273 vaccine and 10 (14.5%) received the ChAdOx1-S vaccine.

Table 1 reports the detailed distribution of demographics and oncological characteristics across the vaccination subgroups. As compared to unvaccinated patients, there was a higher prevalence of patients from the United Kingdom (49.5% vs 31.5%, p = 0.0003) and receiving systemic anticancer therapy at COVID-19 diagnosis (54.5% vs 40.3%, p = 0.0082) among fully vaccinated patients, while no other characteristics

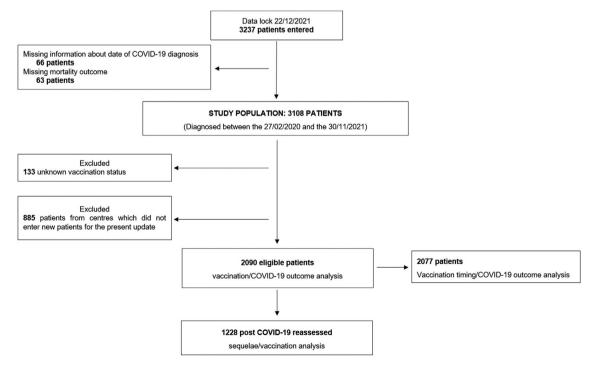


Fig. 1. Study flow diagram.

Table 1
Demographics and oncological characteristics of eligible patients according to the vaccination status.

	Unvaccinated	Fully vaccinated	P-value	$\frac{\text{Partially vaccinated}}{\text{N} = 69 (\%)}$
	N = 1930 (%)	N = 91 (%)		
Country				
United Kingdom	607 (31.5)	45 (49.5)	0.0003	35 (50.7)
Spain	712 (36.9)	32 (35.2)		26 (37.7)
Italy	611 (31.7)	14 (15.4)		8 (11.6)
Sex	` '	` ′		, ,
Male	1043 (54.2)	43 (47.3)	0.1970	35 (50.7)
Females	883 (45.8)	48 (52.7)		34 (49.3)
Missing	4	_ ` ´		_ ` ´
Age				
<65 years	753 (39.2)	44 (48.4)	0.0798	26 (38.8)
≥65 years	1170 (60.8)	47 (51.6)		41 (61.2)
Missing	6	_ ` '		2
Comorbidities				
0-1	1031 (53.4)	48 (52.7)	0.9000	32 (46.4)
>2	899 (46.6)	43 (47.3)		37 (53.6)
Smoking history		()		. ()
Never smokers	818 (50)	34 (41)	0.1095	24 (41.4)
Former/current smokers	819 (50)	49 (59)	0.10,0	34 (58/6)
Missing	293	8		11
Primary Tumour				
Breast	313 (16.4)	16 (17.6)	0.5284	13 (18.8)
Gastrointestinal	385 (20.1)	15 (16.5)		12 (17.4)
Gynaecological/Genito-Urinary	344 (18.0)	21 (23.1)		13 (18.8)
Thoracic	301 (15.7)	15 (16.5)		13 (18.8)
Others	225 (11.8)	13 (14.3)		10 (14.5)
Haematologic	346 (18.1)	11 (12.1)		8 (11.6)
Missing	16	= (1211)		- -
Tumour stage				
Local/loco-regional	833 (47.2)	34 (40)	0.1959	31 (47)
Advanced	933 (52.8)	51 (60)	******	35 (53)
Missing	164	6		3
Tumour status at COVID-19 diagnosis	101	Ü		3
Remission/non measurable disease	679 (35.5)	30 (33)	0.6249	27 (39.7)
Active malignancy	1235 (64.5)	61 (67)	0.0219	41 (60.3)
Missing	16	- -		1
SACT at COVID-19 diagnosis ^a	10			•
No	1103 (59.7)	40 (45.5)	0.0082	33 (53.2)
Yes	746 (40.3)	48 (54.5)	0.0002	29 (46.8)
Missing	81	3		7
Timing of infection	01	3		,
Post-vaccination phase	560 (29.0)	91 (100)		69 (98.6)
Pre-vaccination phase	1370 (71.0)	- (100)		- (50.0)
rie-vaccination phase	13/0 (/1.0)			

SACT: systemic anticancer therapy.

including sex, age, comorbidities burden and tumour features were significantly associated with vaccination status. Of note, primary tumour types were well balanced across vaccination subgroups. Importantly, the great majority of patients diagnosed after the approval of the first SARS-CoV-2 vaccine in the UK were still unvaccinated at COVID-19 diagnosis (560 vs 91/69 fully and partially vaccinated patients, respectively).

3.2. Full vaccination is associated with improvement in COVID-19 morbidity and mortality

The 14-days CFR were significantly lower in fully vaccinated patients (5.5%) in comparison with

unvaccinated patients (20.7%, p = 0.0004). Similarly, the 28-days CFR was significantly lower among fully vaccinated patients compared with unvaccinated patients (13.2% vs 27.4%, p = 0.0028). Compared to unvaccinated patients, the receipt of a full vaccination course was associated with improved morbidity from COVID-19, as showed by the significant reduction in the rates of symptomatic COVID-19 (79.1% vs 88.5%, p = 0.0070), provision of COVID-19-specific therapy (34.9% vs 63.2%, p < 0.0001), acute complications from COVID-19 (28.6% vs 39.4%, p = 0.0379), hospitalizations due to COVID-19 (42.2% vs 52.5%, p = 0.0007) and requirement for oxygen therapy (35.7% vs 52%, p = 0.0036). Table 2 provides a summary of COVID-19

^a Within 4 weeks prior to COVID-19 diagnosis.

Table 2 Summary of COVID-19 related outcomes according to the vaccination status.

	Unvaccinated	Fully vaccinated $N = 91 (\%)$	P-value	$\frac{\text{Partially vaccinated}}{\text{N} = 69 (\%)}$
	N = 1930 (%)			
14-Days case fatality rate				
Alive	1530 (79.3)	86 (94.5)	0.0004	59 (85.5)
Death events	400 (20.7)	5 (5.5)		10 (14.5)
28-Days case fatality rate				
Alive	1401 (72.6)	79 (86.8)	0.0028	55 (79.7)
Death events	529 (27.4)	12 (13.2)		14 (20.3)
COVID-19 symptoms				
No	222 (11.5)	19 (20.9)	0.0070	10 (14.5)
Yes	1708 (88.5)	72 (79.1)		59 (85.5)
Need of COVID-19 oriented therap	y			
No	676 (36.8)	56 (65.1)	< 0.0001	37 (55.2)
Yes	1162 (63.2)	30 (34.9)		30 (44.8)
Missing	92	5		2
Complications from COVID-19				
No	1169 (60.6)	65 (71.4)	0.0379	43 (62.3)
Yes	761 (39.4)	26 (28.6)		26 (37.7)
Hospitalization				
Not required	431 (22.5)	36 (40.0)	0.0007	15 (22.1)
Required due to COVID-19	1006 (52.5)	38 (42.2)		31 (45.6)
Pre-existing	479 (25.0)	16 (17.8)	_	22 (32.4)
Missing	14	1		1
Oxygen therapy				
No	884 (48.0)	54 (64.3)	0.0036	40 (60.6)
Yes	956 (52.0)	30 (35.7)		26 (39.4)
Missing	90	7		3

related outcomes according to the vaccination status, also visualized in Fig. 2A and reported as unadjusted OR in Fig. 3A.

IPTW analysis showed trends towards improvement in COVID-19 outcomes for fully vaccinated patients, however no statistically significant difference according to vaccination status can be confirmed for the 28-days CFR (adjusted odds ratio – AOR 0.32, 95% Confidence Intervals (CI): 0.01–13.34), COVID-19 symptoms (AOR 0.46, 95%CI: 0.02-8.19), need of COVID-19 oriented therapy (AOR 0.29, 95%CI: 0.02-3.65), oxygen therapy (AOR 0.43, 95%CI: 0.03-6.18), complicafrom COVID-19 (AOR 0.48,0.03-7.90) and hospitalization due to COVID-19 (AOR 0.27, 95%CI: 0.02-4.71) (Fig. 3B). Multivariable logistic models for each outcome are reported in Supplementary Table 2.

3.3. Time-dependent characteristics of the relationship between SARS-Cov-2 vaccination and outcomes from COVID-19 infection

In view of the time-dependent efficacy of SARS-Cov-2 vaccines [20], we postulated whether the improvement in COVID-19 outcomes seen in fully vaccinated patients could change as a function of time. The exact dates of all vaccination doses were available for 147 patients. For this analysis 70 patients (47.6%) were considered partially vaccinated, 64 patients (43.5%) fully vaccinated \geq 6 months from the infection and 13 patients (8.9%) fully

vaccinated more than 6 months prior to the date of infection. Among partially vaccinated patients 33 (47.1%) received the BNT162b2 vaccine, 27 (38.6%) received mRNA-1273 and 10 (14.3%) received ChAdOx1-S. Among fully vaccinated patients, 31 (48.4%) received the BNT162b2 vaccine, 17 (26.6%) received mRNA-1273, 13 (20.3%) ChAdOx1-S and 3 (4.7%) Ad.26.COV2.S. Among the patients vaccinated ≥6 months from the infection, 11 (84.6%) received BNT162b2, 1 (7.7%) mRNA-1273 and 1 (7.7%) ChAdOx1-S. As summarized in Supplementary Table 3 and Supplementary Fig. 1, we observed a trend towards incremental improvement in all COVID-19-related outcomes across unvaccinated/ partially vaccinated to fully vaccinated patients, followed by an increasing trend among patients vaccinated ≥6 months from the infection.

3.4. Vaccination against SARS-CoV-2 is associated with a lower prevalence of COVID-19 sequelae in patients with cancer

By the data lock, 1228 (58.8%) of the eligible patients underwent a formal clinical reassessment at participating centres after a median time of 40 days from COVID-19 diagnosis (Inter quartile range: 25–68). Baseline demographics and oncological characteristics stratified by vaccination status are summarized in Supplementary Table 4. Similar to what was reported for the overall population, fully vaccinated patients were more likely from the United Kingdom (46.7% vs 21.9%,

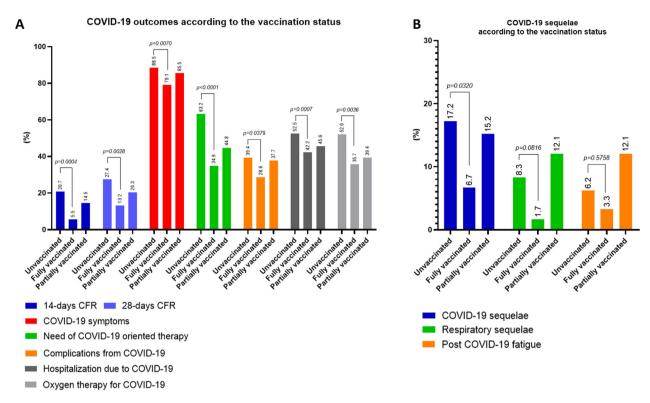


Fig. 2. Summary of COVID-19 outcomes according to vaccination status (A). Patients were categorized as fully vaccinated at the time of COVID-19 diagnoses if they had received two doses for the BNT162b2, mRNA-1273, and ChAdOx1-S vaccines or in case of infection diagnosed at least 28 days after a single dose of the Ad.26.COV2.S vaccine. (B) Summary COVID-19 sequelae analysis according to the vaccination status.

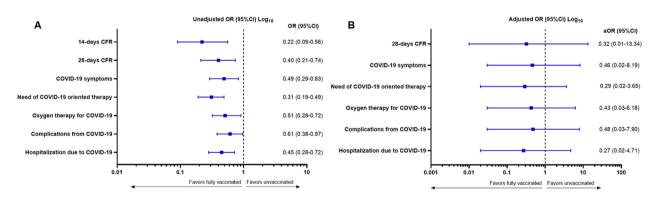


Fig. 3. Forest plot graph reporting the (A) unadjusted odds ratio (OR) and the (B) adjusted odds ratio (aOR) from the Inverse Probability of Treatment Weighting (IPTW) fitted multivariable logistic regression models for each COVID-19 outcomes. The following covariates were included in each model: country (United Kingdom vs Spain vs Italy), biological sex (male vs female), age (\geq 65 vs < 65 years), number of co-morbidities (\geq 2 vs 0–1), tumour status (presence of active vs non-active disease), and the receipt of systemic anticancer therapy (SACT) within 4 weeks of SARS-CoV-2 infection (yes vs no). CFR: case fatality rate. Complete multivariable models for each COVID-19 outcomes are reported in Supplementary Table 2.

p = 0.0003) and were receiving systemic anticancer therapy (SACT) at COVID-19 diagnosis (62.7% vs 47.6%, p = 0.0235). No other feature was associated with vaccination status.

Overall, 199 patients (16.2%) reported at least one COVID-19 sequela. As shown in Fig. 2B, the proportion of patients reporting at least 1 sequela from COVID-19 was significantly lower in fully vaccinated patients compared to unvaccinated controls (6.7% vs 17.2%,

p = 0.0320), with no difference in the distribution of individual type of sequelae across groups (Supplementary Table 5).

4. Discussion

Vaccinal immunity to SARS-CoV-2 has radically changed the natural history of COVID-19. Whilst only partially effective in controlling viral transmission,

especially after the emergence of novel variants of concern, vaccines remain widely effective in reducing the severity of COVID-19 [21–24]. However, their remarkable clinical efficacy has been only partially demonstrated in patients with cancer [12].

Recently, data from the COVID-19 and Cancer Consortium (CCC19) on 54 fully vaccinated patients reported comparable rates of mortality and risk of adverse outcomes from COVID-19 irrespective of vaccination status, highlighting that vaccine protection vaccination may be incomplete in patients with cancer and supporting the need for further investigation in independent cohorts [25].

In this analysis of the OnCovid registry, we documented for the first time that patients who contracted SARS-Cov-2 after full vaccination were characterized by a lower probability of severe COVID-19 and mortality compared to unvaccinated controls.

Univariable analyses demonstrate a reduction in CFR 14- and 28-days post-infection for fully vaccinated patients compared to unvaccinated patients. All indices of COVID-19 morbidity showed a protective effect for fully vaccinated patients, including COVID-19 symptoms, the requirement for COVID-19-oriented therapy and oxygen therapy, complications and hospitalization rates due to COVID-19.

The retrospective design of our study and the relatively low proportion of fully vaccinated subjects by data cut-off underscores the preliminary nature of our findings. Baseline characteristics were comparable across exposed and unexposed groups, lending credence to the view that the improvement in outcomes observed in vaccinated patients may be truly due to SARS-Cov-2 immunity. While primary analyses confirmed our hypothesis, IPTW models yielded non-statistically significant trends towards improvement of outcomes, as a likely result of a largely unbalanced sample size of patients' subgrouping.

In a clinical setting that is derived from solid level I evidence and within the limitation of a registry study, our findings provide a meaningful contribution to the growing body of knowledge demonstrating the ability of SARS-CoV-2 vaccines as a measure to reduce adverse outcomes from the disease. Compelling evidence suggests how SARS-CoV-2 vaccination induces effective immune responses across different tumour types and irrespective of recent exposure to diverse anti-cancer therapies [6-8]. However, protective immunity is not universal in patients with cancer, where evidence of diminished immunogenicity has been shown in patients with haematological malignancies and after treatment with CD20 inhibitors [7,9,10]: a finding that may explain the heterogeneity of results reported by the CCC19, which included 35% of patients with haematological malignancies among those fully vaccinated [25], compared to 12% of our cohort.

Whilst the retrospective nature of our study does not allow us to conclude that the association between full vaccination and improved outcome is truly causal, documentation of CFRs of 5.5% in this subgroup is highly important as it provides an important new benchmark in a more contemporary and clinically relevant estimate of lethality from COVID-19 in the post-vaccinal era [14], as it impressively diverges from the >30% estimates reported during the initial phase of the pandemic across different registries [17,26,27].

Baseline demographics and oncological characteristics, all well balanced across groups, are an unlikely source of bias. The uneven distribution of the country of origin and SACT exposure can be easily explained by country-specific differences in the delivery of immunization campaigns, where vaccines were often offered alongside SACT in some centres.

The evidence that vaccine response may decrease over time [19,20] mirrors our descriptive analysis of vaccination timing, which suggests that the protection provided by SARS-CoV-2 vaccines declines beyond the 6-months landmark. These findings may also be related to the emergence of new variants of concern, such as the B.1.617.2 (delta), which has a higher transmissibility than previous strains and proved to cause a higher rate of breakthrough infections [28,29]. Taken together, these results support the need of prioritizing frail patients for booster doses, to sustain vaccine's immunogenicity over time [30].

Another important question that has not been addressed in any previous study in oncological patients is whether vaccinal immunity may affect the risk of developing COVID-19 sequelae. In a recent analysis of the OnCovid registry, we demonstrated that COVID-19 is detrimental to patients' outcome even beyond the acute phase of the disease. COVID-19 sequalae can affect ~15% of patients, with long-term consequences for their continuity of care and survival [5]. From this perspective, the additional analysis performed among COVID-19 survivors who underwent a clinical reassessment at participating centres suggests that the protection provided by vaccines extends beyond the acute phase, as supported by the reduced incidence of sequelae in fully vaccinated patients.

Our data collection relies on unplanned time intervals to capture vaccine administration, which could lead to determination bias. Moreover, breakthrough infections could have been asymptomatic with an associated risk of diminished reporting and underestimation of the effect. However, hard end points like mortality/severe disease were unlikely affected.

Notably, we did not discuss booster/third doses, as by the data lock only a small minority of breakthrough infections among patients who received it had been reported. However, considering the recently emerged B.1.1.529 (omicron) variant, its different immunogenic profile, and the wave of infections recorded in Europe between December 2021 and January 2022 [31,32], a

dedicated update is currently ongoing, to produce reliable dedicated evidence.

Although preliminary, this study provides novel evidence supporting the clinical efficacy against COVID-19 morbidity, mortality and sequelae of SARS-CoV-2 vaccines in a large real-world population of patients with cancer. Universal SARS-CoV-2 vaccination should remain a goal in the management of patients with cancer during and beyond the COVID-19 pandemic.

Role of the funding source

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Ethical approval and consent to participate

OnCovid was granted central approval by the United Kingdom Health Research Authority (20/HRA/1608) and by the corresponding research ethics committees at each participating institution. Full waiver of consent due to the retrospective nature of the study was granted by the UK HRA in accordance with UK law.

Authors' contributions

All authors contributed to the publication according to the ICMJE guidelines for the authorship. All authors read and approved the submitted version of the manuscript (and any substantially modified version that involves the author's contribution to the study). Each author has agreed both to be personally accountable for the author's own contributions and to ensure that questions related to the accuracy or integrity of any part of the work, even ones in which the author was not personally involved, are appropriately investigated, resolved and the resolution documented in the literature.

Consent for publication

Informed consent was waived by competent authorities due to anonymized nature of patient data and retrospective design of the study.

Availability of data and material

Study data made available upon reasonable request.

Conflict of interest statement

The authors declare the following financial interests/ personal relationships which may be considered as potential competing interests: As corresponding author of the abovementioned manuscript, I declare on behalf of my co-authors the following conflict of interests: David J Pinato received lecture fees from ViiV Healthcare, Bayer Healthcare, BMS, Roche, EISAI, Falk Foundation, travel expenses from BMS and Bayer Healthcare; consulting fees for Mina Therapeutics, EISAI, Roche, DaVolterra and Astra Zeneca; research funding (to institution) from MSD and BMS. Aleix Prat has declared personal honoraria from Pfizer, Roche, MSD Oncology, Eli Lilly, and Daiichi Sankyo; travel, accommodations, and expenses paid by Daiichi Sankyo; research funding from Roche and Novartis; and consulting/advisory role for NanoString Technologies, Amgen, Roche, Novartis, Pfizer and Bristol-Myers Squibb.Matteo Lambertini acted as consultant for Roche, Novartis, Lilly, AstraZeneca, Exact Sciences, MSD, Pfizer, Seagen and received speaker honoraria from Roche, Novartis, Lilly, Pfizer, Takeda, Ipsen and Sandoz outside the submitted work. Joan Brunet has declared consulting/advisory role for MSD and Astra Zeneca. Alessandra Gennari has declared consulting/ advisory role for Roche, MSD, Eli Lilly, Pierre Fabre, EISAI, and Daichii Sankyo; speakers bureau for Eisai, Novartis, Eli Lilly, Roche, Teva, Gentili, Pfizer, Astra Zeneca, Celgene, and Daichii Sankvo: research funds: EISAI, Eli Lilly, and Roche. CMV has received travel grants and other honoraria from BMS, MSD, Novartis and Roche. Gianluca Gaidano has declared consulting/ advisory role for Janssen, Abbvie, Astra-Zeneca and BeiGene, and speaker fees from Janssen and Abbvie.-Lorenza Rimassa received consulting fees from Taiho Oncology, Servier, Amgen, ArQule, AstraZeneca, Basilea, Bayer, BMS, Celgene, Eisai, Exelixis, Genenta, Hengrui, Incyte, Ipsen, IQVIA, Lilly, MSD, Nerviano Medical Sciences, Roche, Sanofi, Zymeworks; lecture fees from AbbVie, Amgen, Bayer, Eisai, Gilead, Incyte, Ipsen, Lilly, Merck Serono, Roche, Sanofi; travel expenses from Ipsen; and institutional research funding from Agios, ARMO BioSciences, AstraZeneca, Bei-Gene, Eisai, Exelixis, Fibrogen, Incyte, Ipsen, Lilly, MSD, Nerviano Medical Sciences, Roche, Zymeworks. Joseph Tabernero reported consulting fees from Array Biopharma, AstraZeneca, Avvinity, Bayer, Boehringer Ingelheim, Chugai, Daiichi Sankyo, F. Hoffmann-La Roche Ltd, Genentech Inc, HalioDX SAS, Hutchison MediPharma International, Ikena Oncology, Inspirna Inc, IQVIA, Lilly, Menarini, Merck Serono, Merus, MSD, Mirati, Neophore, Novartis, Ona Therapeutics, Orion Biotechnology, Peptomyc, Pfizer, Pierre Fabre, Samsung Bioepis, Sanofi, Seattle Genetics, Scandion Oncology, Servier, Sotio Biotech, Taiho, Tessa Therapeutics and TheraMyc. He also reported speaker's fees from Imedex, Medscape Education, MJH Life Sciences, PeerView Institute for Medical Education and Physicians Education Resource (PER). He also declared institutional research support from Amgen Inc. Array Biopharma Inc, AstraZeneca Pharmaceuticals LP. BeiGene, Boehringer Ingelheim, Bristol Myers Squibb, Celgene, Debiopharm International SA, F. Hoffmann-La Roche Ltd, Genentech Inc, HalioDX SAS, Hutchison MediPharma International, Janssen-Cilag SA, MedImmune, Menarini, Merck Health KGAA, Merck Sharp & Dohme, Merus NV, Mirati, Novartis Farmacéutica SA, Pfizer, Pharma Mar, Sanofi Aventis Recherche & Développement, Servier, Taiho Pharma USA Inc. Spanish Association Against Cancer Scientific Foundation and Cancer Research UK. Alessio Cortellini received consulting fees from MSD, BMS, AstraZeneca, Roche; speakers' fee from AstraZeneca, MSD, Novartis and Eisai. All remaining authors have declared no conflicts of interest.London, April 18th, 2022.

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Appendix A. Supplementary data

Supplementary data to this article can be found online at https://doi.org/10.1016/j.ejca.2022.04.036.

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