

Reporting Summary

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Statistics

For all statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.

n/a Confirmed

- The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement
- A statement on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly
- The statistical test(s) used AND whether they are one- or two-sided
Only common tests should be described solely by name; describe more complex techniques in the Methods section.
- A description of all covariates tested
- A description of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons
- A full description of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) AND variation (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)
- For null hypothesis testing, the test statistic (e.g. F , t , r) with confidence intervals, effect sizes, degrees of freedom and P value noted
Give P values as exact values whenever suitable.
- For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings
- For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes
- Estimates of effect sizes (e.g. Cohen's d , Pearson's r), indicating how they were calculated

Our web collection on [statistics for biologists](#) contains articles on many of the points above.

Software and code

Policy information about [availability of computer code](#)

Data collection

No software used for data collection

Data analysis

All statistical analyses were performed in R (<https://www.R-project.org/>), version 3.6. R libraries included Hmisc (version 4.7), glmnet (v3.0-2), randomForestSRC (v 2.9.3). Code is deposited in a public repository (EGAS00001006703) and the link is in the Code Availability statement of the manuscript.

For manuscripts utilizing custom algorithms or software that are central to the research but not yet described in published literature, software must be made available to editors and reviewers. We strongly encourage code deposition in a community repository (e.g. GitHub). See the Nature Portfolio [guidelines for submitting code & software](#) for further information.

Data

Policy information about [availability of data](#)

All manuscripts must include a [data availability statement](#). This statement should provide the following information, where applicable:

- Accession codes, unique identifiers, or web links for publicly available datasets
- A description of any restrictions on data availability
- For clinical datasets or third party data, please ensure that the statement adheres to our [policy](#)

Public datasets that were used during data processing included ExAC database (<http://exac.broadinstitute.org/>) and IDs from COSMIC database (<https://cancer.sanger.ac.uk/cosmic>).

The data and code required to reproduce results are deposited to the European Genome-Phenome Archive under accession number EGAS00001006703, and can be made available upon request. Qualified researchers may request access to individual patient-level data through the clinical study data request platform (<https://vivli.org/>). Further details on Roche's criteria for eligible studies are available at <https://vivli.org/members/ourmembers>. For further details on Roche's Global Policy on the Sharing of Clinical Information and how to request access to related clinical study documents, see https://www.roche.com/research_and_development/who_we_are_how_we_work/clinical_trials/our_commitment_to_data_sharing.htm.

Human research participants

Policy information about [studies involving human research participants and Sex and Gender in Research](#).

Reporting on sex and gender

Gender was not collected in this study, and accordingly we do not use this term in the paper. Please refer to Extended Data Tables 1 and 2 for covariate-relevant population characteristics of the human research participants in the IMpower150 ctDNA study, which includes Sex.

Population characteristics

Patient metadata are included in the manuscript, including tables of population characteristics for both the full IMpower150 population and this retrospective exploratory ctDNA substudy reported in this manuscript (Supp Table ST1). Briefly, IMpower150 patients had stage IV or recurrent metastatic nonsquamous NSCLC for which they had not previously received chemotherapy, a baseline Eastern Cooperative Oncology Group (ECOG) performance-status score of 0 or 1, and tumor tissue available for biomarker testing and if they were eligible to receive bevacizumab; patients with any PD-L1 immunohistochemistry status were eligible. Please refer to Extended Data Tables 1 and 2 for covariate-relevant population characteristics of the human research participants in the IMpower150 ctDNA study, including baseline ECOG score, Age, Sex, Tobacco use, Race, Region, Number of metastatic sites, and PDL1 status.

Recruitment

In the IMpower150 study patients were eligible for inclusion in this retrospective exploratory ctDNA substudy reported in this manuscript if they had plasma samples available for ctDNA testing at both baseline and an early on-treatment timepoint (either cycle 2 day 1 or cycle 3 day 1), as well as PBMCs available in order to perform germline subtraction. This reduced the number of patients in this ctDNA substudy to 466. We expected a survivorship bias in this ctDNA-evaluable population due to our requirement for patients to have samples available after randomization, and while no strong PFS bias was found for ctDNA evaluable versus non-evaluable (HR=0.92 [0.82-1.05]), we did detect a survivorship bias for OS (HR=0.86 [0.75 – 0.99]) (Extended Data Figure ED1a). However, baseline characteristics were similar between the full IMpower150 and ctDNA evaluable population, including baseline ECOG, age, sex, race, region, among others (Supplementary Table ST1). We do not expect this OS survivorship to strongly impact our results, because this exploratory ctDNA substudy is concerned with building ctDNA-based models to predict overall survival, and typically patients who do not have evaluable on-treatment samples available are those who unfortunately had very rapid disease progression, and therefore for which a ctDNA model to predict survival is of limited utility.

Ethics oversight

The study was conducted in accordance with the International Conference on Harmonisation Good Clinical Practice guidelines and with the principles of the Declaration of Helsinki. All patients provided written informed consent, and the protocol was approved by independent ethics committees at each participating site. The study protocol is in the appendix of the primary clinical manuscript for IMpower150 (Socinski et al, NEJM 2018) and can also be found at <https://clinicaltrials.gov/ct2/show/NCT02366143>.

There are 161 institutes/organizations whose ethical committees approved the protocol, including:

- [1] "Copernicus Group Independent Review Board"
- [2] "Western Institutional Review Board"
- [3] "Melbourne Health Human Research Ethics Committee"
- [4] "Tasmania Health and Medical Human Research Ethics Committee"
- [5] "Asociacion Benefica Prisma"
- [6] "Comité Institucional de ética en Investigación Instituto Nacional de Enfermedades Neoplásicas"
- [7] "Singhealth Centralised Institutional Review Board"
- [8] "Institutional Review Board Kaohsiung Medical University Chung-Ho Memorial Hospital"
- [9] "Institutional Review Board Taipei Veterans General Hospital"
- [10] "Institutional Review Board"
- [11] "Institutional Review Board of the Chi Mei Medical Center"
- [12] "The Institutional Review Board of Taichung Veterans General Hospital"
- [13] "Institutional Review Board of Tri-Service General Hospital"
- [14] "Cheng-Hsin General Hospital Institutional Review Board"
- [15] "Consejo de Evaluación Ética de Investigación en Salud - CoEIS"
- [16] "Ministerio de Salud - Provincia de Río Negro"
- [17] "Comité Independiente de Ética en investigación clínica \"Dr. Carlos A. Barclay"
- [18] "Comité de ética en investigación Fundación Oncosalud"
- [19] "Comisión Conjunta de Investigación en Salud (CCIS)"
- [20] "Comité de Ética Centro de Oncología e Investigación Buenos Aires"
- [21] "Comité de Ética Independiente - Fundación Sanatorio"
- [22] "Institutional Review Board of Chang Gung Medical Foundation"
- [23] "Ethikkommission für das Bundesland Salzburg"
- [24] "The Ethics Committee for Clinical Trials of Medicinal Products"
- [25] "Chesapeake Institutional Review Board"
- [26] "St. Luke's Hospital & Health Network IRB"
- [27] "St Charles Medical Center"
- [28] "Saint Luke's Hospital Institutional Review Board"
- [29] "Kantonale Ethikkommission Bern (KEK)"
- [30] "Comite de Etica Cientifico del Servicio de Salud Metropolitano Norte"
- [31] "Comitato Etico Area Vasta Nord Ovest presso Azienda Ospedaliero Universitaria Pisana di Pisa"

- [32] "Comitato Etico Cardarelli-Santobono"
- [33] "Comitato Etico San Martino IST 2"
- [34] "Comitato Etico Lazio 1"
- [35] "COMITATO ETICO DELL'UNIVERSITA' CAMPUS BIO-MEDICO DI ROMA"
- [36] "Medical Research Ethics Committees United"
- [37] "CPP Sud-Méditerranée 2"
- [38] "Sir Charles Gairdner Hospital HREC"
- [39] "Bellberry Human Research Ethics Committee"
- [40] "Cabrini Human Research Ethics Committee"
- [41] "Comitato Etico Catania 1 presso A.O. Universitaria Policlinico Vittorio Emanuele di Catania"
- [42] "Comissão de Ética para a Investigação Clínica - CEIC"
- [43] "Ethikkommission an der Universität Regensburg"
- [44] "Ethikkommission an der Universität Regensburg"
- [45] "Lakeridge Health REB"
- [46] "St. Joseph Mercy Health System Institutional Review Board #2 - Oncology Central IRB"
- [47] "Mercy Saint Vincent Medical Center Institutional Review Board"
- [48] "US Oncology Inc. Institutional Review Board"
- [49] "Western Institutional Review Board (WIRB)"
- [50] "CEIC de la Corporacion Sanitaria del Parc Tauli"
- [51] "CEIC de Cantabria"
- [52] "CEIC de Andalucia (CCEIBA)"
- [53] "CEIC Hospital Universitario La Paz"
- [54] "CEIC Fundación Jiménez Díaz"
- [55] "CEIC Hospital Clínico Universitario de Valencia"
- [56] "CEIC Hospital General Universitario Gregorio Marañón"
- [57] "CEIC Hospital Clinico San Carlos"
- [58] "CEIC Grupo Hospital de Madrid"
- [59] "CEIC Parc de Salut Mar"
- [60] "CEIC Islas Baleares (CEIC-IB)"
- [61] "CEIC de Galicia (CAEI)"
- [62] "CEIC Hospital Clinic de Barcelona"
- [63] "Kaiser Permanente Southern California Institutional Review Board"
- [64] "Research Ethics Committee of National Taiwan University Hospital"
- [65] "Institutional Review Board of Chung Shan Medical University Hospital"
- [66] "CHU de Liège - Comité d'Ethique"
- [67] "CEQ of MI Kryvyi Rih Oncology Dispensary of Dnipropetrovsk Regional Council"
- [68] "CEQ of Treatment and Prevention Institution Volyn Regional Oncology Dispensary"
- [69] "CEQ of MI Dnipropetrovsk City Multifield Clinical Hospital #4 of Dnipropetrovsk Regional Council"
- [70] "CEQ of Poltava Regional Clinical Oncology Dispensary of Poltava Regional Council"
- [71] "Commission on Ethics Questions of Uzhgorod Central City Clinical Hospital"
- [72] "Commission of Ethics Questions on the basis of the Chernivtsi Regional Clinical Oncology Dispensary"
- [73] "CEQ of MI of Zaporizhzhia Regional Council Zaporizhzhia Regional Clinical Oncology Dispensary"
- [74] "CEQ of Transcarpathian Regional Clinical Oncology Dispensary"
- [75] "CEQ of SI Institute of Medical Radiology n.a. S.P. Hryhoriev of NAMS of Ukraine"
- [76] "CEQ of Municipal Noncommercial Institution Regional Center of Oncology"
- [77] "CEQ of Regional Municipal Institution Sumy Regional Clinical Oncology Dispensary"
- [78] "Comitato Etico Azienda Ospedaliera Universitaria Maggiore della Carità"
- [79] "Comitato Etico Dell Universita Cattolica del Sacro Cuore Policlinico Universitario Agostino Gemelli"
- [80] "Comité de Ética em Pesquisa em Seres Humanos da Faculdade de Medicina de São José do Rio Preto"
- [81] "Comitê de Ética em Pesquisa em Seres Humanos do Hospital Socor"
- [82] "Comitê de Ética em Pesquisa em Seres Humanos da Universidade de Ribeirão Preto (UNAERP)"
- [83] "Comitê de Ética em Pesquisa em Seres Humanos da Irmandade da Santa Casa de Londrina"
- [84] "Comitê de Ética em Pesquisa em Seres Humanos da Liga Norte Riograndense Contra o Câncer"
- [85] "Comitê de Ética em Pesquisa Fundação Pio XII Hospital de Câncer de Barretos"
- [86] "Comité de Ética en Investigación de la Facultad de Medicina y Hospital Universitario"
- [87] "Missouri Baptist Medical Center Institutional Review Board"
- [88] "University of California Irvine Institutional Review Board"
- [89] "University of California"
- [90] "Frederick Memorial Hospital Institutional Review Board"
- [91] "Mercy Medical Center IRB"
- [92] "University of Chicago Hospitals Institutional Review Board"
- [93] "Mount Sinai Medical Center IRB"
- [94] "Comité Provincial de Bioética - Ministerio de Salud de la Provincia de Santa Fé"
- [95] "Comite de Etica Investigacion de la Clinica Bajío"
- [96] "Comite de Etica en Investigacion de Mexico Centre for Clinical Research SA de CV"
- [97] "Comitê de Ética em Pesquisa da Universidade de Caxias do Sul"
- [98] "Comitê de Ética em Pesquisa do Hospital de Clínicas de Porto Alegre"
- [99] "Comite de Etica em Pesquisa da Universidade Federal de Sao Paulo - Hospital Sao Paulo"
- [100] "Comitê de Ética em Pesquisa - Hospital Mãe de Deus"
- [101] "Comitê de Ética em Pesquisa da Fundação Antônio Prudente – Hospital do Câncer A. C. Camargo"
- [102] "Ethics Committee for Multi-Centre Trials"
- [103] "Ethics Committee at Clinical Oncology Dispensary"
- [104] "Ethics Committee at City Clinical oncologic dispensary"
- [105] "Ethics Committee at Russian Oncology Research Center n.a. N.N.Blokhin"
- [106] "Ethics Committee at Moscow City Oncology Hospital #62 of Moscow Healthcare Department"
- [107] "Ethics Committee at Volzhskiy regional clinical oncology dispensary #3"

- [108] "CEQ of Ivano-Frankivsk Regional Oncology Dispensary"
- [109] "CEIC Hospital Universitario Insular Materno-Infantil de Las Palmas"
- [110] "CEIC Hospital Universitari Vall d'Hebron"
- [111] "CEIC Hospital Universitario Ramon y Cajal"
- [112] "CEIC Hospital Universitario 12 de Octubre"
- [113] "Eticka komisia Presovskeho samospravného kraja"
- [114] "Eticka komisia Univerzity nemocnica Bratislava"
- [115] "Lithuanian Bioethics Committee"
- [116] "CEIC Hospital Universitari de Bellvitge"
- [117] "Eticka komisia pri Narodnom onkologickom ustave"
- [118] "WIRB Copernicus Group"
- [119] "Kaiser Permanente of Colorado Institutional Review Board"
- [120] "Ethics Committee at Russian Medical Military Academy n.a. S.M.Kirov"
- [121] "Comite de Etica en Investigacion del Instituto Regional de Enfermedades Neoplasicas"
- [122] "National Cheng Kung University Hospital Human Experiment and Ethic Committee"
- [123] "Houston Methodist Research Institute IRB"
- [124] "Ingalls Memorial Hospital IRB"
- [125] "Rush University Medical Center Institutional Review Board"
- [126] "Mercy Health Springfield Communities Institutional Review Board"
- [127] "Mayo Clinic Institutional Review Board"
- [128] "Salus IRB"
- [129] "Mackay Memorial Hospital Institutional Review Board"
- [130] "Commission on Ethics Questions of Vinnytsya Regional Clinical Oncology Dispensary"
- [131] "Comite Etico Cientifico Clinica Santa Maria"
- [132] "BRANY IRB"
- [133] "Yale University Human Research Protection Program"
- [134] "Maimonides Med Ctr Institutional Review Board"
- [135] "Scripps Health Institutional Review Board"
- [136] "University of Texas Health Science Center San Antonio Institutional Review Board"
- [137] "Park Nicollet Institute Institutional Review Board"
- [138] "Eticka komisia NsP Sv. Jakuba"
- [139] "Ethikkommission der Bayerischen Landesärztekammer"
- [140] "CEQ of Kyiv City Clinical Oncological Center"
- [141] "The Institutional Review Board of China Medical University Hospital"
- [142] "Ethics Committee at Private Medical Institution \"Evromedservis\""
- [143] "Comitato di Bioetica dell'AUSL 1 di Sassari"
- [144] "Eticka komisia Onkologicky ustav sv. Alzbety"
- [145] "Ethics Committee at Railway Clinical Hospital JSC RZhD"
- [146] "Sault Area Hospital Research Ethics Board"
- [147] "Toranomon Hospital and Toranomon Hospital Kajigaya IRB"
- [148] "Kitasato University Sagamihara IRB"
- [149] "Niigata Cancer Center Hospital IRB"
- [150] "Kyoto University Hospital IRB"
- [151] "Osaka City University Hospital IRB"
- [152] "National Hospital Organization Toneyama National Hospital IRB"
- [153] "Wakayama Medical University IRB"
- [154] "Kurume University IRB"
- [155] "National Hospital Organization Kyushu Cancer Center"
- [156] "Kanagawa Cardiovascular and Respiratory Center IRB"
- [157] "National Hospital Organization Kyushu Medical Center IRB"
- [158] "National Hospital Organization Shikoku Cancer Center IRB"
- [159] "Miyagi Cancer Center IRB"
- [160] "Kyorin University Hospital IRB"
- [161] "Center Hospital of the National Center for Global Health and Medicine IRB"

Note that full information on the approval of the study protocol must also be provided in the manuscript.

Field-specific reporting

Please select the one below that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.

Life sciences Behavioural & social sciences Ecological, evolutionary & environmental sciences

For a reference copy of the document with all sections, see [nature.com/documents/nr-reporting-summary-flat.pdf](https://www.nature.com/documents/nr-reporting-summary-flat.pdf)

Life sciences study design

All studies must disclose on these points even when the disclosure is negative.

Sample size

Details on the IMpower150 study plan are published elsewhere. No sample size calculations were done for the ctDNA subsidy of IMpower150. The sample size was determined by the number of patients in IMpower150 (n=1201) which had baseline samples run on a 1.25 Mb ctDNA assay (n=1062), and those were subset to those patients with aliquots of C2D1 or C3D1 plasma available (n=910). 566 patients were then

randomly chosen for the ctDNA substudy, only 466 of which had PBMC available for CHIP/germline correction who were included in the final ctDNA analysis population.

Data exclusions Patients were excluded if they did not have baseline plasma and C2D1/C3D1 and PBMC available. Please refer to Figure 1 for sample flow diagram.

Replication Biomarkers measured in patient plasma samples were not replicated. Replication was not possible due to the limited patient samples available. The one experiment that had replication was during assay development in which 63 replicates were run to measure the concordance between the baseline assay and on-treatment assay. This experiment showed high concordance between the replicates ($R^2=0.9961$; Supplementary Figure 1B right). No other experiments were replicated.

Randomization Details on the original IMpower150 study plan including randomization has been published elsewhere and can also be found at <https://clinicaltrials.gov/ct2/show/NCT02366143>. For this exploratory retrospective ctDNA substudy reported in this manuscript, the 466 ctDNA-evaluable patients were split into a training cohort and a testing cohort for model development. The training and test sets were initially chosen based on sequencing batch for the set of 566 patients chosen for the ctDNA substudy (see Figure 1a), where we put sequencing batch1 in training set and then added in patients from later batches to reach the target 50%/50% split. The sequencing lab decided which samples to include in batch 1 without any knowledge about the baseline characteristics, treatment, or clinical outcomes of the patients. We then checked for imbalances and it was found that RACE was not well distributed due to all Asian patients appearing in batch1, and so we moved half of the Asian patients to the test set and replaced these spots in the training data with a random set of patients. As the analysis progressed (in the training subset of data) we decided to add in PBMC correction due to concern over germline/CHIP variants contaminating the ctDNA dataset, which reduced the number of patients to those with PBMC available for correction, giving a final n of 466 patients and a final split of 240/226 patients for train/test. The final training/test sets were well balanced in clinical features and survival outcomes as can be seen in Supplementary Table ST2 and Extended Data Figure ED1f.

Blinding Details about blinding during arm allocation for the IMpower150 study itself can be found in the IMpower150 primary clinical manuscript or at <https://clinicaltrials.gov/ct2/show/NCT02366143>. For this retrospective exploratory ctDNA substudy, our model development analysis was not initiated until after samples were allocated to either the training or testing subgroup. The allocation of training/testing was not performed blinded, as the goal was to ensure that the training and testing subgroups were similar in baseline characteristics. Sample collection was performed prior to this retrospective analysis, and so investigators were blinded during sample collection. Data was generated by a diagnostic company separate from the analysts who performed the model development, and so data generation was also performed by blinded individuals.

Reporting for specific materials, systems and methods

We require information from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, system or method listed is relevant to your study. If you are not sure if a list item applies to your research, read the appropriate section before selecting a response.

Materials & experimental systems

Methods

- n/a Involved in the study
- Antibodies
- Eukaryotic cell lines
- Palaeontology and archaeology
- Animals and other organisms
- Clinical data
- Dual use research of concern

- n/a Involved in the study
- ChIP-seq
- Flow cytometry
- MRI-based neuroimaging

Antibodies

Antibodies used PD-L1 expression on tumor cells or tumor-infiltrating immune cells was analyzed in archival or freshly collected tumor tissue (or both) with the use of a PD-L1 immunohistochemistry assay (Ventana Medical Systems; clone SP142; catalog number N/A; pre-dilute ready to use antibody product at 36ug/5mL)

Validation Ventana PD-L1 (SP142) assay validation can be found at the following link: https://www.accessdata.fda.gov/cdrh_docs/pdf16/p160002c.pdf

Clinical data

Policy information about [clinical studies](#)

All manuscripts should comply with the ICMJE [guidelines for publication of clinical research](#) and a completed [CONSORT checklist](#) must be included with all submissions.

Clinical trial registration NCT02366143

Study protocol The protocol was published with the primary clinical manuscript and can be found here: https://www.nejm.org/doi/suppl/10.1056/NEJMoa1716948/suppl_file/nejmoa1716948_protocol.pdf

Details of clinical data collection can be found in primary clinical manuscript for IMpower150. Briefly, IMpower150 data was collected between March 2015 and December 2019, patients from 26 countries or regions were enrolled at 240 sites. The list of 240 sites is the following:

United States, Arizona
 Ironwood Cancer & Research Centers
 Chandler, Arizona, United States, 85224
 Arizona Oncology Associates
 Flagstaff, Arizona, United States, 86001
 United States, California
 Southern CA Permanente Med Grp
 Bellflower, California, United States
 Marin Cancer Care Inc
 Greenbrae, California, United States, 94904
 Scripps Health
 La Jolla, California, United States, 92037
 Chao Family Comprehensive Cancer Center UCI
 Orange, California, United States, 92868
 United States, Colorado
 Rocky Mountain Cancer Center
 Denver, Colorado, United States, 80218
 Kaiser Permanente
 Lonetree, Colorado, United States, 80124
 United States, Connecticut
 Danbury Hospital
 Danbury, Connecticut, United States, 06810
 Yale Cancer Center
 New Haven, Connecticut, United States, 06520
 United States, Florida
 Holy Cross Hospital Inc
 Fort Lauderdale, Florida, United States, 33308
 Cancer Specialists of North Florida - Baptist South
 Jacksonville, Florida, United States, 32258
 Mount Sinai Medical Center
 Miami Beach, Florida, United States, 33140
 Hematology Oncology Associates of the Treasure Coast
 Port Saint Lucie, Florida, United States, 34952
 United States, Georgia
 Piedmont Cancer Institute, PC
 Atlanta, Georgia, United States, 30318
 United States, Illinois
 Rush University Medical Center
 Chicago, Illinois, United States, 60612
 Univ of Chicago
 Chicago, Illinois, United States, 60637
 Ingalls Memorial Hospital
 Harvey, Illinois, United States, 60426
 Oncology Specialists, S.C.
 Park Ridge, Illinois, United States, 60068
 United States, Iowa
 Hematology-Oncology; Associates of the Quad Cities
 Bettendorf, Iowa, United States, 52722
 United States, Kentucky
 Norton Cancer Institute
 Louisville, Kentucky, United States, 40202
 United States, Maine
 New England Cancer Specialists
 Scarborough, Maine, United States, 04074
 United States, Maryland
 Mercy Medical Center
 Baltimore, Maryland, United States, 21202
 Regional Cancer Care Associates
 Bethesda, Maryland, United States, 20817
 Maryland Oncology Hematology, P.A.
 Columbia, Maryland, United States, 21044
 United States, Michigan
 St. Joseph Mercy Health System
 Ann Arbor, Michigan, United States, 48106
 United States, Minnesota
 St. Luke's Regional Cancer Center
 Duluth, Minnesota, United States, 55805
 Park Nicolett - Frauenshuh Cancer Center
 Saint Louis Park, Minnesota, United States, 55426
 United States, Missouri
 St. Luke's Cancer Institute
 Kansas City, Missouri, United States, 64111

Missouri Baptist Medical Center
 Saint Louis, Missouri, United States, 63131
 United States, Montana
 Billings Clinic
 Billings, Montana, United States, 59102
 Montana Cancer Specialists
 Missoula, Montana, United States, 59802
 United States, Nevada
 Comprehensive Cancer Centers of Nevada
 Henderson, Nevada, United States, 89014
 United States, New Jersey
 Summit Medical Group
 Berkeley Heights, New Jersey, United States, 07922
 Valley Hospital; Oncology Research
 Paramus, New Jersey, United States, 07652
 Regional Cancer Care Associates LLC
 Sewell, New Jersey, United States, 08080
 United States, New York
 Montefiore Medical Center
 Bronx, New York, United States, 10467
 Maimonides Medical Center
 Brooklyn, New York, United States, 11219
 United States, North Carolina
 First Health of the Carolinas
 Pinehurst, North Carolina, United States, 28374
 United States, Ohio
 University of Cincinnati
 Cincinnati, Ohio, United States, 45203-0542
 Mercy St Anne Hospital
 Toledo, Ohio, United States, 43623
 United States, Oregon
 Bend Memorial Clinic
 Bend, Oregon, United States, 97701
 St. Charles Medical Center Bend; Cancer Care Of The Cascades
 Bend, Oregon, United States, 97701
 Willamette Valley Cancer Insitute and Research Center
 Springfield, Oregon, United States, 97477
 United States, Pennsylvania
 St. Luke's Cancer Care Associates
 Bethlehem, Pennsylvania, United States, 18015
 Allegheny Cancer Center
 Pittsburgh, Pennsylvania, United States, 15212
 Univ of Pittsburgh Medical Ctr
 Pittsburgh, Pennsylvania, United States, 15232
 United States, Tennessee
 West Clinic
 Germantown, Tennessee, United States, 38138
 Tennessee Cancer Specialists
 Knoxville, Tennessee, United States, 37920
 United States, Texas
 Houston Methodist Cancer Center
 Houston, Texas, United States, 77030
 Longview Cancer Center
 Longview, Texas, United States, 75601
 University of Texas Health Science Center at San Antonio
 San Antonio, Texas, United States, 78229
 United States, Virginia
 Virginia Cancer Specialists, PC
 Fairfax, Virginia, United States, 22031
 Virginia Oncology Associates
 Norfolk, Virginia, United States, 23502
 Virginia Cancer Institute
 Richmond, Virginia, United States, 23226
 Blue Ridge Cancer Care
 Roanoke, Virginia, United States, 24014
 United States, Washington
 MultiCare Regional Cancer Center - Auburn
 Auburn, Washington, United States, 98002-4117
 Providence Regional Cancer Partnership
 Everett, Washington, United States, 98201
 Virginia Mason Medical Center
 Seattle, Washington, United States, 98101
 Medical Oncology Associates
 Spokane, Washington, United States, 99208
 United States, West Virginia
 West Virginia University; Mary Babb Randolph Can Ctr

Morgantown, West Virginia, United States, 26506
Argentina
Centro Medico Austral
Buenos Aires, Argentina, 1019
Fundación CENIT para la Investigación en Neurociencias
Buenos Aires, Argentina, C1125ABD
Sanatorio Allende
Cordoba, Argentina, X5000JHQ
Centro Oncologico Riojano Integral (CORI)
La Rioja, Argentina, F5300COE
Hospital Provincial del Centenario
Rosario, Argentina, 2000
Fundacion Koriza
Santa Rosa, Argentina, 6300
Centro de Investigacion; Clinica - Clinica Viedma S.A.
Viedma, Argentina, R8500ACE
Australia, New South Wales
Chris O'Brien Lifehouse
Camperdown, New South Wales, Australia, 2050
Concord Repatriation General Hospital
Concord, New South Wales, Australia, 2139
Nepean Cancer Care Centre
Sydney, New South Wales, Australia, 2747
Australia, Queensland
Prince Charles Hospital; Department of Medical Oncology
Chermside, Queensland, Australia, 4032
Townsville Hospital
Townsville, Queensland, Australia, 4810
Princess Alexandra Hospital
Woolloongabba, Queensland, Australia, 4102
Australia, South Australia
Royal Adelaide Hospital
Adelaide, South Australia, Australia, 5000
Adelaide Cancer Centre
Kurralta Park, South Australia, Australia, 5037
Australia, Tasmania
Royal Hobart Hospital
Hobart, Tasmania, Australia, 7000
Launceston General Hospital
Launceston, Tasmania, Australia, 7250
Australia, Victoria
Frankston Hospital
Frankston, Victoria, Australia, 3199
Austin Health
Heidelberg, Victoria, Australia, 3084
Cabrini Hospital Malvern
Malvern, Victoria, Australia, 3144
The Alfred Hospital
Prahan, Victoria, Australia, 3181
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 Hospital Clinic de Barcelona
 Barcelona, Spain, 08036
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 Hospital General Universitario Gregorio Marañón; Servicio de Oncologia
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 Hospital Universitario La Paz
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 Hospital Ramon y Cajal; Servicio de Oncologia
 Madrid, Spain, 28034
 Hospital Clinico San Carlos; Servicio de Oncologia
 Madrid, Spain, 28040
 Hospital Universitario Fundación Jimenez Díaz
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 Hospital Universitario 12 de Octubre
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Outcomes

The two primary end points of IMpower150 were progression-free survival (as assessed by investigators according to RECIST criteria) both among patients in the intention-to-treat population who had a wild-type genotype (WT population; patients with EGFR or ALK genomic alterations were excluded) and among patients in the WT population who had high expression of an effector T-cell (Teff) gene signature in the tumor (Teff-high WT population) and overall survival in the WT population. The ctDNA substudy also used the Overall Survival and radiographic response as assessed by RECIST v1.1. Additional details on the IMpower150 study can be found in the primary clinical manuscript.