# **Supplemental Online Content**

Fu J, Reid SA, French B, et al; COVID-19 and Cancer Consortium (CCC19). Racial disparities in COVID-19 outcomes among Black and White patients with cancer. *JAMA Netw Open*. 2022;5(3):e224304. doi:10.1001/jamanetworkopen.2022.4304

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This supplemental material has been provided by the authors to give readers additional information about their work.

### eAppendix 1. CCC19 Data Collection and Quality Assurance

All information is retrieved from electronic health records and there is no direct contact with patients. With missing data, attempts to inquire further with active providers are pursued for completion of record. Information is abstracted from the electronic health records manually by designees of academic and community institutions. Each case report is assigned a "quality score," which is a numeric metric to define case reports as analytic (0-4) or non-analytic (>4) quality based on data problems classified as minor, moderate, and major. The quality score is elaborated in our prior publication and summarized in eTable 1.<sup>1</sup> Only records meeting a sufficient quality score (0-4 points, i.e., no major problems and at most one moderate problem) are included (eFigure 1).

**eAppendix 2.** Alphabetical List of Participants by Institution that Contributed at Least One Record to the Analysis

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# eAppendix 3. Statistical Analysis Plan

| Approved Project Title            | Racial and Ethnic Inequities and Disparities in Clinical<br>Characteristics and Outcomes of Patients with Cancer and<br>COVID - 19 |
|-----------------------------------|--|
| Approved Project PI               | Julie Fu   |
| Name of the investigator          | Dimpy Shah   |
| completing this survey            | 15   |
| Proposed milestone deadline for   | 9/3/20   |
| this manuscript                   |  |
| Name and emails of (at most) 2    | Sonya Reid; Oscar Serrano  |
| additional project team           |  |
| members who would like to be      |  |
| part of the analysis team for the |  |
| project                           |  |
| 1 (a) Manuscript Title            | Racial and Ethnic Inequities and Disparities in Clinical<br>Characteristics and Outcomes of Patients with Cancer and<br>COVID-19   |
| 1 (b) Provide in the abstract an  | The novel SARS-CoV2 virus and its resulting illness,   |
| informative and balanced          | COVID-19, has led to a global pandemic resulting in over   |
| summary of what was done and      | 12 million cases worldwide and over 3 million cases in the   |
| what will be found                | United States (US).1 Initial reports implicate age, sex, and   |
|                                   | comorbid conditions as critical factors in determining the   |
|                                   | outcome from this illness. Most studies assessing outcomes   |
|                                   | by small sample size. One of the early reports from  |
|                                   | Wuhan China reviewed 28 COVID-19-infected cancer   |
|                                   | nation to with more than half experiencing severe outcomes   |
|                                   | and death in 28% of patients 2 It is postulated that cancer-   |
|                                   | directed treatment may be associated with severe events.   |
|                                   | These observations underscored the severity of COVID-19-   |
|                                   | infected patients with cancer and led to recommendations   |
|                                   | on COVID-19 screening and avoidance or dose  |
|                                   | modification of immunosuppressive treatments in these  |
|                                   | patients.2 Albeit limited, data from the US has  |
|                                   | corroborated worse outcomes following COVID-19 in  |
|                                   | patients with cancer.3,4 Based on recent disease-tracking  |
|                                   | dashboards, COVID-19 has been reported to  |
|                                   | disproportionately affect Blacks at higher rates compared  |
|                                   | to Non-Hispanic Whites (NHW).5 Blacks also have higher   |
|                                   | rates of hospitalization and death after contracting   |
|                                   | COVID-19.6-8 In New York City, Blacks had a  |
|                                   | substantially higher mortality rate (92.3 deaths per   |

|                            | 100,000) compared to Whites (45.2) and Asians (34.5).9  |  |  |
|----------------------------|---|--|--|
|                            | Similarly, in Chicago, Blacks accounted for 50% of COVID-   |  |  |
|                            | 19 cases and nearly 60% of COVID-19 related deaths even<br>though they only account for 30% of the overall  |  |  |
|                            |   |  |  |
|                            | population.10 Similar observations of racial and ethnic   |  |  |
|                            | disparities in impact of COVID-19 have been made in   |  |  |
|                            | Louisiana,11 New Jersey,12 Georgia,13 Michigan14 and  |  |  |
|                            | Connecticut.15 Despite these early reports, there is a paucity of dataon outcomes of COVID-19 infection in  |  |  |
|                            |   |  |  |
|                            | patients with cancer, stratified by race and ethnicity.<br>Factors contributing to these disparities are complex and<br>likely constitue an interplay of socioeconomic status, pre-<br>existing comorbid conditions, cancer status at the time of<br>infection, and access to care. Prior to COVID-19 pandemic,<br>it was well known that Black patients with cancer have the |  |  |
|                            |   |  |  |
|                            |   |  |  |
|                            |   |  |  |
|                            |   |  |  |
|                            |   |  |  |
|                            | highest death rates compared to all other racial and ethnic   |  |  |
|                            | groups of patients with cancer.16 Given the higher rates of   |  |  |
|                            | COVID-19-related mortality reported in minorities in the  |  |  |
|                            | general population, we hypothesize that Black patients<br>with cancer would have significantly worse outcomes than  |  |  |
|                            |   |  |  |
|                            | NHW patients with cancer, after accounting for  |  |  |
|                            | confounding variables.  |  |  |
| State specific objectives, | 1. Racial/Ethnic inequalities and inequities in baseline  |  |  |
| including any prespecified | characteristics and severity of presentation at the   |  |  |
| hypotheses                 | time of COVID-19 Diagnosis (Non-Hispanic Blacks   |  |  |
|                            | (NHB) versus NHW patients).   |  |  |
|                            |   |  |  |
|                            | a. To identify disparities in demographic,  |  |  |
|                            | socioeconomic, clinical characteristics (including  |  |  |
|                            | status of cancer and anti-cancer treatment), and  |  |  |
|                            | ECOG performance status) at the time of COVID-  |  |  |
|                            | 19 diagnosis between minority (NHB) and NHW   |  |  |
|                            | patients with cancer.   |  |  |
|                            |   |  |  |
|                            | D. To describe disparities in initial severity of COVID-  |  |  |
|                            | 19 Infection at the time of presentation between  |  |  |
|                            | minority (INHB) and INHW patients with cancer,  |  |  |
|                            | within the context of racial inequalities.  |  |  |
|                            |   |  |  |
|                            | 2 Racial/Ethnic Disparities in Clinical Complications   |  |  |
|                            | and Outcomes  |  |  |
|                            |   |  |  |
|                            | 1   |  |  |

|                                    | a. To assess disparities in COVID-19 severity (ordinal outcome) and 30-day all-cause mortality rate |  |
|------------------------------------|---|--|
|                                    | cancer, after adjusting for other prognostic  |  |
|                                    | covariates. Hypothesis: Minority patients with  |  |
|                                    | cancer will have higher 30-day all-cause mortality  |  |
|                                    | Tate compared to write patients with cancer.  |  |
|                                    | b. To assess disparities in incidence of clinical   |  |
|                                    | complications (see appendix I) between minority   |  |
|                                    | (NHB) and NHW patients with cancer.<br>Hypothesis: Minority patients with cancer will               |  |
|                                    | have significantly higher rates of clinical   |  |
|                                    | complications compared to NHW patients with   |  |
|                                    | cancer.   |  |
|                                    | c. To assess disparities in rates of hospitalization, ICU   |  |
|                                    | admission, and mechanical ventilation between   |  |
|                                    | minority (NHB) and NHW patients with cancer.  |  |
|                                    | Hypothesis: Minority patients with cancer will  |  |
|                                    | nave significantly higher rates of hospitalization,   |  |
|                                    | compared to NHW patients with cancer.   |  |
| Setting                            | The COVID-19 and Cancer Consortium (CCC19)  |  |
|                                    | (NCT04354701) is the largest international cohort study   |  |
|                                    | examining the clinical characteristics, course of illness, and                                      |  |
|                                    | outcomes of COVID-19 in patients with cancer. The CCC19   |  |
|                                    | database is uniquely positioned to answer these critical  |  |
|                                    | researchers, and policy makers about the burden of this   |  |
|                                    | pandemic on minorities. Data on all patients available at   |  |
|                                    | the time of analysis  |  |
| (a) Give the eligibility criteria, | CCC19 records of all NHB and NHW patients with cancer   |  |
| and the sources and methods of     | and lab-confirmed SARS-CoV-2 diagnosis in US, and   |  |
| selection of participants.         | race/ethnicity data present at the time of the analyses. Each                                       |  |
| Describe memods of follow-up       | NHW): Age > 18 years  |  |
|                                    | Exclusion criteria: Quality score >4  |  |
|                                    | Non-invasive cancers and premalignant conditions  |  |
|                                    | Non-melanoma non-invasive skin cancers (exclude if no   |  |
|                                    | confirmation)   |  |

|   | Incomplete follow-up resulting in unknown COVID-19 severity   |
|---|---|
| (b) For matched studies, give<br>matching criteria and number of<br>exposed and unexposed | Not applicable  |
| Outcomes  | Objective 1a:   |
|   | <ul> <li>baseline demographic, socioeconomic, clinical<br/>characteristics (including status of cancer and anti-cancer<br/>treatment), comorbidity (cardiac, renal, pulmonary,<br/>diabetes), ECOG performance status, and severity of<br/>presentation of COVID-19</li> <li>1b.<br/>overall baseline health status (including clinical laboratory<br/>markers) between NHW and NHB in hospitalized patients</li> </ul> |
|   | <ul> <li>2a:</li> <li>Primary outcome measure will be ordinal variable with following COVID-19 severity (0=uncomplicated, 1=hospitalization, 2=ICU admission, 3=mechanical ventilation, 4=death). No time restriction for recording ordinal outcome</li> <li>Secondary outcome will be 30-day all-cause mortality for multivariable modeling</li> </ul>   |
|   | <ul> <li>2b.</li> <li>Simple summary table stratified by race that gives n(%) for:</li> <li>1. clinical systemic complications (see appendix I)</li> <li>2. total hospitalization</li> <li>3. total mechanical ventilation</li> <li>4. total ICU admission</li> <li>5. overall Death</li> </ul>   |
| Exposures   | Race (Black vs NHW)   |
| Potential confounders   | Age, gender, smoking, obesity, comorbidities (cardio,<br>pulmonary, renal, diabetes), ECOG, cancer type/ status<br>anti-cancer therapy, calendar time, census region of<br>patient's residence  |
|   |   |
| Diagnostic criteria (if applicable)   | N/A   |

| For each variable of interest,<br>give sources of data and details<br>of methods of assessment<br>(measurement). Describe<br>comparability of assessment<br>methods if there is more than<br>one group<br>Explain how the study size was<br>arrived at | CCC19 database  |
|--|---|
| Explain how quantitative<br>variables will be handled in the<br>analyses. If applicable, describe<br>which groupings will be chosen<br>and why   | Age will be treated as continuous variable. Spline plot to<br>identify cutoff thresholds. The remainder of the variables,<br>including lab values, under examination are categorical in<br>nature.  |
| (a) Describe all statistical<br>methods, including those to be<br>used to control for confounding  | <ul> <li>Objective 1 will assess differences in baseline<br/>demographic, socioeconomic, comorbidities, clinical<br/>characteristics (including status of cancer and anti-cancer<br/>treatment), ECOG performance status, and severity of<br/>presentation of COVID-19 between each of the racial<br/>group comparisons. After checking for the accuracy,<br/>integrity, and distribution of the data, all characteristics<br/>and outcomes will be presented using descriptive<br/>statistics. We will provide the median and interquartile<br/>range (IQR) for continuous variables. Counts and<br/>percentages will be used to describe the binary and<br/>categorical variables.</li> <li>1b. Descriptive table restricted to hospitalized patients:<br/>laboratory measurements</li> <li>2a. Primary outcome measure will be ordinal variable and<br/>secondary outcome will be 30-d all-cause mortality for<br/>multivariable modeling. All <i>a priori</i> variables (but not<br/>baseline severity) and significant interactions will be<br/>included in the final MV model.</li> <li>We will use the e value to quantify sensitivity to<br/>unmeasured confounding.</li> <li>We will perform analysis based on inverse probability of<br/>treatment weighted (IPTW) methods. First, we will<br/>estimate propensity scores from a logistic regression</li> </ul> |
|  | treatment weighted (IPTW) methods. First, we will<br>estimate propensity scores from a logistic regression<br>model for which the outcome is a binary indicator of non-   |

| Hispanic Black versus non-Hispanic White race and             |
|---|
| prespecified covariates. For each patient, a weight will be   |
| calculated equal to the reciprocal of the probability of      |
| "receiving the treatment" (that is, race) that was "actually  |
| received." which will be estimated from the propensity        |
| score model   |
| Next we will use graphics and summary statistics to           |
| avaluate the propensity score model. The empirical            |
| distributions of the propensity scores will be stratified by  |
| race will be plotted to evaluate their overlap between        |
| race will be plotted, to evaluate their overlap between       |
| groups. Mean propensity scores will be calculated             |
| stratified by race across quintiles of the propensity scores  |
| in the overall conort, to evaluate balance in the propensity  |
| scores between groups. Unweighted and weighted                |
| absolute standardized mean differences for demographic        |
| and clinical characteristics at COVID-19 diagnosis between    |
| non-Hispanic Black and non-Hispanic White patients will       |
| be calculated, to evaluate whether the two groups were        |
| balanced on their observed characteristics; an absolute       |
| standardized mean difference <0.1 indicated balance.          |
| Finally, we will estimate IPTW differences in COVID-19        |
| severity between non-Hispanic Black and non-Hispanic          |
| White patients from an ordinal logistic regression model      |
| that included an offset for (log) follow-up time. Between-    |
| group IPTW differences in 30-day all-cause mortality will     |
| be estimated from both a logistic regression model (to        |
| estimate odds ratios) and a modified Poisson regression       |
| model (to estimate relative risks). All models will include   |
| race as the sole covariate, weighted by the reciprocal of the |
| probability of "receiving the treatment" (that is, race) that |
| was "actually received," and will use a robust (a.k.a.        |
| sandwich) variance estimator to account for the               |
| uncertainty due to estimation of the weights (and for the     |
| modified Poisson model, to account for misspecification of    |
| the variance structure). Results will be reported as odds     |
| ratios (or relative risks) with 95% confidence intervals.     |
|   |
| Proportional odds assumption will be tested                   |
| 2h  |
| Simple summary table stratified by race that gives $p(%)$     |
| for   |
| 1 clinical systemic complications (see appendix I)            |
| 1. emiliar systemic complications (see appendix 1)            |

|   | 2. total hospitalization  |
|---|---|
|   | 3. total mechanical ventilation   |
|   | 4. total ICU admission  |
|   | 5. overall Death  |
| (b) Describe any methods that                     | We will also examine interaction between  |
| will be used to examine                           | 1. race and all comorbidities (cardio, pulmonary, renal,  |
| subgroups and interactions                        | diabetes), and  |
|   | 2. race and cancer status   |
|   | 3. race and obesity   |
|   |   |
|   | to understand the synergistic impact of these factors on  |
|   | mortality.  |
| (c) Explain how missing data<br>will be addressed | Multiple imputation will be used to impute missing and<br>unknown data for all variables included in the analysis,<br>with some exceptions: unknown ECOG performance score<br>and unknown cancer status will not be imputed and |
|   | values will not be imputed.   |
|   | Imputation will be performed on the largest dataset   |
|   | possible (that is, after removing test cases and other  |
|   | manual exclusions, but before applying specific exclusion   |
|   | criteria). At least 10 imputed datasets will be used.   |
| (d) If applicable, explain how                    | Not applicable  |
| loss to follow-up will be                         |   |
| addressed   |   |
| (e) Describe any sensitivity                      | Not applicable  |
| analyses  |   |
| Complete?   | Complete  |

# Appendix I: CCC19- Black vs White Clinical systemic complication

| Multisystem organ failure* |                         |
|----------------------------|-------------------------|
| Sepsis*                    |                         |
| Bleeding*                  |                         |
| DIC*                       |                         |
| Pulmonary complications*   | Respiratory Failure     |
|                            | Pneumonitis             |
|                            | ARDS                    |
|                            | Pulmonary embolism (PE) |
|                            | Pleural effusion        |
|                            | Empyema                 |

|                                 | Other, None, Unknown (?)                     |
|---------------------------------|--|
| Cardiovascular complications*   | Hypotension                                  |
|                                 | Myocardial Infarction                        |
|                                 | Other cardiac ischemia                       |
|                                 | Atrial fibrillation                          |
|                                 | Ventricular fibrillation                     |
|                                 | Other cardiac arrhythmia                     |
|                                 | Cardiomyopathy                               |
|                                 | Congestive heart failure (CHF)               |
|                                 | Pulmonary embolism (PE)                      |
|                                 | Deep venous thrombosis (DVT)                 |
|                                 | Superficial venous thrombosis (SVT)          |
|                                 | Cerebrovascular accident (CVA; stroke)       |
|                                 | Thrombosis, NOS                              |
|                                 | Other, None, Unknown (?)                     |
| Gastrointestinal complications* | Acute hepatic injury                         |
|                                 | Ascites                                      |
|                                 | Bowel obstruction                            |
|                                 | Bowel perforation                            |
|                                 | Ileus  |
|                                 | Peritonitis                                  |
|                                 | Other, None, Unknown (?)                     |
| Other complications*            | Acute Kidney injury (e.g. dialysis-the later |
|                                 | my addition)                                 |
|                                 | Seizures                                     |
|                                 | Gangrene                                     |
|                                 | Other, None, Unknown (?)                     |
| Supplemental O2 required (Y/N)* |  |
| Coinfections (Y/N)*             |  |
| Blood transfusion (Y/N)*        |  |

\*Included in CCC19 data entry, no free text

SAP Finalization Date: 12/08/2020 SAP Revision Date: 07/15/2021

### eAppendix 4. Statistical Methods

### Regression models

Adjusted odds ratios (ORs) for COVID-19 severity were estimated from multivariable ordinal logistic regression models.<sup>2</sup> Because the ordinal outcome was assessed over patients' total follow-up period, the model included an offset for (log) follow-up time. Adjusted ORs and relative risks (RRs) for 30-day mortality were estimated from logistic and modified Poisson regression models, respectively.<sup>3</sup> In addition to models minimally adjusted for age and sex, we included all pre-specified covariates in fully adjusted models, given a sufficient number of events (and corresponding degrees of freedom) to enable full multivariable models. Coefficients and standard errors from models with different levels of adjustment, variance inflation factors, and clinical judgement were used to assess model stability. Exploratory analyses with smoothing splines were used to determine the association of age (as a continuous variable) with outcomes, which appeared non-linear (eFigure 2). Linear and guadratic terms for age (centered at 40 years) provided an adequate fit. All other covariates were categorical and were adjusted for using indicator variables for each category other than the reference category. These specifications reflected the assumed functional form for covariates. Note that these unweighted models quantified conditional differences in outcomes between non-Hispanic Black and non-Hispanic White patients, conditional on covariate values.

Upon revision, we performed analyses based on inverse probability of treatment weighted (IPTW) methods.<sup>4</sup> While some authors advocate for the use of methods based on causal inference to assess disparities,<sup>5</sup> others do not recommend these methods when the exposure of interest is intrinsic and not modifiable, which therefore does not allow a meaningful definition for counterfactual outcomes.<sup>6</sup> Because race as recorded in medical records and utilized in this analysis is a social and political construct, it is in theory a modifiable risk factor.<sup>7</sup>

First, we estimated propensity scores from a logistic regression model for which the outcome was a binary indicator of non-Hispanic Black versus non-Hispanic White race and the minimum sufficient adjustment set of covariates<sup>5</sup> including age, sex, region of patient residence, smoking status, obesity, cardiovascular and pulmonary comorbidities, renal disease, diabetes mellitus, type of malignancy, ECOG performance status, cancer status, timing and modality of anti-cancer therapy, and month of COVID-19 diagnosis, region of patient's residence, and calendar time, and without (primary) and with (sensitivity) insurance (with missing or unknown included as an "unknown" category). For each patient, a weight was calculated equal to the reciprocal of the probability of "receiving the treatment" (that is, race) that was "actually received," which was estimated from the propensity score model.

Next, we used graphics and summary statistics to evaluate the propensity score model.<sup>8</sup> The empirical distributions of the propensity scores stratified by race were plotted, to evaluate their overlap between groups. Mean propensity scores were calculated stratified by race across quintiles of the propensity scores in the overall cohort, to evaluate balance in the propensity scores between groups. Unweighted and weighted absolute standardized mean differences for demographic and clinical characteristics at COVID-19 diagnosis between non-Hispanic Black and non-Hispanic White patients were calculated, to evaluate whether the two groups were balanced on their observed characteristics; an absolute standardized mean difference <0.1 indicated balance.

Finally, we estimated IPTW differences in COVID-19 severity between non-Hispanic Black and non-Hispanic White patients from an ordinal logistic regression model that included an offset for (log) follow-up time. Between-group IPTW differences in 30-day all-cause mortality were estimated from both a logistic regression model (to estimate odds ratios) and a modified Poisson regression model (to estimate relative risks).<sup>3</sup> All models included race as the sole covariate, were weighted by the reciprocal of the probability of "receiving the treatment" (that is, race) that was "actually received," and used a robust (a.k.a. sandwich) variance estimator to account for the uncertainty due to estimation of the weights (and for the modified Poisson model, to account for misspecification of the variance structure). Results were reported as odds ratios (or relative risks) with 95% confidence intervals. Note that these weighted models quantified marginal differences in outcomes between non-Hispanic Black and non-Hispanic White patients.

# Proportional odds assumption

We evaluated the proportional odds assumption by fitting a set of univariable logistic regression models for all possible cut points of the ordinal COVID-19 severity outcome, with:

| Death from any cause:               | 4 |
|-------------------------------------|---|
| Received mechanical ventilation:    | 3 |
| Admitted to an intensive care unit: | 2 |
| Admitted to the hospital:           | 1 |
| No complications:                   | 0 |

That is, for each covariate, we fit a univariable logistic regression model with an offset for (log) follow-up time for each of the four binary outcomes of 4 versus <4,  $\geq$ 3 versus <3,  $\geq$ 2 versus <2, and  $\geq$ 1 versus 0.<sup>9</sup> From each logistic regression model, we obtained the estimated logits (i.e., the log odds of the outcome) for all levels of the covariate. The estimated logits obtained from the 4 versus <4,  $\geq$ 3 versus <3, and  $\geq$ 2 versus <2 models were compared to those obtained from the  $\geq$ 1 versus 0 model via subtraction, plotted, and visually inspected. If the proportional odds assumption was satisfied, then these logit differences would be similar (that is, "proportional") across all covariate levels. There did not appear to be systematic violations of the proportional odds assumption (eFigure 3), including for race; there was a suggestion that the assumption might not be satisfied for Eastern Cooperative Oncology Group (ECOG) performance status.

# Missing data

Missing or unknown data for prognostic factors and other covariates could arise due to respondent non-response for optional survey questions or a response of unknown; an unknown category was provided for all survey questions. Therefore, we assumed that any missing or unknown data were, at worst, missing at random (i.e., missingness depends on observed data only); these missing or unknown data were imputed as described below. However, unknown ECOG performance status and cancer status could be due to the lack of reassessment after initiating new anti-cancer therapy, mixed findings on scans, and lack of surveillance, among other reasons. Therefore, unknown status could be related to unobserved data (that is, missing not at random), and not appropriate to impute. Instead, unknown ECOG performance status and unknown cancer status were included as "unknown" categories.

Multiple imputation using additive regression, bootstrapping, and predictive mean matching was used to impute missing and unknown data.<sup>10</sup> The imputation model included separate binary

variables for components of the ordinal COVID-19 severity outcome (hospital admission, intensive care unit admission, receipt of mechanical ventilation, all-cause mortality), race and ethnicity, other prognostic factors (age, sex, region of patient residence, smoking status, obesity, cardiovascular and pulmonary comorbidities, renal disease, diabetes mellitus, type of malignancy, Eastern Cooperative Oncology Group (ECOG) performance status, cancer status, timing and modality of anti-cancer therapy, and month of COVID-19 diagnosis), and anti-COVID-19 treatments (remdesivir, hydroxychloroquine, corticosteroids, and other). Because rates of missingness were <5% for all variables considered (Table 1), we generated 10 imputed datasets. Results were combined across these imputed datasets using Rubin's rules. Imputation was performed on the full dataset prior to applying exclusion criteria (n=4,965).

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<sup>a</sup> Non-analytic records are case reports that did not meet data-quality metrics (eTable 1).

eFigure 2. COVID-19 Severity by Age<sup>a</sup>



<sup>a</sup> Levels of ordinal COVID-19 severity are: 0, none of the following complications; 1, admitted to the hospital; 2, admitted to an intensive care unit; 3, received mechanical ventilation; 4, died from any cause. Points are jittered vertically to enhance legibility. Red lines and shaded regions represent LOESS smoothers and 95% confidence bands, respectively.

# **eFigure 3.** Differences in Outcome Log Odds Between Univariable Logistic Regression Models for All Possible Cutpoints of the Ordinal COVID-19 Severity Outcome, Relative to the $\geq$ 1 Versus 0 Comparison<sup>a</sup>



ECOG, Eastern Cooperative Oncology Group.

<sup>a</sup> Levels of ordinal COVID-19 severity are: 0, none of the following complications; 1, admitted to the hospital; 2, admitted to an intensive care unit; 3, received mechanical ventilation; 4, died from any cause.

### eFigure 4. Distribution and Summary Statistics for Propensity Scores<sup>a</sup>



|                         | Mean prop     | ensity score   |
|-------------------------|---------------|----------------|
| Percentile <sup>a</sup> | Black         | White          |
| 0-<20 <sup>th</sup>     | 0.1139        | 0.1025         |
| 20-<40 <sup>th</sup>    | 0.2181        | 0.2117         |
| 40-<60 <sup>th</sup>    | 0.2969        | 0.2938         |
| 60-<80 <sup>th</sup>    | 0.3841        | 0.3817         |
| 80–100 <sup>th</sup>    | 0.5396        | 0.5177         |
| a Dorooptiloo (         | i o quintiloc | ) of proposity |

<sup>a</sup> Percentiles (i.e., quintiles) of propensity scores in the total cohort.

<sup>a</sup> Propensity scores were estimated from a logistic regression model for race that included age, sex, region of patient residence, smoking status, obesity, cardiovascular and pulmonary comorbidities, renal disease, diabetes mellitus, type of malignancy, Eastern Cooperative Oncology Group performance status, cancer status, timing and modality of anti-cancer therapy, and month of COVID-19 diagnosis.

#### □ Non-Hispanic Black □ Non-Hispanic White

**eFigure 5.** Absolute Standardized Mean Differences for Demographic and Clinical Characteristics at COVID-19 Diagnosis Between Non-Hispanic Black and Non-Hispanic White Patients<sup>a</sup>



ECOG, Eastern Cooperative Oncology Group.

<sup>a</sup> Weighted absolute standardized mean differences were weighted by the reciprocal of the probability of "receiving the treatment" (that is, race) that was "actually received," which was estimated from a propensity score model for race that included age, sex, region of patient residence, smoking status, obesity, cardiovascular and pulmonary comorbidities, renal disease, diabetes mellitus, type of malignancy, Eastern Cooperative Oncology Group performance status, cancer status, timing and modality of anti-cancer therapy, and month of COVID-19 diagnosis, and without (primary) and with (sensitivity) insurance.

# eTable 1. Metrics for Data Quality

| Quality score | Definition   |
|---------------|--|
| 0             | No problems identified   |
| 1             | 1 minor problem  |
| 2             | 2 minor problems   |
| 3             | 3 minor problems or 1 moderate problem   |
| 4             | 4 minor problems or 1 moderate problem and 1 minor problem                     |
| 5             | 5 minor problems or 1 moderate problem and 2 minor problems or 1 major problem |
| ≥6            | As above with additional problems  |

Minor problems were valued at 1 point, moderate problems at 3 points, and major problems at 5 points. Reports with a quality score of >4 were excluded from the analysis.

# eTable 2. Patients on Multimodal Anticancer Therapy

|               | Targeted | Endocrine | Immunotherapy | Local | Other |
|---------------|----------|-----------|---------------|-------|-------|
| Cyto          | 812      | 812       | 613           | 686   | 527   |
| Targeted      | -        | 741       | 590           | 713   | 465   |
| Endocrine     | -        | -         | 502           | 591   | 368   |
| Immunotherapy | -        | -         | -             | 452   | 188   |
| Local         | -        | -         | -             | -     | 330   |

| Total      | Non-Hispanic Black  | Non-Hispanic White  |
|------------|---|---|
| (N = 3506) | (N = 1068)  | (N = 2438)  |
|            |   |   |
| 707 (20)   | 232 (22)  | 475 (19)  |
| 593 (17)   | 211 (20)  | 382 (16)  |
| 436 (12)   | 137 (13)  | 299 (12)  |
| 293 (8)    | 72 (7)  | 221 (9)   |
| 287 (8)    | 85 (8)  | 202 (8)   |
| 223 (6)    | 66 (6)  | 157 (6)   |
| 168 (5)    | 50 (5)  | 118 (5)   |
| 153 (4)    | 6 (1)   | 147 (6)   |
| 103 (3)    | 28 (3)  | 75 (3)  |
| 79 (2)     | 17 (2)  | 62 (3)  |
| 51 (1)     | 8 (1)   | 43 (2)  |
| 38 (1)     | 17 (2)  | 21 (1)  |
|            |   |   |
| 428 (12)   | 106 (10)  | 322 (13)  |
| 166 (5)    | 81 (8)  | 85 (3)  |
| 152 (4)    | 43 (4)  | 109 (4)   |
| 11 (<1)    | 3 (<1)  | 8 (<1)  |
|            | Total $(N = 3506)$ 707 (20)         593 (17)         436 (12)         293 (8)         287 (8)         223 (6)         168 (5)         153 (4)         103 (3)         79 (2)         51 (1)         38 (1)         428 (12)         166 (5)         152 (4)         11 (<1) | TotalNon-Hispanic Black $(N = 3506)$ $(N = 1068)$ 707 (20)232 (22)593 (17)211 (20)436 (12)137 (13)293 (8)72 (7)287 (8)85 (8)223 (6)66 (6)168 (5)50 (5)153 (4)6 (1)103 (3)28 (3)79 (2)17 (2)51 (1)8 (1)38 (1)17 (2)428 (12)106 (10)166 (5)81 (8)152 (4)43 (4)11 (<1) |

# eTable 3. Type of Malignancy

Data presented as n (%). Categories are not mutually exclusive.

### eTable 4. Laboratory Measurements Among Hospitalized Patients<sup>a</sup>

|                                  | Total      | Non-Hispanic Black | Non-Hispanic White |
|----------------------------------|------------|--------------------|--------------------|
|                                  | (N = 3506) | (N = 1068)         | (N = 2438)         |
| Absolute lymphocyte count, n (%) |            |                    |                    |
| Low <sup>b</sup>                 | 973 (48)   | 329 (47)           | 644 (48)           |
| Normal                           | 626 (31)   | 248 (36)           | 378 (28)           |
| High <sup>c</sup>                | 56 (3)     | 12 (2)             | 44 (3)             |
| Missing/unknown                  | 371 (18)   | 107 (15)           | 264 (20)           |
| Absolute neutrophil count, n (%) |            |                    |                    |
| Low <sup>c</sup>                 | 145 (7)    | 52 (7)             | 93 (7)             |
| Normal                           | 1207 (60)  | 449 (65)           | 758 (57)           |
| High <sup>c</sup>                | 346 (17)   | 101 (15)           | 245 (18)           |
| Missing/unknown                  | 328 (16)   | 94 (14)            | 234 (18)           |
| Platelet count, n (%)            |            |                    |                    |
| Low <sup>c</sup>                 | 530 (26)   | 160 (23)           | 370 (28)           |
| Normal                           | 1177 (58)  | 438 (63)           | 739 (56)           |
| High <sup>c</sup>                | 83 (4)     | 26 (4)             | 57 (4)             |
| Missing/unknown                  | 236 (12)   | 72 (10)            | 164 (12)           |
| Creatinine, n (%)                |            |                    |                    |
| Normal                           | 1016 (50)  | 292 (42)           | 724 (54)           |
| Abnormal <sup>c</sup>            | 789 (39)   | 339 (49)           | 450 (34)           |
| Missing/unknown                  | 221 (11)   | 65 (9)             | 156 (12)           |
| D-dimer, n (%)                   |            |                    |                    |
| Normal                           | 159 (8)    | 48 (7)             | 111 (8)            |
| Abnormal <sup>c</sup>            | 954 (47)   | 364 (52)           | 590 (44)           |
| Missing/unknown                  | 913 (45)   | 284 (41)           | 629 (47)           |
| Troponin, n (%)                  |            |                    |                    |
| Normal                           | 693 (34)   | 252 (36)           | 441 (33)           |
| Abnormal <sup>c</sup>            | 474 (23)   | 151 (22)           | 323 (24)           |

| Missing/unknown              | 859 (42)  | 293 (42) | 566 (43) |
|------------------------------|-----------|----------|----------|
| Lactate dehydrogenase, n (%) |           |          |          |
| Normal                       | 244 (12)  | 74 (11)  | 170 (13) |
| Abnormal <sup>c</sup>        | 808 (40)  | 316 (45) | 492 (37) |
| Missing/unknown              | 974 (48)  | 306 (44) | 668 (50) |
| C-reactive protein, n (%)    |           |          |          |
| Normal                       | 83 (4)    | 30 (4)   | 53 (4)   |
| Abnormal <sup>c</sup>        | 1041 (51) | 390 (56) | 651 (49) |
| Missing/unknown              | 902 (45)  | 276 (40) | 626 (47) |

Data presented as n (%).

<sup>a</sup> Respondents were instructed to report the earliest measured laboratory measurements during COVID-19 course. Except for low absolute lymphocyte count (ALC), which was centrally defined as ALC < 1500/µL, ascertainment of upper and lower limits of normal was left to the discretion of respondents. Laboratory measurements were summarized among hospitalized patients only due to common clinical practice to avoid a laboratory blood draw for outpatients.

<sup>b</sup> Low absolute lymphocyte count is defined as less than 1500/uL.

<sup>c</sup> As defined by the reporting institution's normal laboratory value ranges.

|                                       | Total          |           | Non-Hispanic Black |                    | Non-Hispanic White |                    |
|---------------------------------------|----------------|-----------|--------------------|--------------------|--------------------|--------------------|
|                                       | N <sup>a</sup> | n (%)     | N <sup>a</sup>     | n <sup>b</sup> (%) | N <sup>a</sup>     | n <sup>b</sup> (%) |
| Cardiovascular complications          |                |           |                    |                    |                    |                    |
| Hypotension                           | 3373           | 401 (12)  | 1026               | 151 (15)           | 2347               | 250 (11)           |
| Myocardial infarction                 | 3365           | 47 (1)    | 1024               | 20 (2)             | 2341               | 27 (1)             |
| Other cardiac ischemia                | 3365           | 31 (1)    | 1024               | 7 (1)              | 2341               | 24 (1)             |
| Atrial fibrillation                   | 3372           | 206 (6)   | 1025               | 50 (5)             | 2347               | 156 (7)            |
| Ventricular fibrillation              | 3364           | 14 (<1)   | 1024               | 7 (1)              | 2340               | 7 (<1)             |
| Other cardiac arrhythmia              | 3366           | 90 (3)    | 1026               | 34 (3)             | 2340               | 56 (2)             |
| Cardiomyopathy                        | 3365           | 24 (1)    | 1024               | 7 (1)              | 2341               | 17 (1)             |
| Congestive heart failure              | 3365           | 113 (3)   | 1024               | 35 (3)             | 2341               | 78 (3)             |
| Pulmonary embolism                    | 3440           | 81 (2)    | 1048               | 30 (3)             | 2392               | 51 (2)             |
| Deep venous thrombosis                | 3365           | 68 (2)    | 1024               | 25 (2)             | 2341               | 43 (2)             |
| Superficial venous thrombosis         | 3365           | 11 (<1)   | 1024               | <5 (<1)            | 2341               | 7 (<1)             |
| Cerebrovascular accident              | 3365           | 40 (1)    | 1024               | 18 (2)             | 2341               | 22 (1)             |
| Thrombosis, NOS                       | 3395           | 24 (1)    | 1034               | 7 (1)              | 2361               | 17 (1)             |
| Pulmonary complications               |                |           |                    |                    |                    |                    |
| Respiratory failure                   | 3438           | 1002 (29) | 1051               | 357 (34)           | 2387               | 645 (27)           |
| Pneumonitis or pneumonia <sup>c</sup> | 3420           | 440 (13)  | 1044               | 127 (12)           | 2376               | 313 (13)           |
| ARDS                                  | 3425           | 275 (8)   | 1046               | 116 (11)           | 2379               | 159 (7)            |
| Pulmonary embolism                    | 3440           | 81 (2)    | 1048               | 30 (3)             | 2392               | 51 (2)             |
| Pleural effusion                      | 3422           | 135 (4)   | 1043               | 45 (4)             | 2379               | 90 (4)             |
| Empyema                               | 3418           | 8 (<1)    | 1043               | <5 (<1)            | 2375               | 5 (<1)             |
| Gastrointestinal complications        |                |           |                    |                    |                    |                    |
| Acute hepatic injury                  | 3334           | 84 (3)    | 1015               | 36 (4)             | 2319               | 48 (2)             |
| Ascites                               | 3334           | 18 (1)    | 1016               | 8 (1)              | 2318               | 10 (<1)            |
| Bowel obstruction                     | 3333           | 15 (<1)   | 1015               | 5 (<1)             | 2318               | 10 (<1)            |
| Bowel perforation                     | 3332           | 6 (<1)    | 1015               | 0 (0)              | 2317               | 6 (<1)             |
| lleus                                 | 3332           | 12 (<1)   | 1015               | 6 (1)              | 2317               | 6 (<1)             |
| Peritonitis                           | 3332           | 6 (<1)    | 1015               | <5 (<1)            | 2317               | 5 (<1)             |

| eTable 5. Rates of Cardiovascular, Pulmonary, and Gastrointe | estinal Complications (N = 3506) |
|--|----------------------------------|
|--|----------------------------------|

ARDS, Acute respiratory distress syndrome; NOS, not otherwise specified.

<sup>a</sup> Number of patients with non-missing data.
 <sup>b</sup> Groups with fewer than 5 patients were masked (i.e., <5) to minimize the risk of re-identification as per CCC19 policy.</li>

<sup>c</sup> These are collected as separate complications but given the difficulty in radiographically distinguishing pneumonia from pneumonitis, they are combined here.

| COVI          | D-19 severity | 30-day mortality |               |               |               |
|---------------|---------------|------------------|---------------|---------------|---------------|
| Odds<br>Ratio | CI            | Odds Ratio       | CI            | Relative Risk | СІ            |
| 1.13          | (1.02 - 1.24) | 1.18             | (0.95 - 1.47) | 1.16          | (0.96 - 1.39) |

# eTable 6. Inverse Probability Treatment Weighting (IPTW) With Insurance Added