

(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization
International Bureau



(10) International Publication Number

WO 2019/222657 A1

(43) International Publication Date
21 November 2019 (21.11.2019)

(51) International Patent Classification:

G01N 33/50 (2006.01) *C12Q 1/6869* (2018.01)
A61P 35/00 (2006.01) *C12N 15/07* (2006.01)

EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV,
MC, MK, MT, NL, NO, PL, PT, RO, RS, SE, SI, SK, SM,
TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW,
KM, ML, MR, NE, SN, TD, TG).

(21) International Application Number:

PCT/US2019/032914

Published:

— with international search report (Art. 21(3))

(22) International Filing Date:

17 May 2019 (17.05.2019)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

62/673,516 18 May 2018 (18.05.2018) US
62/795,900 23 January 2019 (23.01.2019) US

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(81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AO, AT, AU, AZ, BA, BB, BG, BH, BN, BR, BW, BY, BZ, CA, CH, CL, CN, CO, CR, CU, CZ, DE, DJ, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IR, IS, JO, JP, KE, KG, KH, KN, KP, KR, KW, KZ, LA, LC, LK, LR, LS, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PA, PE, PG, PH, PL, PT, QA, RO, RS, RU, RW, SA, SC, SD, SE, SG, SK, SL, SM, ST, SV, SY, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW.

(84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LR, LS, MW, MZ, NA, RW, SD, SL, ST, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, RU, TJ, TM), European (AL, AT, BE, BG, CH, CY, CZ, DE, DK,

(54) Title: CELL-FREE DNA FOR ASSESSING AND/OR TREATING CANCER

(57) Abstract: This document relates to methods and materials for assessed, monitored, and/or treated mammals (e.g., humans) having cancer. For example, methods and materials for identifying a mammal as having cancer (e.g., a localized cancer) are provided. For example, methods and materials for assessing, monitoring, and/or treating a mammal having cancer are provided.

WO 2019/222657 A1

CELL-FREE DNA FOR ASSESSING AND/OR TREATING CANCER

CROSS-REFERENCE TO RELATED APPLICATIONS

This application claims the benefit of U.S. Patent Application Serial No. 62/673,516, filed on May 18, 2018, and claims the benefit of U.S. Patent Application Serial No. 5 62/795,900, filed on January 23, 2019. The disclosure of the prior applications are considered part of (and are incorporated by reference in) the disclosure of this application.

STATEMENT REGARDING FEDERAL FUNDING

This invention was made with U.S. government support under grant No. CA121113 from the National Institutes of Health. The U.S. government has certain rights in the 10 invention.

BACKGROUND

1. Technical Field

This document relates to methods and materials for assessing and/or treating mammals (e.g., humans) having cancer. For example, this document provides methods and 15 materials for identifying a mammal as having cancer (e.g., a localized cancer). For example, this document provides methods and materials for monitoring and/or treating a mammal having cancer.

2. Background Information

Much of the morbidity and mortality of human cancers world-wide is a result of the 20 late diagnosis of these diseases, where treatments are less effective (Torre et al., 2015 *CA Cancer J Clin* 65:87; and World Health Organization, 2017 *Guide to Cancer Early Diagnosis*). Unfortunately, clinically proven biomarkers that can be used to broadly diagnose and treat patients are not widely available (Mazzucchelli, 2000 *Advances in clinical pathology* 4:111; Ruibal Morell, 1992 *The International journal of biological markers* 7:160; 25 Galli et al., 2013 *Clinical chemistry and laboratory medicine* 51:1369; Sikaris, 2011 *Heart, lung & circulation* 20:634; Lin et al., 2016 in *Screening for Colorectal Cancer: A Systematic*

Review for the U.S. Preventive Services Task Force. (Rockville, MD); Wanebo et al., 1978 N Engl J Med 299:448; and Zauber, 2015 *Dig Dis Sci* 60:681).

SUMMARY

Recent analyses of cell-free DNA suggests that such approaches may provide new avenues for early diagnosis (Phallen et al., 2017 *Sci Transl Med* 9; Cohen et al., 2018 *Science* 359:926; Alix-Panabieres et al., 2016 *Cancer discovery* 6:479; Siravegna et al., 2017 *Nature reviews. Clinical oncology* 14:531; Haber et al., 2014 *Cancer discovery* 4:650; Husain et al., 2017 *JAMA* 318:1272; and Wan et al., 2017 *Nat Rev Cancer* 17:223).

This document provides methods and materials for determining a cell free DNA (cfDNA) fragmentation profile in a mammal (e.g., in a sample obtained from a mammal). In some cases, determining a cfDNA fragmentation profile in a mammal can be used for identifying a mammal as having cancer. For example, cfDNA fragments obtained from a mammal (e.g., from a sample obtained from a mammal) can be subjected to low coverage whole-genome sequencing, and the sequenced fragments can be mapped to the genome (e.g., in non-overlapping windows) and assessed to determine a cfDNA fragmentation profile. This document also provides methods and materials for assessing and/or treating mammals (e.g., humans) having, or suspected of having, cancer. In some cases, this document provides methods and materials for identifying a mammal as having cancer. For example, a sample (e.g., a blood sample) obtained from a mammal can be assessed to determine if the mammal has cancer based, at least in part, on the cfDNA fragmentation profile. In some cases, this document provides methods and materials for monitoring and/or treating a mammal having cancer. For example, one or more cancer treatments can be administered to a mammal identified as having cancer (e.g., based, at least in part, on a cfDNA fragmentation profile) to treat the mammal.

Described herein is a non-invasive method for the early detection and localization of cancer. cfDNA in the blood can provide a non-invasive diagnostic avenue for patients with cancer. As demonstrated herein, **DNA Evaluation of Fragments for early Interception (DELFI)** was developed and used to evaluate genome-wide fragmentation patterns of cfDNA of 236 patients with breast, colorectal, lung, ovarian, pancreatic, gastric, or bile duct cancers as well as 245 healthy individuals. These analyses revealed that cfDNA profiles of healthy

individuals reflected nucleosomal fragmentation patterns of white blood cells, while patients with cancer had altered fragmentation profiles. DELFI had sensitivities of detection ranging from 57% to >99% among the seven cancer types at 98% specificity and identified the tissue of origin of the cancers to a limited number of sites in 75% of cases. Assessing cfDNA (e.g., 5 using DELFI) can provide a screening approach for early detection of cancer, which can increase the chance for successful treatment of a patient having cancer. Assessing cfDNA (e.g., using DELFI) can also provide an approach for monitoring cancer, which can increase the chance for successful treatment and improved outcome of a patient having cancer. In addition, a cfDNA fragmentation profile can be obtained from limited amounts of cfDNA and 10 using inexpensive reagents and/or instruments.

In general, one aspect of this document features methods for determining a cfDNA fragmentation profile of a mammal. The methods can include, or consist essentially of, processing cfDNA fragments obtained from a sample obtained from the mammal into sequencing libraries, subjecting the sequencing libraries to whole genome sequencing (e.g., 15 low-coverage whole genome sequencing) to obtain sequenced fragments, mapping the sequenced fragments to a genome to obtain windows of mapped sequences, and analyzing the windows of mapped sequences to determine cfDNA fragment lengths. The mapped sequences can include tens to thousands of windows. The windows of mapped sequences can be non-overlapping windows. The windows of mapped sequences can each include 20 about 5 million base pairs. The cfDNA fragmentation profile can be determined within each window. The cfDNA fragmentation profile can include a median fragment size. The cfDNA fragmentation profile can include a fragment size distribution. The cfDNA fragmentation profile can include a ratio of small cfDNA fragments to large cfDNA fragments in the windows of mapped sequences. The cfDNA fragmentation profile can be over the whole 25 genome. The cfDNA fragmentation profile can be over a subgenomic interval (e.g., an interval in a portion of a chromosome).

In another aspect, this document features methods for identifying a mammal as having cancer. The methods can include, or consist essentially of, determining a cfDNA fragmentation profile in a sample obtained from a mammal, comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile, and identifying the 30 mammal as having cancer when the cfDNA fragmentation profile in the sample obtained

from the mammal is different from the reference cfDNA fragmentation profile. The reference cfDNA fragmentation profile can be a cfDNA fragmentation profile of a healthy mammal. The reference cfDNA fragmentation profile can be generated by determining a cfDNA fragmentation profile in a sample obtained from the healthy mammal. The reference
5 DNA fragmentation pattern can be a reference nucleosome cfDNA fragmentation profile. The cfDNA fragmentation profiles can include a median fragment size, and a median fragment size of the cfDNA fragmentation profile can be shorter than a median fragment size of the reference cfDNA fragmentation profile. The cfDNA fragmentation profiles can include a fragment size distribution, and a fragment size distribution of the cfDNA
10 fragmentation profile can differ by at least 10 nucleotides as compared to a fragment size distribution of the reference cfDNA fragmentation profile. The cfDNA fragmentation profiles can include position dependent differences in fragmentation patterns, including a ratio of small cfDNA fragments to large cfDNA fragments, where a small cfDNA fragment can be 100 base pairs (bp) to 150 bp in length and a large cfDNA fragments can be 151 bp to
15 220 bp in length, and where a correlation of fragment ratios in the cfDNA fragmentation profile can be lower than a correlation of fragment ratios of the reference cfDNA fragmentation profile. The cfDNA fragmentation profiles can include sequence coverage of small cfDNA fragments, large cfDNA fragments, or of both small and large cfDNA fragments, across the genome. The cancer can be colorectal cancer, lung cancer, breast
20 cancer, bile duct cancer, pancreatic cancer, gastric cancer, or ovarian cancer. The step of comparing can include comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile in windows across the whole genome. The step of comparing can include comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile over a subgenomic interval (e.g., an interval in a portion of a chromosome). The
25 mammal can have been previously administered a cancer treatment to treat the cancer. The cancer treatment can be surgery, adjuvant chemotherapy, neoadjuvant chemotherapy, radiation therapy, hormone therapy, cytotoxic therapy, immunotherapy, adoptive T cell therapy, targeted therapy, or any combinations thereof. The method also can include administering to the mammal a cancer treatment (e.g., surgery, adjuvant chemotherapy,
30 neoadjuvant chemotherapy, radiation therapy, hormone therapy, cytotoxic therapy, immunotherapy, adoptive T cell therapy, targeted therapy, or any combinations thereof). The

mammal can be monitored for the presence of cancer after administration of the cancer treatment.

In another aspect, this document features methods for treating a mammal having cancer. The methods can include, or consist essentially of, identifying the mammal as having cancer, where the identifying includes determining a cfDNA fragmentation profile in a sample obtained from the mammal, comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile, and identifying the mammal as having cancer when the cfDNA fragmentation profile obtained from the mammal is different from the reference cfDNA fragmentation profile; and administering a cancer treatment to the mammal. The mammal can be a human. The cancer can be colorectal cancer, lung cancer, breast cancer, gastric cancers, pancreatic cancers, bile duct cancers, or ovarian cancer. The cancer treatment can be surgery, adjuvant chemotherapy, neoadjuvant chemotherapy, radiation therapy, hormone therapy, cytotoxic therapy, immunotherapy, adoptive T cell therapy, targeted therapy, or combinations thereof. The reference cfDNA fragmentation profile can be a cfDNA fragmentation profile of a healthy mammal. The reference cfDNA fragmentation profile can be generated by determining a cfDNA fragmentation profile in a sample obtained from a healthy mammal. The reference DNA fragmentation pattern can be a reference nucleosome cfDNA fragmentation profile. The cfDNA fragmentation profile can include a median fragment size, where a median fragment size of the cfDNA fragmentation profile is shorter than a median fragment size of the reference cfDNA fragmentation profile. The cfDNA fragmentation profile can include a fragment size distribution, where a fragment size distribution of the cfDNA fragmentation profile differs by at least 10 nucleotides as compared to a fragment size distribution of the reference cfDNA fragmentation profile. The cfDNA fragmentation profile can include a ratio of small cfDNA fragments to large cfDNA fragments in the windows of mapped sequences, where a small cfDNA fragment is 100 bp to 150 bp in length, where a large cfDNA fragments is 151 bp to 220 bp in length, and where a correlation of fragment ratios in the cfDNA fragmentation profile is lower than a correlation of fragment ratios of the reference cfDNA fragmentation profile. The cfDNA fragmentation profile can include the sequence coverage of small cfDNA fragments in windows across the genome. The cfDNA fragmentation profile can include the sequence coverage of large cfDNA fragments in windows across the genome. The cfDNA fragmentation profile can

include the sequence coverage of small and large cfDNA fragments in windows across the genome. The step of comparing can include comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile over the whole genome. The step of comparing can include comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile over a subgenomic interval. The mammal can have previously been administered a cancer treatment to treat the cancer. The cancer treatment can be surgery, adjuvant chemotherapy, neoadjuvant chemotherapy, radiation therapy, hormone therapy, cytotoxic therapy, immunotherapy, adoptive T cell therapy, targeted therapy, or combinations thereof. The method also can include monitoring the mammal for the presence of cancer after administration of the cancer treatment.

Unless otherwise defined, all technical and scientific terms used herein have the same meaning as commonly understood by one of ordinary skill in the art to which this invention pertains. Although methods and materials similar or equivalent to those described herein can be used to practice the invention, suitable methods and materials are described below. All publications, patent applications, patents, and other references mentioned herein are incorporated by reference in their entirety. In case of conflict, the present specification, including definitions, will control. In addition, the materials, methods, and examples are illustrative only and not intended to be limiting.

The details of one or more embodiments of the invention are set forth in the accompanying drawings and the description below. Other features, objects, and advantages of the invention will be apparent from the description and drawings, and from the claims.

DESCRIPTION OF THE DRAWINGS

Figure 1. Schematic of an exemplary DELFI approach. Blood is collected from a cohort of healthy individuals and patients with cancer. Nucleosome protected cfDNA is extracted from the plasma fraction, processed into sequencing libraries, examined through whole genome sequencing, mapped to the genome, and analyzed to determine cfDNA fragment profiles in different windows across the genome. Machine learning approaches are used to categorize individuals as healthy or as having cancer and to identify the tumor tissue of origin using genome-wide cfDNA fragmentation patterns.

Figure 2. Simulations of non-invasive cancer detection based on number of alterations analyzed and tumor-derived cfDNA fragment distributions. Monte Carlo simulations were performed using different numbers of tumor-specific alterations to evaluate the probability of detecting cancer alterations in cfDNA at the indicated fraction of tumor-derived molecules. The simulations were performed assuming an average of 2000 genome equivalents of cfDNA and the requirement of five or more observations of any alteration. These analyses indicate that increasing the number of tumor-specific alterations improves the sensitivity of detection of circulating tumor DNA.

Figure 3. Tumor-derived cfDNA fragment distributions. Cumulative density functions of cfDNA fragment lengths of 42 loci containing tumor-specific alterations from 30 patients with breast, colorectal, lung, or ovarian cancer are shown with 95% confidence bands (blue). Lengths of mutant cfDNA fragments were significantly different in size compared to wild-type cfDNA fragments (red) at these loci.

Figures 4A and 4B. Tumor-derived cfDNA GC content and fragment length. A, GC content was similar for mutated and non-mutated fragments. B, GC content was not correlated to fragment length.

Figure 5. Germline cfDNA fragment distributions. Cumulative density functions of fragment lengths of 44 loci containing germline alterations (non-tumor derived) from 38 patients with breast, colorectal, lung, or ovarian cancer are shown with 95% confidence bands. Fragments with germline mutations (blue) were comparable in length to wild-type cfDNA fragment lengths (red).

Figure 6. Hematopoietic cfDNA fragment distributions. Cumulative density functions of fragment lengths of 41 loci containing hematopoietic alterations (non-tumor derived) from 28 patients with breast, colorectal, lung, or ovarian cancer are shown with 95% confidence bands. After correction for multiple testing, there were no significant differences ($\alpha=0.05$) in the size distributions of mutated hematopoietic cfDNA fragments (blue) and wild-type cfDNA fragments (red).

Figures 7A – 7F. cfDNA fragmentation profiles in healthy individuals and patients with cancer. A, Genome-wide cfDNA fragmentation profiles (defined as the ratio of short to long fragments) from ~9x whole genome sequencing are shown in 5 Mb bins for 30 healthy individuals (top) and 8 lung cancer patients (bottom). B, An analysis of healthy cfDNA

(top), lung cancer cfDNA (middle), and healthy lymphocyte (bottom) fragmentation profiles and lymphocyte profiles from chromosome 1 at 1 Mb resolution. The healthy lymphocyte profiles were scaled with a standard deviation equal to that of the median healthy cfDNA profiles. Healthy cfDNA patterns closely mirrored those in healthy lymphocytes while lung
5 cancer cfDNA profiles were more varied and differed from both healthy and lymphocyte profiles. C, Smoothed median distances between adjacent nucleosome centered at zero using 100 kb bins from healthy cfDNA (top) and nuclease-digested healthy lymphocytes (middle) are depicted together with the first eigenvector for the genome contact matrix obtained through previously reported Hi-C analyses of lymphoblastoid cells (bottom). Healthy
10 cfDNA nucleosome distances closely mirrored those in nuclease-digested lymphocytes as well as those from lymphoblastoid Hi-C analyses. cfDNA fragmentation profiles from healthy individuals (n=30) had high correlations while patients with lung cancer had lower correlations to median fragmentation profiles of lymphocytes (D), healthy cfDNA (E), and lymphocyte nucleosome (F) distances.

15 Figure 8. Density of cfDNA fragment lengths in healthy individuals and patients with lung cancer. cfDNA fragments lengths are shown for healthy individuals (n=30, gray) and patients with lung cancer (n=8, blue).

Figures 9A and 9B. Subsampling of whole genome sequence data for analysis of cfDNA fragmentation profiles. A, High coverage (9x) whole-genome sequencing data were
20 subsampled to 2x, 1x, 0.5x, 0.2x, and 0.1x fold coverage. Mean centered genome-wide fragmentation profiles in 5 Mb bins for 30 healthy individuals and 8 patients with lung cancer are depicted for each subsampled fold coverage with median profiles shown in blue. B, Pearson correlation of subsampled profiles to initial profile at 9x coverage for healthy individuals and patients with lung cancer.

25 Figure 10. cfDNA fragmentation profiles and sequence alterations during therapy. Detection and monitoring of cancer in serial blood draws from NSCLC patients (n=19) undergoing treatment with targeted tyrosine kinase inhibitors (black arrows) was performed using targeted sequencing (top) and genome-wide fragmentation profiles (bottom). For each case, the vertical axis of the lower panel displays -1 times the correlation of each sample to
30 the median healthy cfDNA fragmentation profile. Error bars depict confidence intervals from binomial tests for mutant allele fractions and confidence intervals calculated using

Fisher transformation for genome-wide fragmentation profiles. Although the approaches analyze different aspects of cfDNA (whole genome compared to specific alterations) the targeted sequencing and fragmentation profiles were similar for patients responding to therapy as well as those with stable or progressive disease. As fragmentation profiles reflect both genomic and epigenomic alterations, while mutant allele fractions only reflect individual mutations, mutant allele fractions alone may not reflect the absolute level of correlation of fragmentation profiles to healthy individuals.

Figures 11A – 11C. cfDNA fragmentation profiles in healthy individuals and patients with cancer. A, Fragmentation profiles (bottom) in the context of tumor copy number changes (top) in a colorectal cancer patient where parallel analyses of tumor tissue were performed. The distribution of segment means and integer copy numbers are shown at top right in the indicated colors. Altered fragmentation profiles were present in regions of the genome that were copy neutral and were further affected in regions with copy number changes. B, GC adjusted fragmentation profiles from 1-2x whole genome sequencing for healthy individuals and patients with cancer are depicted per cancer type using 5 Mb windows. The median healthy profile is indicated in black and the 98% confidence band is shown in gray. For patients with cancer, individual profiles are colored based on their correlation to the healthy median. C, Windows are indicated in orange if more than 10% of the cancer samples had a fragment ratio more than three standard deviations from the median healthy fragment ratio. These analyses highlight the multitude of position dependent alterations across the genome in cfDNA of individuals with cancer.

Figures 12A and 12B. Profiles of cfDNA fragment lengths in copy neutral regions in healthy individuals and one patient with colorectal cancer. A, The fragmentation profile in 211 copy neutral windows in chromosomes 1-6 for 25 randomly selected healthy individuals (gray). For a patient with colorectal cancer (CGCRC291) with an estimated mutant allele fraction of 20%, the cancer fragment length profile was diluted to an approximate 10% tumor contribution (blue). A and B, While the marginal densities of the fragment profiles for the healthy samples and cancer patient show substantial overlap (A, right), the fragmentation profiles are different as can be seen visualization of the fragmentation profiles (A, left) and by the separation of the colorectal cancer patient from the healthy samples in a principal component analysis (B).

Figures 13A and 13B. Genome-wide GC correction of cfDNA fragments. To estimate and control for the effects of GC content on sequencing coverage, coverage in non-overlapping 100kb genomic windows was calculated across the autosomes. For each window, the average GC of the aligned fragments was calculated. A, Loess smoothing of raw coverage (top row) for two randomly selected healthy subjects (CGPLH189 and CGPLH380) and two cancer patients (CGLLU161 and CGPLBR24) with undetectable aneuploidy (PA score < 2.35). After subtracting the average coverage predicted by the loess model, the residuals were rescaled to the median autosomal coverage (bottom row). As fragment length may also result in coverage biases, this GC correction procedure was performed separately for short (≤ 150 bp) and long (≥ 151 bp) fragments. While the 100 kb bins on chromosome 19 (blue points) consistently have less coverage than predicted by the loess model, we did not implement a chromosome-specific correction as such an approach would remove the effects of chromosomal copy number on coverage. B, Overall, a limited correlation was found between short or long fragment coverage and GC content after correction among healthy subjects and cancer patients with a PA score <3.

Figure 14. Schematic of machine learning model. Gradient tree boosting machine learning was used to examine whether cfDNA can be categorized as having characteristics of a cancer patient or healthy individual. The machine learning model included fragmentation size and coverage characteristics in windows throughout the genome, as well as chromosomal arm and mitochondrial DNA copy numbers. A 10-fold cross validation approach was employed in which each sample is randomly assigned to a fold and 9 of the folds (90% of the data) are used for training and one fold (10% of the data) is used for testing. The prediction accuracy from a single cross validation is an average over the 10 possible combinations of test and training sets. As this prediction accuracy can reflect bias from the initial randomization of patients, the entire procedure was repeat, including the randomization of patients to folds, 10 times. For all cases, feature selection and model estimation were performed on training data and were validated on test data and the test data were never used for feature selection. Ultimately, a DELFI score was obtained that could be used to classify individuals as likely healthy or having cancer.

Figure 15. Distribution of AUCs across the repeated 10-fold cross-validation. The 25th, 50th, and 75th percentiles of the 100 AUCs for the cohort of 215 healthy individuals and 208 patients with cancer are indicated by dashed lines.

Figures 16A and 16B. Whole-genome analyses of chromosomal arm copy number changes and mitochondrial genome representation. A, Z scores for each autosome arm are depicted for healthy individuals (n=215) and patients with cancer (n=208). The vertical axis depicts normal copy at zero with positive and negative values indicating arm gains and losses, respectively. Z scores greater than 50 or less than -50 are thresholded at the indicated values. B, The fraction of reads mapping to the mitochondrial genome is depicted for healthy individuals and patients with cancer.

Figures 17A and 17B. Detection of cancer using DELFI. A, Receiver operator characteristics for detection of cancer using cfDNA fragmentation profiles and other genome-wide features in a machine learning approach are depicted for a cohort of 215 healthy individuals and 208 patients with cancer (DELF1, AUC = 0.94), with ≥ 95% specificity shaded in blue. Machine learning analyses of chromosomal arm copy number (Chr copy number (ML)), and mitochondrial genome copy number (mtDNA), are shown in the indicated colors. B, Analyses of individual cancers types using the DELFI-combined approach had AUCs ranging from 0.86 to >0.99.

Figure 18. DELFI detection of cancer by stage. Receiver operator characteristics for detection of cancer using cfDNA fragmentation profiles and other genome-wide features in a machine learning approach are depicted for a cohort of 215 healthy individuals and each stage of 208 patients with cancer with ≥ 95% specificity shaded in blue.

Figure 19. DELFI tissue of origin prediction. Receiver operator characteristics for DELFI tissue prediction of bile duct, breast, colorectal, gastric, lung, ovarian, and pancreatic cancers are depicted. In order to increase sample sizes within cancer type classes, cases detected with a 90% specificity were included, and the lung cancer cohort was supplemented with the addition of baseline cfDNA data from 18 lung cancer patients with prior treatment (see, e.g., Shen *et al.*, 2018 *Nature*, 563:579–583).

Figure 20. Detection of cancer using DELFI and mutation-based cfDNA approaches. DELFI (green) and targeted sequencing for mutation identification (blue) were performed independently in a cohort of 126 patients with breast, bile duct, colorectal, gastric, lung, or

ovarian cancers. The number of individuals detected by each approach and in combination are indicated for DELFI detection with a specificity of 98%, targeted sequencing specificity at >99%, and a combined specificity of 98%. ND indicates not detected.

DETAILED DESCRIPTION

5 This document provides methods and materials for determining a cfDNA fragmentation profile in a mammal (e.g., in a sample obtained from a mammal). As used herein, the terms “fragmentation profile,” “position dependent differences in fragmentation patterns,” and “differences in fragment size and coverage in a position dependent manner across the genome” are equivalent and can be used interchangeably. In some cases,
10 determining a cfDNA fragmentation profile in a mammal can be used for identifying a mammal as having cancer. For example, cfDNA fragments obtained from a mammal (e.g., from a sample obtained from a mammal) can be subjected to low coverage whole-genome sequencing, and the sequenced fragments can be mapped to the genome (e.g., in non-overlapping windows) and assessed to determine a cfDNA fragmentation profile. As
15 described herein, a cfDNA fragmentation profile of a mammal having cancer is more heterogeneous (e.g., in fragment lengths) than a cfDNA fragmentation profile of a healthy mammal (e.g., a mammal not having cancer). As such, this document also provides methods and materials for assessing, monitoring, and/or treating mammals (e.g., humans) having, or suspected of having, cancer. In some cases, this document provides methods and materials
20 for identifying a mammal as having cancer. For example, a sample (e.g., a blood sample) obtained from a mammal can be assessed to determine the presence and, optionally, the tissue of origin of the cancer in the mammal based, at least in part, on the cfDNA fragmentation profile of the mammal. In some cases, this document provides methods and materials for monitoring a mammal as having cancer. For example, a sample (e.g., a blood sample)
25 obtained from a mammal can be assessed to determine the presence of the cancer in the mammal based, at least in part, on the cfDNA fragmentation profile of the mammal. In some cases, this document provides methods and materials for identifying a mammal as having cancer, and administering one or more cancer treatments to the mammal to treat the mammal. For example, a sample (e.g., a blood sample) obtained from a mammal can be assessed to
30 determine if the mammal has cancer based, at least in part, on the cfDNA fragmentation

profile of the mammal, and one or more cancer treatments can be administered to the mammal.

A cfDNA fragmentation profile can include one or more cfDNA fragmentation patterns. A cfDNA fragmentation pattern can include any appropriate cfDNA fragmentation pattern. Examples of cfDNA fragmentation patterns include, without limitation, median fragment size, fragment size distribution, ratio of small cfDNA fragments to large cfDNA fragments, and the coverage of cfDNA fragments. In some cases, a cfDNA fragmentation pattern includes two or more (e.g., two, three, or four) of median fragment size, fragment size distribution, ratio of small cfDNA fragments to large cfDNA fragments, and the coverage of cfDNA fragments. In some cases, cfDNA fragmentation profile can be a genome-wide cfDNA profile (e.g., a genome-wide cfDNA profile in windows across the genome). In some cases, cfDNA fragmentation profile can be a targeted region profile. A targeted region can be any appropriate portion of the genome (e.g., a chromosomal region). Examples of chromosomal regions for which a cfDNA fragmentation profile can be determined as described herein include, without limitation, a portion of a chromosome (e.g., a portion of 2q, 4p, 5p, 6q, 7p, 8q, 9q, 10q, 11q, 12q, and/or 14q) and a chromosomal arm (e.g., a chromosomal arm of 8q, 13q, 11q, and/or 3p). In some cases, a cfDNA fragmentation profile can include two or more targeted region profiles.

In some cases, a cfDNA fragmentation profile can be used to identify changes (e.g., alterations) in cfDNA fragment lengths. An alteration can be a genome-wide alteration or an alteration in one or more targeted regions/loci. A target region can be any region containing one or more cancer-specific alterations. Examples of cancer-specific alterations, and their chromosomal locations, include, without limitation, those shown in Table 3 (Appendix C) and those shown in Table 6 (Appendix F). In some cases, a cfDNA fragmentation profile can be used to identify (e.g., simultaneously identify) from about 10 alterations to about 500 alterations (e.g., from about 25 to about 500, from about 50 to about 500, from about 100 to about 500, from about 200 to about 500, from about 300 to about 500, from about 10 to about 400, from about 10 to about 300, from about 10 to about 200, from about 10 to about 100, from about 10 to about 50, from about 20 to about 400, from about 30 to about 300, from about 40 to about 200, from about 50 to about 100, from about 20 to about 100, from about 25 to about 75, from about 50 to about 250, or from about 100 to about 200, alterations).

In some cases, a cfDNA fragmentation profile can be used to detect tumor-derived DNA. For example, a cfDNA fragmentation profile can be used to detect tumor-derived DNA by comparing a cfDNA fragmentation profile of a mammal having, or suspected of having, cancer to a reference cfDNA fragmentation profile (e.g., a cfDNA fragmentation profile of a healthy mammal and/or a nucleosomal DNA fragmentation profile of healthy cells from the mammal having, or suspected of having, cancer). In some cases, a reference cfDNA fragmentation profile is a previously generated profile from a healthy mammal. For example, methods provided herein can be used to determine a reference cfDNA fragmentation profile in a healthy mammal, and that reference cfDNA fragmentation profile can be stored (e.g., in a computer or other electronic storage medium) for future comparison to a test cfDNA fragmentation profile in mammal having, or suspected of having, cancer. In some cases, a reference cfDNA fragmentation profile (e.g., a stored cfDNA fragmentation profile) of a healthy mammal is determined over the whole genome. In some cases, a reference cfDNA fragmentation profile (e.g., a stored cfDNA fragmentation profile) of a healthy mammal is determined over a subgenomic interval.

In some cases, a cfDNA fragmentation profile can be used to identify a mammal (e.g., a human) as having cancer (e.g., a colorectal cancer, a lung cancer, a breast cancer, a gastric cancer, a pancreatic cancer, a bile duct cancer, and/or an ovarian cancer).

A cfDNA fragmentation profile can include a cfDNA fragment size pattern. cfDNA fragments can be any appropriate size. For example, cfDNA fragment can be from about 50 base pairs (bp) to about 400 bp in length. As described herein, a mammal having cancer can have a cfDNA fragment size pattern that contains a shorter median cfDNA fragment size than the median cfDNA fragment size in a healthy mammal. A healthy mammal (e.g., a mammal not having cancer) can have cfDNA fragment sizes having a median cfDNA fragment size from about 166.6 bp to about 167.2 bp (e.g., about 166.9 bp). In some cases, a mammal having cancer can have cfDNA fragment sizes that are, on average, about 1.28 bp to about 2.49 bp (e.g., about 1.88 bp) shorter than cfDNA fragment sizes in a healthy mammal. For example, a mammal having cancer can have cfDNA fragment sizes having a median cfDNA fragment size of about 164.11 bp to about 165.92 bp (e.g., about 165.02 bp).

A cfDNA fragmentation profile can include a cfDNA fragment size distribution. As described herein, a mammal having cancer can have a cfDNA size distribution that is more

variable than a cfDNA fragment size distribution in a healthy mammal. In some case, a size distribution can be within a targeted region. A healthy mammal (e.g., a mammal not having cancer) can have a targeted region cfDNA fragment size distribution of about 1 or less than about 1. In some cases, a mammal having cancer can have a targeted region cfDNA

5 fragment size distribution that is longer (e.g., 10, 15, 20, 25, 30, 35, 40, 45, 50 or more bp longer, or any number of base pairs between these numbers) than a targeted region cfDNA fragment size distribution in a healthy mammal. In some cases, a mammal having cancer can have a targeted region cfDNA fragment size distribution that is shorter (e.g., 10, 15, 20, 25, 30, 35, 40, 45, 50 or more bp shorter, or any number of base pairs between these numbers)

10 than a targeted region cfDNA fragment size distribution in a healthy mammal. In some cases, a mammal having cancer can have a targeted region cfDNA fragment size distribution that is about 47 bp smaller to about 30 bp longer than a targeted region cfDNA fragment size distribution in a healthy mammal. In some cases, a mammal having cancer can have a targeted region cfDNA fragment size distribution of, on average, a 10, 11, 12, 13, 14, 15, 15,

15 17, 18, 19, 20 or more bp difference in lengths of cfDNA fragments. For example, a mammal having cancer can have a targeted region cfDNA fragment size distribution of, on average, about a 13 bp difference in lengths of cfDNA fragments. In some case, a size distribution can be a genome-wide size distribution. A healthy mammal (e.g., a mammal not having cancer) can have very similar distributions of short and long cfDNA fragments

20 genome-wide. In some cases, a mammal having cancer can have, genome-wide, one or more alterations (e.g., increases and decreases) in cfDNA fragment sizes. The one or more alterations can be any appropriate chromosomal region of the genome. For example, an alteration can be in a portion of a chromosome. Examples of portions of chromosomes that can contain one or more alterations in cfDNA fragment sizes include, without limitation,

25 portions of 2q, 4p, 5p, 6q, 7p, 8q, 9q, 10q, 11q, 12q, and 14q. For example, an alteration can be across a chromosome arm (e.g., an entire chromosome arm).

A cfDNA fragmentation profile can include a ratio of small cfDNA fragments to large cfDNA fragments and a correlation of fragment ratios to reference fragment ratios. As used herein, with respect to ratios of small cfDNA fragments to large cfDNA fragments, a small

30 cfDNA fragment can be from about 100 bp in length to about 150 bp in length. As used herein, with respect to ratios of small cfDNA fragments to large cfDNA fragments, a large

cfDNA fragment can be from about 151 bp in length to 220 bp in length. As described herein, a mammal having cancer can have a correlation of fragment ratios (e.g., a correlation of cfDNA fragment ratios to reference DNA fragment ratios such as DNA fragment ratios from one or more healthy mammals) that is lower (e.g., 2-fold lower, 3-fold lower, 4-fold 5 lower, 5-fold lower, 6-fold lower, 7-fold lower, 8-fold lower, 9-fold lower, 10-fold lower, or more) than in a healthy mammal. A healthy mammal (e.g., a mammal not having cancer) can have a correlation of fragment ratios (e.g., a correlation of cfDNA fragment ratios to reference DNA fragment ratios such as DNA fragment ratios from one or more healthy mammals) of about 1 (e.g., about 0.96). In some cases, a mammal having cancer can have a 10 correlation of fragment ratios (e.g., a correlation of cfDNA fragment ratios to reference DNA fragment ratios such as DNA fragment ratios from one or more healthy mammals) that is, on average, about 0.19 to about 0.30 (e.g., about 0.25) lower than a correlation of fragment ratios (e.g., a correlation of cfDNA fragment ratios to reference DNA fragment ratios such as DNA fragment ratios from one or more healthy mammals) in a healthy mammal.

15 A cfDNA fragmentation profile can include coverage of all fragments. Coverage of all fragments can include windows (e.g., non-overlapping windows) of coverage. In some cases, coverage of all fragments can include windows of small fragments (e.g., fragments from about 100 bp to about 150 bp in length). In some cases, coverage of all fragments can include windows of large fragments (e.g., fragments from about 151 bp to about 220 bp in 20 length).

In some cases, a cfDNA fragmentation profile can be used to identify the tissue of origin of a cancer (e.g., a colorectal cancer, a lung cancer, a breast cancer, a gastric cancer, a pancreatic cancer, a bile duct cancer, or an ovarian cancer). For example, a cfDNA fragmentation profile can be used to identify a localized cancer. When a cfDNA 25 fragmentation profile includes a targeted region profile, one or more alterations described herein (e.g., in Table 3 (Appendix C) and/or in Table 6 (Appendix F)) can be used to identify the tissue of origin of a cancer. In some cases, one or more alterations in chromosomal regions can be used to identify the tissue of origin of a cancer.

A cfDNA fragmentation profile can be obtained using any appropriate method. In 30 some cases, cfDNA from a mammal (e.g., a mammal having, or suspected of having, cancer) can be processed into sequencing libraries which can be subjected to whole genome

sequencing (e.g., low-coverage whole genome sequencing), mapped to the genome, and analyzed to determine cfDNA fragment lengths. Mapped sequences can be analyzed in non-overlapping windows covering the genome. Windows can be any appropriate size. For example, windows can be from thousands to millions of bases in length. As one non-limiting 5 example, a window can be about 5 megabases (Mb) long. Any appropriate number of windows can be mapped. For example, tens to thousands of windows can be mapped in the genome. For example, hundreds to thousands of windows can be mapped in the genome. A cfDNA fragmentation profile can be determined within each window. In some cases, a cfDNA fragmentation profile can be obtained as described in Example 1. In some cases, a 10 cfDNA fragmentation profile can be obtained as shown in Figure 1.

In some cases, methods and materials described herein also can include machine learning. For example, machine learning can be used for identifying an altered fragmentation profile (e.g., using coverage of cfDNA fragments, fragment size of cfDNA fragments, coverage of chromosomes, and mtDNA).

15 In some cases, methods and materials described herein can be the sole method used to identify a mammal (e.g., a human) as having cancer (e.g., a colorectal cancer, a lung cancer, a breast cancer, a gastric cancer, a pancreatic cancer, a bile duct cancer, and/or an ovarian cancer). For example, determining a cfDNA fragmentation profile can be the sole method used to identify a mammal as having cancer.

20 In some cases, methods and materials described herein can be used together with one or more additional methods used to identify a mammal (e.g., a human) as having cancer (e.g., a colorectal cancer, a lung cancer, a breast cancer, a gastric cancer, a pancreatic cancer, a bile duct cancer, and/or an ovarian cancer). Examples of methods used to identify a mammal as having cancer include, without limitation, identifying one or more cancer-specific sequence alterations, identifying one or more chromosomal alterations (e.g., aneuploidies and 25 rearrangements), and identifying other cfDNA alterations. For example, determining a cfDNA fragmentation profile can be used together with identifying one or more cancer-specific mutations in a mammal's genome to identify a mammal as having cancer. For example, determining a cfDNA fragmentation profile can be used together with identifying 30 one or more aneuploidies in a mammal's genome to identify a mammal as having cancer.

In some aspects, this document also provides methods and materials for assessing, monitoring, and/or treating mammals (e.g., humans) having, or suspected of having, cancer. In some cases, this document provides methods and materials for identifying a mammal as having cancer. For example, a sample (e.g., a blood sample) obtained from a mammal can be assessed to determine if the mammal has cancer based, at least in part, on the cfDNA fragmentation profile of the mammal. In some cases, this document provides methods and materials for identifying the location (e.g., the anatomic site or tissue of origin) of a cancer in a mammal. For example, a sample (e.g., a blood sample) obtained from a mammal can be assessed to determine the tissue of origin of the cancer in the mammal based, at least in part, on the cfDNA fragmentation profile of the mammal. In some cases, this document provides methods and materials for identifying a mammal as having cancer, and administering one or more cancer treatments to the mammal to treat the mammal. For example, a sample (e.g., a blood sample) obtained from a mammal can be assessed to determine if the mammal has cancer based, at least in part, on the cfDNA fragmentation profile of the mammal, and administering one or more cancer treatments to the mammal. In some cases, this document provides methods and materials for treating a mammal having cancer. For example, one or more cancer treatments can be administered to a mammal identified as having cancer (e.g., based, at least in part, on the cfDNA fragmentation profile of the mammal) to treat the mammal. In some cases, during or after the course of a cancer treatment (e.g., any of the cancer treatments described herein), a mammal can undergo monitoring (or be selected for increased monitoring) and/or further diagnostic testing. In some cases, monitoring can include assessing mammals having, or suspected of having, cancer by, for example, assessing a sample (e.g., a blood sample) obtained from the mammal to determine the cfDNA fragmentation profile of the mammal as described herein, and changes in the cfDNA fragmentation profiles over time can be used to identify response to treatment and/or identify the mammal as having cancer (e.g., a residual cancer).

Any appropriate mammal can be assessed, monitored, and/or treated as described herein. A mammal can be a mammal having cancer. A mammal can be a mammal suspected of having cancer. Examples of mammals that can be assessed, monitored, and/or treated as described herein include, without limitation, humans, primates such as monkeys, dogs, cats, horses, cows, pigs, sheep, mice, and rats. For example, a human having, or suspected of

having, cancer can be assessed to determine a cfDNA fragmentation profiled as described herein and, optionally, can be treated with one or more cancer treatments as described herein.

Any appropriate sample from a mammal can be assessed as described herein (e.g., assessed for a DNA fragmentation pattern). In some cases, a sample can include DNA (e.g., genomic DNA). In some cases, a sample can include cfDNA (e.g., circulating tumor DNA (ctDNA)). In some cases, a sample can be fluid sample (e.g., a liquid biopsy). Examples of samples that can contain DNA and/or polypeptides include, without limitation, blood (e.g., whole blood, serum, or plasma), amnion, tissue, urine, cerebrospinal fluid, saliva, sputum, broncho-alveolar lavage, bile, lymphatic fluid, cyst fluid, stool, ascites, pap smears, breast milk, and exhaled breath condensate. For example, a plasma sample can be assessed to determine a cfDNA fragmentation profiled as described herein.

A sample from a mammal to be assessed as described herein (e.g., assessed for a DNA fragmentation pattern) can include any appropriate amount of cfDNA. In some cases, a sample can include a limited amount of DNA. For example, a cfDNA fragmentation profile can be obtained from a sample that includes less DNA than is typically required for other cfDNA analysis methods, such as those described in, for example, Phallen et al., 2017 *Sci Transl Med* 9; Cohen et al., 2018 *Science* 359:926; Newman et al., 2014 *Nat Med* 20:548; and Newman et al., 2016 *Nat Biotechnol* 34:547).

In some cases, a sample can be processed (e.g., to isolate and/or purify DNA and/or polypeptides from the sample). For example, DNA isolation and/or purification can include cell lysis (e.g., using detergents and/or surfactants), protein removal (e.g., using a protease), and/or RNA removal (e.g., using an RNase). As another example, polypeptide isolation and/or purification can include cell lysis (e.g., using detergents and/or surfactants), DNA removal (e.g., using a DNase), and/or RNA removal (e.g., using an RNase).

A mammal having, or suspected of having, any appropriate type of cancer can be assessed (e.g., to determine a cfDNA fragmentation profile) and/or treated (e.g., by administering one or more cancer treatments to the mammal) using the methods and materials described herein. A cancer can be any stage cancer. In some cases, a cancer can be an early stage cancer. In some cases, a cancer can be an asymptomatic cancer. In some cases, a cancer can be a residual disease and/or a recurrence (e.g., after surgical resection and/or after cancer therapy). A cancer can be any type of cancer. Examples of types of

cancers that can be assessed, monitored, and/or treated as described herein include, without limitation, colorectal cancers, lung cancers, breast cancers, gastric cancers, pancreatic cancers, bile duct cancers, and ovarian cancers.

When treating a mammal having, or suspected of having, cancer as described herein, 5 the mammal can be administered one or more cancer treatments. A cancer treatment can be any appropriate cancer treatment. One or more cancer treatments described herein can be administered to a mammal at any appropriate frequency (e.g., once or multiple times over a period of time ranging from days to weeks). Examples of cancer treatments include, without limitation adjuvant chemotherapy, neoadjuvant chemotherapy, radiation therapy, hormone 10 therapy, cytotoxic therapy, immunotherapy, adoptive T cell therapy (e.g., chimeric antigen receptors and/or T cells having wild-type or modified T cell receptors), targeted therapy such as administration of kinase inhibitors (e.g., kinase inhibitors that target a particular genetic lesion, such as a translocation or mutation), (e.g. a kinase inhibitor, an antibody, a bispecific antibody), signal transduction inhibitors, bispecific antibodies or antibody fragments (e.g., 15 BiTEs), monoclonal antibodies, immune checkpoint inhibitors, surgery (e.g., surgical resection), or any combination of the above. In some cases, a cancer treatment can reduce the severity of the cancer, reduce a symptom of the cancer, and/or to reduce the number of cancer cells present within the mammal.

In some cases, a cancer treatment can include an immune checkpoint inhibitor. Non- 20 limiting examples of immune checkpoint inhibitors include nivolumab (Opdivo), pembrolizumab (Keytruda), atezolizumab (tecentriq), avelumab (bavencio), durvalumab (imfinzi), ipilimumab (yervoy). See, e.g., Pardoll (2012) Nat. Rev Cancer 12: 252-264; Sun et al. (2017) Eur Rev Med Pharmacol Sci 21(6): 1198-1205; Hamanishi et al. (2015) J. Clin. Oncol. 33(34): 4015-22; Brahmer et al. (2012) N Engl J Med 366(26): 2455-65; Ricciuti et 25 al. (2017) J. Thorac Oncol. 12(5): e51-e55; Ellis et al. (2017) Clin Lung Cancer pii: S1525-7304(17)30043-8; Zou and Awad (2017) Ann Oncol 28(4): 685-687; Sorscher (2017) N Engl J Med 376(10): 996-7; Hui et al. (2017) Ann Oncol 28(4): 874-881; Vansteenkiste et al. (2017) Expert Opin Biol Ther 17(6): 781-789; Hellmann et al. (2017) Lancet Oncol. 18(1): 31-41; Chen (2017) J. Chin Med Assoc 80(1): 7-14.

30 In some cases, a cancer treatment can be an adoptive T cell therapy (e.g., chimeric antigen receptors and/or T cells having wild-type or modified T cell receptors). See, e.g.,

Rosenberg and Restifo (2015) Science 348(6230): 62-68; Chang and Chen (2017) Trends Mol Med 23(5): 430-450; Yee and Lizee (2016) Cancer J. 23(2): 144-148; Chen et al. (2016) Oncoimmunology 6(2): e1273302; US 2016/0194404; US 2014/0050788; US 2014/0271635; US 9,233,125; incorporated by reference in their entirety herein.

5 In some cases, a cancer treatment can be a chemotherapeutic agent. Non-limiting examples of chemotherapeutic agents include: amsacrine, azacitidine, axathioprine, bevacizumab (or an antigen-binding fragment thereof), bleomycin, busulfan, carboplatin, capecitabine, chlorambucil, cisplatin, cyclophosphamide, cytarabine, dacarbazine, daunorubicin, docetaxel, doxifluridine, doxorubicin, epirubicin, erlotinib hydrochlorides, 10 etoposide, fludarabine, floxuridine, fludarabine, fluorouracil, gemcitabine, hydroxyurea, idarubicin, ifosfamide, irinotecan, lomustine, mechlorethamine, melphalan, mercaptopurine, methotrexate, mitomycin, mitoxantrone, oxaliplatin, paclitaxel, pemetrexed, procarbazine, all-trans retinoic acid, streptozocin, tafluposide, temozolomide, teniposide, tioguanine, topotecan, uramustine, valrubicin, vinblastine, vincristine, vindesine, vinorelbine, and 15 combinations thereof. Additional examples of anti-cancer therapies are known in the art; see, e.g. the guidelines for therapy from the American Society of Clinical Oncology (ASCO), European Society for Medical Oncology (ESMO), or National Comprehensive Cancer Network (NCCN).

When monitoring a mammal having, or suspected of having, cancer as described 20 herein (e.g., based, at least in part, on the cfDNA fragmentation profile of the mammal), the monitoring can be before, during, and/or after the course of a cancer treatment. Methods of monitoring provided herein can be used to determine the efficacy of one or more cancer treatments and/or to select a mammal for increased monitoring. In some cases, the monitoring can include identifying a cfDNA fragmentation profile as described herein. For 25 example, a cfDNA fragmentation profile can be obtained before administering one or more cancer treatments to a mammal having, or suspected or having, cancer, one or more cancer treatments can be administered to the mammal, and one or more cfDNA fragmentation profiles can be obtained during the course of the cancer treatment. In some cases, a cfDNA fragmentation profile can change during the course of cancer treatment (e.g., any of the 30 cancer treatments described herein). For example, a cfDNA fragmentation profile indicative that the mammal has cancer can change to a cfDNA fragmentation profile indicative that the

mammal does not have cancer. Such a cfDNA fragmentation profile change can indicate that the cancer treatment is working. Conversely, a cfDNA fragmentation profile can remain static (e.g., the same or approximately the same) during the course of cancer treatment (e.g., any of the cancer treatments described herein). Such a static cfDNA fragmentation profile 5 can indicate that the cancer treatment is not working. In some cases, the monitoring can include conventional techniques capable of monitoring one or more cancer treatments (e.g., the efficacy of one or more cancer treatments). In some cases, a mammal selected for increased monitoring can be administered a diagnostic test (e.g., any of the diagnostic tests disclosed herein) at an increased frequency compared to a mammal that has not been selected 10 for increased monitoring. For example, a mammal selected for increased monitoring can be administered a diagnostic test at a frequency of twice daily, daily, bi-weekly, weekly, bi-monthly, monthly, quarterly, semi-annually, annually, or any at frequency therein. In some cases, a mammal selected for increased monitoring can be administered a one or more additional diagnostic tests compared to a mammal that has not been selected for increased 15 monitoring. For example, a mammal selected for increased monitoring can be administered two diagnostic tests, whereas a mammal that has not been selected for increased monitoring is administered only a single diagnostic test (or no diagnostic tests). In some cases, a mammal that has been selected for increased monitoring can also be selected for further diagnostic testing. Once the presence of a tumor or a cancer (e.g., a cancer cell) has been 20 identified (e.g., by any of the variety of methods disclosed herein), it may be beneficial for the mammal to undergo both increased monitoring (e.g., to assess the progression of the tumor or cancer in the mammal and/or to assess the development of one or more cancer biomarkers such as mutations), and further diagnostic testing (e.g., to determine the size and/or exact location (e.g., tissue of origin) of the tumor or the cancer). In some cases, one 25 or more cancer treatments can be administered to the mammal that is selected for increased monitoring after a cancer biomarker is detected and/or after the cfDNA fragmentation profile of the mammal has not improved or deteriorated. Any of the cancer treatments disclosed herein or known in the art can be administered. For example, a mammal that has been selected for increased monitoring can be further monitored, and a cancer treatment can be 30 administered if the presence of the cancer cell is maintained throughout the increased monitoring period. Additionally or alternatively, a mammal that has been selected for

increased monitoring can be administered a cancer treatment, and further monitored as the cancer treatment progresses. In some cases, after a mammal that has been selected for increased monitoring has been administered a cancer treatment, the increased monitoring will reveal one or more cancer biomarkers (e.g., mutations). In some cases, such one or more 5 cancer biomarkers will provide cause to administer a different cancer treatment (e.g., a resistance mutation may arise in a cancer cell during the cancer treatment, which cancer cell harboring the resistance mutation is resistant to the original cancer treatment).

When a mammal is identified as having cancer as described herein (e.g., based, at least in part, on the cfDNA fragmentation profile of the mammal), the identifying can be 10 before and/or during the course of a cancer treatment. Methods of identifying a mammal as having cancer provided herein can be used as a first diagnosis to identify the mammal (e.g., as having cancer before any course of treatment) and/or to select the mammal for further diagnostic testing. In some cases, once a mammal has been determined to have cancer, the mammal may be administered further tests and/or selected for further diagnostic testing. In 15 some cases, methods provided herein can be used to select a mammal for further diagnostic testing at a time period prior to the time period when conventional techniques are capable of diagnosing the mammal with an early-stage cancer. For example, methods provided herein for selecting a mammal for further diagnostic testing can be used when a mammal has not been diagnosed with cancer by conventional methods and/or when a mammal is not known to 20 harbor a cancer. In some cases, a mammal selected for further diagnostic testing can be administered a diagnostic test (e.g., any of the diagnostic tests disclosed herein) at an increased frequency compared to a mammal that has not been selected for further diagnostic testing. For example, a mammal selected for further diagnostic testing can be administered a diagnostic test at a frequency of twice daily, daily, bi-weekly, weekly, bi-monthly, monthly, 25 quarterly, semi-annually, annually, or any at frequency therein. In some cases, a mammal selected for further diagnostic testing can be administered a one or more additional diagnostic tests compared to a mammal that has not been selected for further diagnostic testing. For example, a mammal selected for further diagnostic testing can be administered two diagnostic tests, whereas a mammal that has not been selected for further diagnostic 30 testing is administered only a single diagnostic test (or no diagnostic tests). In some cases, the diagnostic testing method can determine the presence of the same type of cancer (e.g.,

having the same tissue or origin) as the cancer that was originally detected (e.g., based, at least in part, on the cfDNA fragmentation profile of the mammal). Additionally or alternatively, the diagnostic testing method can determine the presence of a different type of cancer as the cancer that was original detected. In some cases, the diagnostic testing method 5 is a scan. In some cases, the scan is a computed tomography (CT), a CT angiography (CTA), a esophagram (a Barium swallow), a Barium enema, a magnetic resonance imaging (MRI), a PET scan, an ultrasound (e.g., an endobronchial ultrasound, an endoscopic ultrasound), an X-ray, a DEXA scan. In some cases, the diagnostic testing method is a physical examination, such as an anoscopy, a bronchoscopy (e.g., an autofluorescence bronchoscopy, a white-light 10 bronchoscopy, a navigational bronchoscopy), a colonoscopy, a digital breast tomosynthesis, an endoscopic retrograde cholangiopancreatography (ERCP), an ensophagogastroduodenoscopy, a mammography, a Pap smear, a pelvic exam, a positron emission tomography and computed tomography (PET-CT) scan. In some cases, a mammal that has been selected for further diagnostic testing can also be selected for increased 15 monitoring. Once the presence of a tumor or a cancer (e.g., a cancer cell) has been identified (e.g., by any of the variety of methods disclosed herein), it may be beneficial for the mammal to undergo both increased monitoring (e.g., to assess the progression of the tumor or cancer in the mammal and/or to assess the development of one or more cancer biomarkers such as mutations), and further diagnostic testing (e.g., to determine the size and/or exact location of 20 the tumor or the cancer). In some cases, a cancer treatment is administered to the mammal that is selected for further diagnostic testing after a cancer biomarker is detected and/or after the cfDNA fragmentation profile of the mammal has not improved or deteriorated. Any of the cancer treatments disclosed herein or known in the art can be administered. For example, a mammal that has been selected for further diagnostic testing can be administered a further 25 diagnostic test, and a cancer treatment can be administered if the presence of the tumor or the cancer is confirmed. Additionally or alternatively, a mammal that has been selected for further diagnostic testing can be administered a cancer treatment, and can be further monitored as the cancer treatment progresses. In some cases, after a mammal that has been selected for further diagnostic testing has been administered a cancer treatment, the 30 additional testing will reveal one or more cancer biomarkers (e.g., mutations). In some cases, such one or more cancer biomarkers (e.g., mutations) will provide cause to administer a

different cancer treatment (e.g., a resistance mutation may arise in a cancer cell during the cancer treatment, which cancer cell harboring the resistance mutation is resistant to the original cancer treatment).

The invention will be further described in the following examples, which do not limit
5 the scope of the invention described in the claims.

EXAMPLES

Example 1: Cell-free DNA fragmentation in patients with cancer

Analyses of cell free DNA have largely focused on targeted sequencing of specific genes. Such studies permit detection of a small number of tumor-specific alterations in
10 patients with cancer and not all patients, especially those with early stage disease, have detectable changes. Whole genome sequencing of cell-free DNA can identify chromosomal abnormalities and rearrangements in cancer patients but detection of such alterations has been challenging in part due to the difficulty in distinguishing a small number of abnormal from normal chromosomal changes (Leary et al., 2010 *Sci Transl Med* 2:20ra14; and Leary et
15 al., 2012 *Sci Transl Med* 4:162ra154). Other efforts have suggested nucleosome patterns and chromatin structure may be different between cancer and normal tissues, and that cfDNA in patients with cancer may result in abnormal cfDNA fragment size as well as position (Snyder et al., 2016 *Cell* 164:57; Jahr et al., 2001 *Cancer Res* 61:1659; Ivanov et al., 2015 *BMC Genomics* 16(Suppl 13):S1). However, the amount of sequencing needed for nucleosome
20 footprint analyses of cfDNA is impractical for routine analyses.

The sensitivity of any cell-free DNA approach depends on the number of potential alterations examined as well as the technical and biological limitations of detecting such changes. As a typical blood sample contains ~2000 genome equivalents of cfDNA per milliliter of plasma (Phallen et al., 2017 *Sci Transl Med* 9), the theoretical limit of detection
25 of a single alteration can be no better than one in a few thousand mutant to wild-type molecules. An approach that detects a larger number of alterations in the same number of genome equivalents would be more sensitive for detecting cancer in the circulation. Monte Carlo simulations show that increasing the number of potential abnormalities detected from only a few to tens or hundreds can potentially improve the limit of detection by orders of

magnitude, similar to recent probability analyses of multiple methylation changes in cfDNA (Figure 2).

This study presents a novel method called DELFI for detection of cancer and further identification of tissue of origin using whole genome sequencing (Figure 1). The approach 5 uses cfDNA fragmentation profiles and machine learning to distinguish patterns of healthy blood cell DNA from tumor-derived DNA and to identify the primary tumor tissue. DELFI was used for a retrospective analysis of cfDNA from 245 healthy individuals and 236 patients with breast, colorectal, lung, ovarian, pancreatic, gastric, or bile duct cancers, with most patients exhibiting localized disease. Assuming this approach had sensitivity ≥ 0.80 for 10 discriminating cancer patients from healthy individuals while maintaining a specificity of 0.95, a study of at least 200 cancer patients would enable estimation of the true sensitivity with a margin of error of 0.06 at the desired specificity of 0.95 or greater.

Materials and Methods

Patient and sample characteristics

15 Plasma samples from healthy individuals and plasma and tissue samples from patients with breast, lung, ovarian, colorectal, bile duct, or gastric cancer were obtained from ILSBio/Bioreclamation, Aarhus University, Herlev Hospital of the University of Copenhagen, Hvidovre Hospital, the University Medical Center of the University of Utrecht, the Academic Medical Center of the University of Amsterdam, the Netherlands Cancer 20 Institute, and the University of California, San Diego. All samples were obtained under Institutional Review Board approved protocols with informed consent for research use at participating institutions. Plasma samples from healthy individuals were obtained at the time of routine screening, including for colonoscopies or Pap smears. Individuals were considered healthy if they had no previous history of cancer and negative screening results.

25 Plasma samples from individuals with breast, colorectal, gastric, lung, ovarian, pancreatic, and bile duct cancer were obtained at the time of diagnosis, prior to tumor resection or therapy. Nineteen lung cancer patients analyzed for change in cfDNA fragmentation profiles across multiple time points were undergoing treatment with anti-EGFR or anti-ERBB2 therapy (see, e.g., Phallen *et al.*, 2019 *Cancer Research* 15, 1204-30 1213). Clinical data for all patients included in this study are listed in Table 1 (Appendix A).

Gender was confirmed through genomic analyses of X and Y chromosome representation. Pathologic staging of gastric cancer patients was performed after neoadjuvant therapy. Samples where the tumor stage was unknown were indicated as stage X or unknown.

Nucleosomal DNA purification

5 Viable frozen lymphocytes were elutriated from leukocytes obtained from a healthy male (C0618) and female (D0808-L) (Advanced Biotechnologies Inc., Eldersburg, MD). Aliquots of 1×10^6 cells were used for nucleosomal DNA purification using EZ Nucleosomal DNA Prep Kit (Zymo Research, Irvine, CA). Cells were initially treated with 100 µl of Nuclei Prep Buffer and incubated on ice for 5 minutes. After centrifugation at 200g for 5
10 minutes, supernatant was discarded and pelleted nuclei were treated twice with 100µl of Atlantis Digestion Buffer or with 100 µl of micrococcal nuclease (MN) Digestion Buffer. Finally, cellular nucleic DNA was fragmented with 0.5U of Atlantis dsDNase at 42°C for 20
15 minutes or 1.5U of MNase at 37°C for 20 minutes. Reactions were stopped using 5X MN Stop Buffer and DNA was purified using Zymo-Spin™ IIC Columns. Concentration and quality of eluted cellular nucleic DNA were analyzed using the Bioanalyzer 2100 (Agilent Technologies, Santa Clara, CA).

Sample preparation and sequencing of cfDNA

Whole blood was collected in EDTA tubes and processed immediately or within one day after storage at 4°C, or was collected in Streck tubes and processed within two days of
20 collection for three cancer patients who were part of the monitoring analysis. Plasma and cellular components were separated by centrifugation at 800g for 10 min at 4°C. Plasma was centrifuged a second time at 18,000g at room temperature to remove any remaining cellular debris and stored at -80°C until the time of DNA extraction. DNA was isolated from plasma using the Qiagen Circulating Nucleic Acids Kit (Qiagen GmbH) and eluted in LoBind tubes
25 (Eppendorf AG). Concentration and quality of cfDNA were assessed using the Bioanalyzer 2100 (Agilent Technologies).

NGS cfDNA libraries were prepared for whole genome sequencing and targeted sequencing using 5 to 250 ng of cfDNA as described elsewhere (see, e.g., Phallen *et al.*, 2017 *Sci Transl Med* 9:eaan2415). Briefly, genomic libraries were prepared using the NEBNext
30 DNA Library Prep Kit for Illumina [New England Biolabs (NEB)] with four main

modifications to the manufacturer's guidelines: (i) The library purification steps used the on-bead AMPure XP approach to minimize sample loss during elution and tube transfer steps (see, e.g., Fisher *et al.*, 2011 *Genome Biol* 12:R1); (ii) NEBNext End Repair, A-tailing, and adapter ligation enzyme and buffer volumes were adjusted as appropriate to accommodate
5 the on-bead AMPure XP purification strategy; (iii) a pool of eight unique Illumina dual index adapters with 8-base pair (bp) barcodes was used in the ligation reaction instead of the standard Illumina single or dual index adapters with 6- or 8-bp barcodes, respectively; and (iv) cfDNA libraries were amplified with Phusion Hot Start Polymerase.

Whole genome libraries were sequenced directly. For targeted libraries, capture was
10 performed using Agilent SureSelect reagents and a custom set of hybridization probes targeting 58 genes (see, e.g., Phallen *et al.*, 2017 *Sci Transl Med* 9:eaan2415) per the manufacturer's guidelines. The captured library was amplified with Phusion Hot Start Polymerase (NEB). Concentration and quality of captured cfDNA libraries were assessed on the Bioanalyzer 2100 using theDNA1000 Kit (Agilent Technologies). Targeted libraries
15 were sequenced using 100-bp paired-end runs on the Illumina HiSeq 2000/2500 (Illumina).

Analyses of targeted sequencing data from cfDNA

Analyses of targeted NGS data for cfDNA samples was performed as described elsewhere (see, e.g., Phallen *et al.*, 2017 *Sci Transl Med* 9:eaan2415). Briefly, primary processing was completed using Illumina CASAVA (Consensus Assessment of Sequence
20 and Variation) software (version 1.8), including demultiplexing and masking of dual-index adapter sequences. Sequence reads were aligned against the human reference genome (version hg18 or hg19) using NovoAlign with additional realignment of select regions using the Needleman-Wunsch method (see, e.g., Jones *et al.*, 2015 *Sci Transl Med* 7:283ra53). The positions of the sequence alterations have not been affected by the different genome builds.
25 Candidate mutations, consisting of point mutations, small insertions, and deletions, were identified using VariantDx (see, e.g., Jones *et al.*, 2015 *Sci Transl Med* 7:283ra53) (Personal Genome Diagnostics, Baltimore, MD) across the targeted regions of interest.

To analyze the fragment lengths of cfDNA molecules, each read pair from a cfDNA molecule was required to have a Phred quality score ≥ 30 . All duplicate ctDNA fragments,
30 defined as having the same start, end, and index barcode were removed. For each mutation,

only fragments for which one or both of the read pairs contained the mutated (or wild-type) base at the given position were included. This analysis was done using the R packages Rsamtools and GenomicAlignments.

For each genomic locus where a somatic mutation was identified, the lengths of 5 fragments containing the mutant allele were compared to the lengths of fragments of the wild-type allele. If more than 100 mutant fragments were identified, Welch's two-sample t-test was used to compare the mean fragment lengths. For loci with fewer than 100 mutant fragments, a bootstrap procedure was implemented. Specifically, replacement N fragments containing the wild-type allele, where N denotes the number of fragments with the mutation, 10 were sampled. For each bootstrap replicate of wild type fragments their median length was computed. The p-value was estimated as the fraction of bootstrap replicates with a median wild-type fragment length as or more extreme than the observed median mutant fragment length.

Analyses of whole genome sequencing data from cfDNA

15 Primary processing of whole genome NGS data for cfDNA samples was performed using Illumina CASAVA (Consensus Assessment of Sequence and Variation) software (version 1.8.2), including demultiplexing and masking of dual-index adapter sequences. Sequence reads were aligned against the human reference genome (version hg19) using ELAND.

20 Read pairs with a MAPQ score below 30 for either read and PCR duplicates were removed. hg19 autosomes were tiled into 26,236 adjacent, non-overlapping 100 kb bins. Regions of low mappability, indicated by the 10% of bins with the lowest coverage, were removed (see, e.g., Fortin *et al.*, 2015 *Genome Biol* 16:180), as were reads falling in the Duke blacklisted regions (see, e.g.,
25 hgdownload.cse.ucsc.edu/goldenpath/hg19/encodeDCC/wgEncodeMapability/). Using this approach, 361 Mb (13%) of the hg19 reference genome was excluded, including centromeric and telomeric regions. Short fragments were defined as having a length between 100 and 150 bp and long fragments were defined has having a length between 151 and 220 bp.

To account for biases in coverage attributable to GC content of the genome, the 30 locally weighted smoother loess with span $\frac{3}{4}$ was applied to the scatterplot of average

fragment GC versus coverage calculated for each 100kb bin. This loess regression was performed separately for short and long fragments to account for possible differences in GC effects on coverage in plasma by fragment length (see, e.g., Benjamini *et al.*, 2012 *Nucleic Acids Res* 40:e72). The predictions for short and long coverage explained by GC from the 5 loess model were subtracted, obtaining residuals for short and long that were uncorrelated with GC. The residuals were returned to the original scale by adding back the genome-wide median short and long estimates of coverage. This procedure was repeated for each sample to account for possible differences in GC effects on coverage between samples. To further reduce the feature space and noise, the total GC-adjusted coverage in 5 Mb bins was 10 calculated.

To compare the variability of fragment lengths from healthy subjects to fragments in patients with cancer, the standard deviation of the short to long fragmentation profiles for each individual was calculated. The standard deviations in the two groups were compared by a Wilcoxon rank sum test.

15 *Analyses of chromosome arm copy number changes*

To develop arm-level statistics for copy number changes, an approach for aneuploidy detection in plasma as described elsewhere (see, e.g., Leary *et al.*, 2012 *Sci Transl Med* 4:162ra154) was adopted. This approach divides the genome into non-overlapping 50KB bins for which GC-corrected log₂ read depth was obtained after correction by loess with span 20 3/4. This loess-based correction is comparable to the approach outlined above, but is evaluated on a log₂ scale to increase robustness to outliers in the smaller bins and does not stratify by fragment length. To obtain an arm-specific Z-score for copy number changes, the mean GC-adjusted read depth for each arm (GR) was centered and scaled by the average and standard deviation, respectively, of GR scores obtained from an independent set of 50 25 healthy samples.

Analyses of mitochondrial-aligned reads from cfDNA

Whole genome sequence reads that initially mapped to the mitochondrial genome were extracted from bam files and realigned to the hg19 reference genome in end-to-end mode with Bowtie2 as described elsewhere (see, e.g., Langmead *et al.*, 2012 *Nat Methods* 30 9:357-359). The resulting aligned reads were filtered such that both mates aligned to the

mitochondrial genome with MAPQ ≥ 30 . The number of fragments mapping to the mitochondrial genome was counted and converted to a percentage of the total number of fragments in the original bam files.

Prediction model for cancer classification

To distinguish healthy from cancer patients using fragmentation profiles, a stochastic gradient boosting model was used (gbm; see, e.g., Friedman *et al.*, 2001 *Ann Stat* 29:1189-1232; and Friedman *et al.*, 2002 *Comput Stat Data An* 38:367-378). GC-corrected total and short fragment coverage for all 504 bins were centered and scaled for each sample to have mean 0 and unit standard deviation. Additional features included Z-scores for each of the 39 autosomal arms and mitochondrial representation (log₁₀-transformed proportion of reads mapped to the mitochondria). To estimate the prediction error of this approach, 10-fold cross-validation was used as described elsewhere (see, e.g., Efron *et al.*, 1997 *J Am Stat Assoc* 92, 548-560). Feature selection, performed only on the training data in each cross-validation run, removed bins that were highly correlated (correlation > 0.9) or had near zero variance. Stochastic gradient boosted machine learning was implemented using the R package gbm package with parameters n.trees=150, interaction.depth=3, shrinkage=0.1, and n.minobsinside=10. To average over the prediction error from the randomization of patients to folds, the 10-fold cross validation procedure was repeated 10 times. Confidence intervals for sensitivity fixed at 98% and 95% specificity were obtained from 2000 bootstrap replicates.

Prediction model for tumor tissue of origin classification

For samples correctly classified as cancer patients at 90% specificity (n = 174), a separate stochastic gradient boosting model was trained to classify the tissue of origin. To account for the small number of lung samples used for prediction, 18 cfDNA baseline samples from late stage lung cancer patients were included from the monitoring analyses. Performance characteristics of the model were evaluated by 10-fold cross-validation repeated 10 times. This gbm model was trained using the same features as in the cancer classification model. As previously described, features that displayed correlation above 0.9 to each other or had near zero variance were removed within each training dataset during cross-validation.

The tissue class probabilities were averaged across the 10 replicates for each patient and the class with the highest probability was taken as the predicted tissue.

Analyses of nucleosomal DNA from human lymphocytes and cfDNA

From the nuclease treated lymphocytes, fragment sizes were analyzed in 5 Mb bins as described for whole genome cfDNA analyses. A genome-wide map of nucleosome positions was constructed from the nuclease treated lymphocyte cell-lines. This approach identified local biases in the coverage of circulating fragments, indicating a region protected from degradation. A “Window positioning score” (WPS) was used to score each base pair in the genome (see, e.g., Snyder et al., 2016 *Cell* 164:57). Using a sliding window of 60bp centered around each base, the WPS was calculated as the number of fragments completely spanning the window minus the number of fragments with only one end in the window. Since fragments arising from nucleosomes have a median length of 167 bp, a high WPS indicated a possible nucleosomic position. WPS scores were centered at zero using a running median and smoothed using a Kolmogorov-Zurbenko filter (see, e.g., Zurbenko, *The spectral analysis of time series*. North-Holland series in statistics and probability; Elsevier, New York, NY, 1986). For spans of positive WPS between 50 and 450 bp, a nucleosome peak was defined as the set of base pairs with a WPS above the median in that window. The calculation of nucleosome positions for cfDNA from 30 healthy individuals with sequence coverage of 9x was determined in the same manner as for lymphocyte DNA. To ensure that nucleosomes in healthy cfDNA were representative, a consensus track of nucleosomes was defined consisting only of nucleosomes identified in two or more individuals. Median distances between adjacent nucleosomes were calculated from the consensus track.

Monte Carlo simulation of detection sensitivity

A Monte Carlo simulation was used to estimate the probability of detecting a molecule with a tumor-derived alteration. Briefly, 1 million molecules were generated from a multinomial distribution. For a simulation with m alterations, wild-type molecules were simulated with probability p and each of the m tumor alterations were simulated with probability $(1-p)/m$. Next, $g * m$ molecules were sampled randomly with replacement, where g denotes the number of genome equivalents in 1 ml of plasma. If a tumor alteration was sampled s or more times, the sample was classified as cancer-derived. The simulation was

repeated 1000 times, estimating the probability that the *in silico* sample would be correctly classified as cancer by the mean of the cancer indicator. Setting $g = 2000$ and $s = 5$, the number of tumor alterations was varied by powers of 2 from 1 to 256 and the fraction of tumor-derived molecules from 0.0001% to 1%.

5 *Statistical analyses*

All statistical analyses were performed using R version 3.4.3. The R packages caret (version 6.0-79) and gbm (version 2.1-4) were used to implement the classification of healthy versus cancer and tissue of origin. Confidence intervals from the model output were obtained with the pROC (version 1.13) R package (see, e.g., Robin *et al.*, 2011 *BMC bioinformatics* 12:77). Assuming the prevalence of undiagnosed cancer cases in this population is high (1 or 2 cases per 100 healthy), a genomic assay with a specificity of 0.95 and sensitivity of 0.8 would have useful operating characteristics (positive predictive value of 0.25 and negative predictive value near 1). Power calculations suggest that an analysis of more than 200 cancer patients and an approximately equal number of healthy controls, enable 15 an estimation of the sensitivity with a margin of error of 0.06 at the desired specificity of 0.95 or greater.

Data and Code Availability

Sequence data utilized in this study have been deposited at the European Genome-phenome Archive under study accession nos. EGAS00001003611 and EGAS00001002577. 20 Code for analyses is available at github.com/Cancer-Genomics/delfi_scripts.

Results

DELFI allows simultaneous analysis of a large number of abnormalities in cfDNA through genome-wide analysis of fragmentation patterns. The method is based on low coverage whole genome sequencing and analysis of isolated cfDNA. Mapped sequences are 25 analyzed in non-overlapping windows covering the genome. Conceptually, windows may range in size from thousands to millions of bases, resulting in hundreds to thousands of windows in the genome. 5 Mb windows were used for evaluating cfDNA fragmentation patterns as these would provide over 20,000 reads per window even at a limited amount of 1-2x genome coverage. Within each window, the coverage and size distribution of cfDNA

fragments was examined. This approach was used to evaluate the variation of genome-wide fragmentation profiles in healthy and cancer populations (Table 1; Appendix A). The genome-wide pattern from an individual can be compared to reference populations to determine if the pattern is likely healthy or cancer-derived. As genome-wide profiles reveal 5 positional differences associated with specific tissues that may be missed in overall fragment size distributions, these patterns may also indicate the tissue source of cfDNA.

The fragmentation size of cfDNA was focused on as it was found that cancer-derived cfDNA molecules may be more variable in size than cfDNA derived from non-cancer cells. cfDNA fragments from targeted regions that were captured and sequenced at high coverage 10 (43,706 total coverage, 8,044 distinct coverage) from patients with breast, colorectal, lung or ovarian cancer (Table 1 (Appendix A), Table 2 (Appendix B), and Table 3 (Appendix C)) were initially examined. Analyses of loci containing 165 tumor-specific alterations from 81 patients (range of 1-7 alterations per patient) revealed an average absolute difference of 6.5 bp (95% CI, 5.4-7.6 bp) between lengths of median mutant and wild-type cfDNA fragments 15 (Fig. 3, Table 3 (Appendix C)). The median size of mutant cfDNA fragments ranged from 30 bases smaller at chromosome 3 position 41,266,124 to 47 bases larger at chromosome 11 position 108,117,753 than the wild-type sequences at these regions (Table 3; Appendix C). GC content was similar for mutated and non-mutated fragments (Fig. 4a), and there was no correlation between GC content and fragment length (Fig. 4b). Similar analyses of 44 20 germline alterations from 38 patients identified median cfDNA size differences of less than 1 bp between fragment lengths of different alleles (Fig. 5, Table 3 (Appendix C)).

Additionally, 41 alterations related to clonal hematopoiesis were identified through a previous sequence comparison of DNA from plasma, buffy coat, and tumors of the same individuals. Unlike tumor-derived fragments, there were no significant differences between 25 fragments with hematopoietic alterations and wild type fragments (Fig. 6, Table 3 (Appendix C)). Overall, cancer-derived cfDNA fragment lengths were significantly more variable compared to non-cancer cfDNA fragments at certain genomic regions ($p<0.001$, variance ratio test). It was hypothesized that these differences may be due to changes in higher-order chromatin structure as well as other genomic and epigenomic abnormalities in cancer and 30 that cfDNA fragmentation in a position-specific manner could therefore serve as a unique biomarker for cancer detection.

As targeted sequencing only analyzes a limited number of loci, larger-scale genome-wide analyses to detect additional abnormalities in cfDNA fragmentation were investigated. cfDNA was isolated from ~4 ml of plasma from 8 lung cancer patients with stage I-III disease , as well as from 30 healthy individuals (Table 1 (Appendix A), Table 4 (Appendix 5 D), and Table 5 (Appendix E)). A high efficiency approach was used to convert cfDNA to next generation sequencing libraries and performed whole genome sequencing at ~9x coverage (Table 4; Appendix D). Overall cfDNA fragment lengths of healthy individuals were larger, with a median fragment size of 167.3 bp, while patients with cancer had median fragment sizes of 163.8 (p<0.01, Welch's t-test) (Table 5; Appendix E). To examine 10 differences in fragment size and coverage in a position dependent manner across the genome, sequenced fragments were mapped to their genomic origin and fragment lengths were evaluated in 504 windows that were 5 Mb in size, covering ~2.6 Gb of the genome. For each window, the fraction of small cfDNA fragments (100 to 150 bp in length) to larger cfDNA 15 fragments (151 to 220 bp) as well as overall coverage were determined and used to obtain genome-wide fragmentation profiles for each sample.

Healthy individuals had very similar fragmentation profiles throughout the genome (Fig. 7 and Fig. 8). To examine the origins of fragmentation patterns normally observed in cfDNA, nuclei were isolated from elutriated lymphocytes of two healthy individuals and treated with DNA nucleases to obtain nucleosomal DNA fragments. Analyses of cfDNA 20 patterns in observed healthy individuals revealed a high correlation to lymphocyte nucleosomal DNA fragmentation profiles (Fig. 7b and 7d) and nucleosome distances (Fig. 7c and 7f). Median distances between nucleosomes in lymphocytes were correlated to open (A) and closed (B) compartments of lymphoblastoid cells as revealed using the Hi-C method (see, e.g., Lieberman-Aiden *et al.*, 2009 *Science* 326:289-293; and Fortin *et al.*, 2015 25 *Genome Biol* 16:180) for examining the three-dimensional architecture of genomes (Fig. 7c). These analyses suggest that the fragmentation patterns of normal cfDNA are the result of nucleosomal DNA patterns that largely reflect the chromatin structure of normal blood cells.

In contrast to healthy cfDNA, patients with cancer had multiple distinct genomic 30 differences with increases and decreases in fragment sizes at different regions (Fig. 7a and 7b). Similar to our observations from targeted analyses, there was also greater variation in fragment lengths genome-wide for patients with cancer compared to healthy individuals.

To determine whether cfDNA fragment length patterns could be used to distinguish patients with cancer from healthy individuals, genome-wide correlation analyses were performed of the fraction of short to long cfDNA fragments for each sample compared to the median fragment length profile calculated from healthy individuals (Fig. 7a, 7b, and 7e).

- 5 While the profiles of cfDNA fragments were remarkably consistent among healthy individuals (median correlation of 0.99), the median correlation of genome-wide fragment ratios among cancer patients was 0.84 (0.15 lower, 95% CI 0.07-0.50, p<0.001, Wilcoxon rank sum test; Table 5 (Appendix E)). Similar differences were observed when comparing fragmentation profiles of cancer patients to fragmentation profiles or nucleosome distances in
10 healthy lymphocytes (Fig. 7c, 7d, and 7f). To account for potential biases in the fragmentation profiles attributable to GC content, a locally weighted smoother was applied independently to each sample and found that differences in fragmentation profiles between healthy individuals and cancer patients remained after this adjustment (median correlation of cancer patientsto healthy = 0.83) (Table 5; Appendix E).

- 15 Subsampling analyses of whole genome sequence data was performed at 9x coverage from cfDNA of patients with cancer at ~2x, ~1x, ~0.5x, ~0.2x, and ~0.1x genome coverage, and it was determined that altered fragmentation profiles were readily identified even at 0.5x genome coverage (Fig. 9). Based on these observations, whole genome sequencing was performed with coverage of 1-2x to evaluate whether fragmentation profiles may change
20 during the course of targeted therapy in a manner similar to monitoring of sequence alterations. cfDNA from 19 non-small cell lung cancer patients including 5 with partial radiographic response, 8 with stable disease, 4 with progressive disease, and 2 with unmeasurable disease, during the course of anti-EGFR or anti-ERBB2 therapy was evaluated (Table 6; Appendix F). As shown in Fig. 10, the degree of abnormality in the fragmentation profiles during therapy closely matched levels of EGFR or ERBB2 mutant allele fractions as determined using targeted sequencing (Spearman correlation of mutant allele fractions to fragmentation profiles = 0.74). This correlation is remarkable as genome-wide and mutation-based methods are orthogonal and examine different cfDNA alterations that may be suppressed in these patients due to prior therapy. Notably all cases that had progression free
25 survival of six or more months displayed a drop of or had extremely low levels of ctDNA after initiation of therapy as determined by fragmentation profiles, while cases with poor
30

clinical outcome had increases in ctDNA. These results demonstrate the feasibility of fragmentation analyses for detecting the presence of tumor-derived cfDNA, and suggests that such analyses may also be useful for quantitative monitoring of cancer patients during treatment.

5 The fragmentation profiles were examined in the context of known copy number changes in a patient where parallel analyses of tumor tissue were obtained. These analyses demonstrated that altered fragmentation profiles were present in regions of the genome that were copy neutral and that these may be further affected in regions with copy number changes (Fig. 11a and Fig. 12a). Position dependent differences in fragmentation patterns
10 could be used to distinguish cancer-derived cfDNA from healthy cfDNA in these regions (Fig. 12a, b), while overall cfDNA fragment size measurements would have missed such differences (Fig. 12a).

These analyses were extended to an independent cohort of cancer patients and healthy individuals. Whole genome sequencing of cfDNA at 1-2x coverage from a total of 208
15 patients with cancer, including breast (n=54), colorectal (n=27), lung (n=12), ovarian (n=28), pancreatic (n=34), gastric (n=27), or bile duct cancers (n=26), as well as 215 individuals without cancer was performed (Table 1 (Appendix A) and Table 4 (Appendix D)). All cancer patients were treatment naïve and the majority had resectable disease (n=183). After
20 GC adjustment of short and long cfDNA fragment coverage (Fig. 13a), coverage and size characteristics of fragments in windows throughout the genome were examined (Fig. 11b, Table 4 (Appendix D) and Table 7 (Appendix G)). Genome-wide correlations of coverage to GC content were limited and no differences in these correlations between cancer patients and healthy individuals were observed (Fig. 13b). Healthy individuals had highly concordant fragmentation profiles, while patients with cancer had high variability with decreased
25 correlation to the median healthy profile (Table 7; Appendix G). An analysis of the most commonly altered fragmentation windows in the genome among cancer patients revealed a median of 60 affected windows across the cancer types analyzed, highlighting the multitude of position dependent alterations in fragmentation of cfDNA in individuals with cancer (Fig. 11c).

30 To determine if position dependent fragmentation changes can be used to detect individuals with cancer, a gradient tree boosting machine learning model was implemented to

examine whether cfDNA can be categorized as having characteristics of a cancer patient or healthy individual and estimated performance characteristics of this approach by ten-fold cross validation repeated ten times (Figs. 14 and 15). The machine learning model included GC-adjusted short and long fragment coverage characteristics in windows throughout the genome. A machine learning classifier for copy number changes from chromosomal arm dependent features rather than a single score was also developed (Fig. 16a and Table 8 (Appendix H)) and mitochondrial copy number changes were also included (Fig. 16b) as these could also help distinguish cancer from healthy individuals. Using this implementation of DELFI, a score was obtained that could be used to classify patients as healthy or having cancer. 152 of the 208 cancer patients were detected (73% sensitivity, 95% CI 67%-79%) while four of the 215 healthy individuals were misclassified (98% specificity) (Table 9). At a threshold of 95% specificity, 80% of patients with cancer were detected (95% CI, 74%-85%), including 79% of resectable (stage I – III) patients (145 of 183) and 82% of metastatic (stage IV) patients (18 out of 22) (Table 9). Receiver operator characteristic analyses for detection of patients with cancer had an AUC of 0.94 (95% CI 0.92 – 0.96), ranged among cancer types from 0.86 for pancreatic cancer to ≥ 0.99 for lung and ovarian cancers (Figs. 17a and 17b), and had AUCs ≥ 0.92 across all stages (Fig. 18). The DELFI classifier score did not differ with age among either cancer patients or healthy individuals (Table 1; Appendix A).

Table 9. DELFI performance for cancer detection.

	Individuals analyzed	95% specificity			98% specificity			
		Individuals detected	Sensitivity	95% CI	Individuals detected	Sensitivity	95% CI	
Healthy	215	10	-	-	4	-	-	
Cancer	208	166	80%	74%-85%	152	73%	67%-79%	
Type	Breast	54	38	70%	56%-82%	31	57%	43%-71%
	Bile duct	26	23	88%	70%-98%	21	81%	61%-93%
	Colorectal	27	22	81%	62%-94%	19	70%	50%-86%
	Gastric	27	22	81%	62%-94%	22	81%	62%-94%
	Lung	12	12	100%	74%-100%	12	100%	74%-100%
	Ovarian	28	25	89%	72%-98%	25	89%	72%-98%
	Pancreatic	34	24	71%	53%-85%	22	65%	46%-80%
Stage	I	41	30	73%	53%-86%	28	68%	52%-82%
	II	109	85	78%	69%-85%	78	72%	62%-80%
	III	33	30	91%	76%-98%	26	79%	61%-91%
	IV	22	18	82%	60%-95%	17	77%	55%-92%
	0, X	3	3	100%	29%-100%	3	100%	29%-100%

To assess the contribution of fragment size and coverage, chromosome arm copy

5 number, or mitochondrial mapping to the predictive accuracy of the model, the repeated 10-fold cross-validation procedure was implemented to assess performance characteristics of these features in isolation. It was observed that fragment coverage features alone (AUC = 0.94) were nearly identical to the classifier that combined all features (AUC = 0.94) (Fig. 17a). In contrast, analyses of chromosomal copy number changes had lower performance
10 (AUC = 0.88) but were still more predictive than copy number changes based on individual scores (AUC=0.78) or mitochondrial mapping (AUC = 0.72) (Fig. 17a). These results suggest that fragment coverage is the major contributor to our classifier. Including all features in the prediction model may contribute in a complementary fashion for detection of patients with cancer as they can be obtained from the same genome sequence data.

15 As fragmentation profiles reveal regional differences in fragmentation that may differ between tissues, a similar machine learning approach was used to examine whether cfDNA patterns could identify the tissue of origin of these tumors. It was found that this approach had a 61% accuracy (95% CI 53%-67%), including 76% for breast, 44% for bile duct, 71% for colorectal, 67% for gastric, 53% for lung, 48% for ovarian, and 50% for pancreatic

cancers (Fig. 19, Table 10). The accuracy increased to 75% (95% CI 69%-81%) when considering assigning patients with abnormal cfDNA to one of two sites of origin (Table 10). For all tumor types, the classification of the tissue of origin by DELFI was significantly higher than determined by random assignment ($p<0.01$, binomial test, Table 10).

Table 10. DELFI tissue of origin prediction

Cancer Type	Patients Detected*	Top Prediction		Top Two Predictions		Random Assignment	
		Patients	Accuracy (95% CI)	Patients	Accuracy (95% CI)	Patients	Accuracy
Breast	42	32	76% (61%-88%)	38	91% (77%-97%)	9	22%
Bile Duct	23	10	44% (23%-66%)	15	65% (43%-84%)	3	12%
Colorectal	24	17	71% (49%-87%)	19	79% (58%-93%)	3	12%
Gastric	24	16	67% (45%-84%)	19	79% (58%-93%)	3	12%
Lung	30	16	53% (34%-72%)	23	77% (58%-90%)	2	6%
Ovarian	27	13	48% (29%-68%)	16	59% (38%-78%)	4	14%
Pancreatic	24	12	50% (29%-71%)	16	67% (45%-84%)	3	12%
Total	194	116	61% (53%-67%)	146	75% (69%-81%)	26	13%

*Patients detected are based on DELFI detection at 90% specificity. Lung cohort includes additional lung cancer patients with prior therapy.

As cancer-specific sequence alterations can be used to identify patients with cancer, it was evaluated whether combining DELFI with this approach could increase the sensitivity of cancer detection (Fig. 20). An analysis of cfDNA from a subset of the treatment naïve cancer 5 patients using both DELFI and targeted sequencing revealed that 82% (103 of 126) of patients had fragmentation profile alterations, while 66% (83 of 126) had sequence alterations. Over 89% of cases with mutant allele fractions >1% were detected by DELFI while for cases with mutant allele fractions <1% the fraction detected by DELFI was 80%, including for cases that were undetectable using targeted sequencing (Table 7; Appendix G).
10 When these approaches were used together, the combined sensitivity of detection increased to 91% (115 of 126 patients) with a specificity of 98% (Fig. 20).

Overall, genome-wide cfDNA fragmentation profiles are different between cancer patients and healthy individuals. The variability in fragment lengths and coverage in a position dependent manner throughout the genome may explain the apparently contradictory 15 observations of previous analyses of cfDNA at specific loci or of overall fragment sizes. In patients with cancer, heterogeneous fragmentation patterns in cfDNA appear to be a result of mixtures of nucleosomal DNA from both blood and neoplastic cells. These studies provide a method for simultaneous analysis of tens to potentially hundreds of tumor-specific abnormalities from minute amounts of cfDNA, overcoming a limitation that has precluded 20 the possibility of more sensitive analyses of cfDNA. DELFI analyses detected a higher fraction of cancer patients than previous cfDNA analysis methods that have focused on sequence or overall fragmentation sizes (see, e.g., Phallen *et al.*, 2017 *Sci Transl Med* 9:eaan2415; Cohen *et al.*, 2018 *Science* 359:926; Newman *et al.*, 2014 *Nat Med* 20:548; Bettegowda *et al.*, 2014 *Sci Transl Med* 6:224ra24; Newman *et al.*, 2016 *Nat Biotechnol* 25 34:547). As demonstrated in this Example, combining DELFI with analyses of other cfDNA alterations may further increase the sensitivity of detection. As fragmentation profiles appear related to nucleosomal DNA patterns, DELFI may be used for determining the primary source of tumor-derived cfDNA. The identification of the source of circulating tumor DNA in over half of patients analyzed may be further improved by including clinical 30 characteristics, other biomarkers, including methylation changes, and additional diagnostic approaches (Ruibal Morell, 1992 *The International journal of biological markers* 7:160;

Galli et al., 2013 *Clinical chemistry and laboratory medicine* 51:1369; Sikaris, 2011 *Heart, lung & circulation* 20:634; Cohen et al., 2018 *Science* 359:926). Finally, this approach requires only a small amount of whole genome sequencing, without the need for deep sequencing typical of approaches that focus on specific alterations. The performance
5 characteristics and limited amount of sequencing needed for DELFI suggests that our approach could be broadly applied for screening and management of patients with cancer.

These results demonstrate that genome-wide cfDNA fragmentation profiles are different between cancer patients and healthy individuals. As such, cfDNA fragmentation profiles can have important implications for future research and applications of non-invasive
10 approaches for detection of human cancer.

Other Embodiments

It is to be understood that while the invention has been described in conjunction with the detailed description thereof, the foregoing description is intended to illustrate and not limit the scope of the invention, which is defined by the scope of the appended claims. Other
15 aspects, advantages, and modifications are within the scope of the following claims.

APPENDIX A: Table 1. Summary of patients and samples analyzed

Patient	Patient Type	Sample Type	Timepoint	Histopathological Diagnosis				Referred to Diagnosis	Location of biopsies at Diagnosis	Volume of Plasma (ml)	cDNA (ng/ml)	Extracted DNA	Whole Genome Library	Fragment Profile Analysis	
				Age at Diagnosis	Gender	Stage	TNM Staging								
CGR1439	Healthy	cDNA	Preoperative treatment naïve	73	F	IA	IA	NA	NA	NA	4.6	14.73	NA	NA	NA
CGR1439	Healthy	cDNA	Preoperative treatment naïve	50	M	IA	IA	NA	NA	NA	4.6	12.14	Y	Y	N
CGR1439	Healthy	cDNA, surgical resectate	Preoperative treatment naïve	58	M	IA	IA	NA	NA	NA	4.5	8.88	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	57	F	IA	IA	NA	NA	NA	4.5	8.37	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	54	M	IA	IA	NA	NA	NA	4.5	8.39	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.5	5.27	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.4	3.79	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.4	9.56	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.4	5.46	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.4	20.31	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	58	M	IA	IA	NA	NA	NA	4.3	12.01	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	62	M	IA	IA	NA	NA	NA	4.4	4.79	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	64	M	IA	IA	NA	NA	NA	4.4	7.70	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	64	M	IA	IA	NA	NA	NA	4.4	6.26	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	64	M	IA	IA	NA	NA	NA	4.4	13.01	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	64	M	IA	IA	NA	NA	NA	4.4	11.13	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	57	F	IA	IA	NA	NA	NA	4.4	2.05	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	64	M	IA	IA	NA	NA	NA	4.3	4.41	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	59	M	IA	IA	NA	NA	NA	4.2	6.38	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	59	M	IA	IA	NA	NA	NA	4.2	7.28	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	57	M	IA	IA	NA	NA	NA	4.2	5.46	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	75	F	IA	IA	NA	NA	NA	4.0	13.20	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	50	M	IA	IA	NA	NA	NA	4.0	10.20	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.0	5.18	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.0	3.89	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.0	4.4	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.0	6.91	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	50	F	IA	IA	NA	NA	NA	4.1	3.20	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.1	5.55	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	50	F	IA	IA	NA	NA	NA	4.5	8.18	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	50	F	IA	IA	NA	NA	NA	4.5	2.18	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.5	5.06	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.5	10.20	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.5	11.73	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	50	F	IA	IA	NA	NA	NA	4.5	10.98	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.5	10.36	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.5	10.17	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	54	M	IA	IA	NA	NA	NA	4.6	14.30	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	51	M	IA	IA	NA	NA	NA	4.6	12.32	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.6	5.42	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.6	2.85	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	52	F	IA	IA	NA	NA	NA	4.7	1.86	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	70	M	IA	IA	NA	NA	NA	4.7	12.87	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	62	M	IA	IA	NA	NA	NA	4.6	2.42	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.6	2.34	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	65	M	IA	IA	NA	NA	NA	4.6	8.95	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	65	M	IA	IA	NA	NA	NA	4.6	6.15	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	63	M	IA	IA	NA	NA	NA	4.6	3.44	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	63	M	IA	IA	NA	NA	NA	4.6	18.97	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	54	M	IA	IA	NA	NA	NA	4.6	8.50	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	52	F	IA	IA	NA	NA	NA	4.6	15.23	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.6	4.2	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.6	5.98	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.6	6.33	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.6	4.85	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.6	4.29	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.6	3.44	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	M	IA	IA	NA	NA	NA	4.6	12.12	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	65	M	IA	IA	NA	NA	NA	4.6	3.65	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	65	M	IA	IA	NA	NA	NA	4.6	16.62	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	10.53	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	12.87	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	7.42	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	7.30	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	3.77	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	10.85	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	5.62	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	4.4	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	2.32	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	4.38	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	3.24	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	3.97	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	5.29	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	3.11	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	5.92	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	16.94	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	6.52	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	8.25	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	5.24	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	2.27	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	2.42	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	4.75	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	10.17	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	2.95	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	25.15	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	7.28	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	6.18	Y	Y	N
CGR1439	Healthy	cDNA	Preoperative treatment naïve	55	F	IA	IA	NA	NA	NA	4.6	15.21	Y	Y	N

Patient	Patient Type	Sample Type	Clinical Data							Pathological Findings						Treatment Response		
			Age at Diagnosis	Timepoint	Gender	Stage	TNM Staging	Site of Primary Tumor	Histopathological Diagnoses	Degree of Differentiation	Location of Metastases & Diagnosis	Volume of Plasma (ml)	cT/tN/tM Input (mg/ml)	Estimated cT/tN/tM Input (mg)	Whole Genome Analysis	Targeted Profiling	Fragment Profiling	Tag-based Analysis
CSP-PA117	Bile Duct Cancer	ctDNA	66	I	NA	I	NA	Itra-Paracite/Bile Duct	Intra-Apocrine/Bile Duct	3.4	2.25	Y	N	Y	Y	Y	Y	
CSP-PA118	Bile Duct Cancer	ctDNA	62	I	NA	I	NA	Itra-Duct	Intra-Duct	3.6	9.05	32.85	N	Y	Y	Y	N	
CSP-PA122	Bile Duct Cancer	ctDNA	63	I	NA	I	NA	Itra-Duct	Intra-Duct	3.6	6.64	27.17	N	Y	Y	Y	N	
CSP-PA124	Bile Duct Cancer	ctDNA	58	I	NA	I	NA	Itra-Duct	Intra-Duct	2.7	8.31	23.11	N	Y	Y	Y	N	
CSP-PA125	Bile Duct Cancer	ctDNA	50	I	NA	I	NA	Itra-Duct	Intra-Duct	4.2	6.56	26.31	N	Y	Y	Y	N	
CSP-PA126	Bile Duct Cancer	ctDNA	71	I	NA	I	NA	Itra-Duct	Intra-Duct	3.0	26.60	79.80	N	Y	Y	Y	N	
CSP-PA127	Bile Duct Cancer	ctDNA	67	I	NA	I	NA	Itra-Duct	Intra-Duct	3.9	5.91	22.67	N	Y	Y	Y	N	
CSP-PA128	Bile Duct Cancer	ctDNA	55	I	NA	I	NA	Itra-Duct	Intra-Duct	4.6	27.97	127.97	N	Y	Y	Y	N	
CSP-PA129	Bile Duct Cancer	ctDNA	62	I	NA	I	NA	Itra-Duct	Intra-Duct	4.0	4.34	16.36	N	Y	Y	Y	N	
CSP-PA130	Bile Duct Cancer	ctDNA	60	I	NA	I	NA	Itra-Duct	Intra-Duct	4.7	28.34	126.50	N	Y	Y	Y	N	
CSP-PA131	Bile Duct Cancer	ctDNA	52	I	NA	I	NA	Itra-Duct	Intra-Duct	3.5	62.95	32.95	N	Y	Y	Y	N	
CSP-PA132	Bile Duct Cancer	ctDNA	71	I	NA	I	NA	Itra-Duct	Intra-Duct	3.9	52.98	95.48	N	Y	Y	Y	N	
CSP-PA133	Bile Duct Cancer	ctDNA	68	I	NA	I	NA	Itra-Duct	Intra-Duct	4.1	24.22	4.22	N	Y	Y	Y	N	
CSP-PA135	Bile Duct Cancer	ctDNA	89	I	NA	I	NA	Itra-Duct	Intra-Duct	3.9	23.23	23.23	N	Y	Y	Y	N	
CSP-PA136	Bile Duct Cancer	ctDNA	55	I	NA	I	NA	Itra-Duct	Intra-Duct	4.0	5.75	23.75	N	Y	Y	Y	N	
CSP-PA137	Bile Duct Cancer	ctDNA	100	I	NA	I	NA	Itra-Duct	Intra-Duct	4.0	14.89	57.59	N	Y	Y	Y	N	
CSP-PA138	Bile Duct Cancer	ctDNA	60	I	NA	I	NA	Itra-Duct	Intra-Duct	4.0	1.30	5.00	N	Y	Y	Y	N	
CSP-PA14	Pancreatic Cancer	ctDNA	68	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	2.8	44.64	123.92	N	Y	Y	Y	N	
CSP-PA140	Bile Duct Cancer	ctDNA	76	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	1.92	38.48	N	Y	Y	Y	N	
CSP-PA141	Bile Duct Cancer	ctDNA	70	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	25.72	78.16	N	Y	Y	Y	N	
CSP-PA143	Pancreatic Cancer	ctDNA	88	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.5	7.54	34.74	N	Y	Y	Y	N	
CSP-PA145	Bile Duct Cancer	ctDNA	72	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.9	16.48	50.48	N	Y	Y	Y	N	
CSP-PA146	Pancreatic Cancer	ctDNA	42	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	13.32	53.20	N	Y	Y	Y	N	
CSP-PA147	Bile Duct Cancer	ctDNA	58	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	12.38	49.52	N	Y	Y	Y	N	
CSP-PA148	Pancreatic Cancer	ctDNA	65	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	1.92	39.00	N	Y	Y	Y	N	
CSP-PA149	Bile Duct Cancer	ctDNA	75	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	1.92	39.00	N	Y	Y	Y	N	
CSP-PA150	Pancreatic Cancer	ctDNA	67	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	16.82	67.28	N	Y	Y	Y	N	
CSP-PA152	Pancreatic Cancer	ctDNA	54	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	8.71	34.57	N	Y	Y	Y	N	
CSP-PA156	Pancreatic Cancer	ctDNA	60	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	6.97	27.85	N	Y	Y	Y	N	
CSP-PA158	Bile Duct Cancer	ctDNA	78	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	3.41	16.33	N	Y	Y	Y	N	
CSP-PA159	Pancreatic Cancer	ctDNA	67	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	0.74	18.00	N	Y	Y	Y	N	
CSP-PA160	Pancreatic Cancer	ctDNA	75	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	6.01	24.03	N	Y	Y	Y	N	
CSP-PA161	Pancreatic Cancer	ctDNA	63	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	3.16	12.50	N	Y	Y	Y	N	
CSP-PA162	Pancreatic Cancer	ctDNA	67	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	21.83	87.32	N	Y	Y	Y	N	
CSP-PA163	Pancreatic Cancer	ctDNA	58	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	5.75	23.13	N	Y	Y	Y	N	
CSP-PA164	Pancreatic Cancer	ctDNA	85	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	11.73	46.08	N	Y	Y	Y	N	
CSP-PA165	Pancreatic Cancer	ctDNA	73	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	4.78	23.92	N	Y	Y	Y	N	
CSP-PA166	Pancreatic Cancer	ctDNA	79	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	9.72	38.88	N	Y	Y	Y	N	
CSP-PA167	Pancreatic Cancer	ctDNA	67	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	1.72	7.68	N	Y	Y	Y	N	
CSP-PA168	Pancreatic Cancer	ctDNA	75	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	2.0	35.97	71.97	N	Y	Y	Y	N	
CSP-PA169	Pancreatic Cancer	ctDNA	63	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	2.2	39.97	83.94	N	Y	Y	Y	N	
CSP-PA170	Pancreatic Cancer	ctDNA	67	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	2.5	5.26	12.98	N	Y	Y	Y	N	
CSP-PA171	Pancreatic Cancer	ctDNA	55	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	14.48	57.48	N	Y	Y	Y	N	
CSP-PA172	Pancreatic Cancer	ctDNA	74	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	6.87	26.61	N	Y	Y	Y	N	
CSP-PA173	Pancreatic Cancer	ctDNA	54	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	3.41	13.63	N	Y	Y	Y	N	
CSP-PA174	Pancreatic Cancer	ctDNA	72	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.2	9.72	39.24	N	Y	Y	Y	N	
CSP-PA175	Pancreatic Cancer	ctDNA	60	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	4.74	19.76	N	Y	Y	Y	N	
CSP-PA176	Pancreatic Cancer	ctDNA	64	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	11.49	34.97	N	Y	Y	Y	N	
CSP-PA177	Pancreatic Cancer	ctDNA	72	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	25.88	77.64	N	Y	Y	Y	N	
CSP-PA178	Pancreatic Cancer	ctDNA	71	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	4.96	19.84	N	Y	Y	Y	N	
CSP-PA179	Pancreatic Cancer	ctDNA	60	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	2.5	23.19	57.97	N	Y	Y	Y	N	
CSP-PA180	Pancreatic Cancer	ctDNA	46	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	14.48	44.48	N	Y	Y	Y	N	
CSP-PA181	Pancreatic Cancer	ctDNA	74	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	6.87	26.61	N	Y	Y	Y	N	
CSP-PA182	Pancreatic Cancer	ctDNA	59	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	3.41	13.63	N	Y	Y	Y	N	
CSP-PA183	Pancreatic Cancer	ctDNA	73	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.2	9.72	39.24	N	Y	Y	Y	N	
CSP-PA184	Pancreatic Cancer	ctDNA	55	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	2.0	35.97	83.94	N	Y	Y	Y	N	
CSP-PA185	Pancreatic Cancer	ctDNA	70	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	2.2	41.67	93.34	N	Y	Y	Y	N	
CSP-PA186	Pancreatic Cancer	ctDNA	64	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	2.5	5.24	12.98	N	Y	Y	Y	N	
CSP-PA187	Pancreatic Cancer	ctDNA	60	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	56.20	168.60	N	Y	Y	Y	N	
CSP-PA188	Pancreatic Cancer	ctDNA	77	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	41.67	93.34	N	Y	Y	Y	N	
CSP-PA189	Pancreatic Cancer	ctDNA	60	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	25.88	77.64	N	Y	Y	Y	N	
CSP-PA190	Pancreatic Cancer	ctDNA	77	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	3.41	13.63	N	Y	Y	Y	N	
CSP-PA191	Gastric Cancer	ctDNA	88	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	10.35	31.05	N	Y	Y	Y	N	
CSP-PA192	Gastric Cancer	ctDNA	72	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	24.33	73.00	N	Y	Y	Y	N	
CSP-PA193	Gastric Cancer	ctDNA	84	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	4.22	12.61	N	Y	Y	Y	N	
CSP-PA194	Gastric Cancer	ctDNA	82	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	11.49	34.97	N	Y	Y	Y	N	
CSP-PA195	Gastric Cancer	ctDNA	97	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.5	57.11	162.33	N	Y	Y	Y	N	
CSP-PA196	Gastric Cancer	ctDNA	78	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	48.89	117.22	N	Y	Y	Y	N	
CSP-PA197	Gastric Cancer	ctDNA	75	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	8.53	21.33	N	Y	Y	Y	N	
CSP-PA198	Gastric Cancer	ctDNA	70	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	3.57	9.41	N	Y	Y	Y	N	
CSP-PA199	Gastric Cancer	ctDNA	76	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	7.44	18.68	N	Y	Y	Y	N	
CSP-PA200	Gastric Cancer	ctDNA	74	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	5.56	13.92	N	Y	Y	Y	N	
CSP-PA201	Gastric Cancer	ctDNA	73	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	4.55	11.38	N	Y	Y	Y	N	
CSP-PA202	Gastric Cancer	ctDNA	78	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	8.75	21.88	N	Y	Y	Y	N	
CSP-PA203	Gastric Cancer	ctDNA	70	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	15.82	40.00	N	Y	Y	Y	N	
CSP-PA204	Gastric Cancer	ctDNA	79	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	3.57	9.41	N	Y	Y	Y	N	
CSP-PA205	Gastric Cancer	ctDNA	76	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	23.91	73.00	N	Y	Y	Y	N	
CSP-PA206	Gastric Cancer	ctDNA	74	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	15.82	40.00	N	Y	Y	Y	N	
CSP-PA207	Gastric Cancer	ctDNA	75	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	3.57	9.41	N	Y	Y	Y	N	
CSP-PA208	Gastric Cancer	ctDNA	72	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	4.55	11.38	N	Y	Y	Y	N	
CSP-PA209	Gastric Cancer	ctDNA	73	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	4.0	8.75	21.88	N	Y	Y	Y	N	
CSP-PA210	Gastric Cancer	ctDNA	78	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	15.82	40.00	N	Y	Y	Y	N	
CSP-PA211	Gastric Cancer	ctDNA	76	I	NA	I	NA	Itra-Hepatocytic/Ect	Itra-Hepatocytic/Ect	3.0	3.57	9.41	N	Y	Y	Y	N	

2) A complete date not available or not applicable for heating individual

APPENDIX B: Table 2. Summary of targeted cfDNA analyses

Patient	Patient Type	Timepoint	Fragment Profile	Mutation Analysis	Read Length	Bases in Target Region	Bases Mapped to Genome	Target Regions	Percent Mapped to Target Regions	Total Coverage	Distinct Coverage
CGCRC291	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7501485600	3771369756	50%	43345	10358
CGCRC292	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	6736035200	3098869573	46%	36448	8603
CGCRC293	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	6300244000	2818734206	45%	33117	5953
CGCRC294	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7756672600	3911796750	50%	46016	12071
CGCRC295	Colorectal Cancer	Preoperative, Treatment naïve	Y	N	100	80930	8240602000	3478059753	42%	40787	5826
CGCRC296	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	5718856500	2380545356	51%	33912	10180
CGCRC297	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7550261600	371722432	49%	43545	5870
CGCRC298	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	12501036400	6096383764	49%	71196	9617
CGCRC299	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7812662900	4121568690	53%	48098	10338
CGCRC300	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	864090300	396295136	46%	46354	5756
CGCRC301	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7538758100	3695480348	49%	43024	6618
CGCRC302	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8573658300	4349420574	51%	51006	13793
CGCRC303	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	5224046400	2505714343	48%	29365	8372
CGCRC304	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	5716212600	2942170530	51%	34462	10208
CGCRC305	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7213884100	3726963480	52%	45516	8589
CGCRC306	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7075579700	3562441899	50%	41507	7372
CGCRC307	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7572887100	3492191519	46%	40793	9680
CGCRC308	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7945738000	3895908386	49%	45224	11809
CGCRC309	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8457934800	3920793611	46%	47336	10739
CGCRC310	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	904735680500	4678812441	52%	54713	11139
CGCRC311	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	681281627100	3276833864	50%	38234	6044
CGCRC312	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7683294200	3316719187	43%	38652	4622
CGCRC313	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	5874099230	2886148722	49%	33821	6506
CGCRC314	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	6883148860	3382767492	49%	39414	6664
CGCRC315	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7492752600	3775556651	50%	44034	8666
CGCRC316	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8168472600	5533867153	52%	64693	14296
CGCRC317	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7086877600	368949216	52%	43538	10344
CGCRC318	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	688041100	3326367413	48%	39077	11571
CGCRC319	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	7485342900	39825767483	53%	47327	10502
CGCRC320	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	70587653200	408688	50%	10198	
CGCRC321	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	1068472600	3633396399	50%	43065	
CGCRC322	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	720269500	3758323705	52%	44580	3243
CGCRC323	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8767144700	4199126527	48%	49731	6336
CGCRC324	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	777169160	3944576280	51%	46518	5014
CGCRC325	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	4084901201	48308	51%	6151	
CGCRC326	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8657346400	4334410573	50%	51390	7551
CGCRC327	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7399611760	3800666199	51%	45083	8092
CGCRC328	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8029493700	4179383804	52%	49380	5631
CGCRC329	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7938863600	4085556110	52%	48397	3808
CGCRC330	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7214688600	3706543098	51%	43805	3014
CGCRC331	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8857346400	4334410573	50%	51390	
CGCRC332	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7399611760	3800666199	51%	44580	
CGCRC333	Colorectal Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	6942167630	3028232737	48%	36828	
CGCRC334	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	8182668200	2383173431	29%	28233	7973
CGCRC335	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	7448272600	3925056341	53%	46679	5582
CGCRC336	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	8863159200	2968208527	42%	4141	
CGCRC337	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	8478811600	3425540889	40%	40328	
CGCRC338	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	7434818400	30467514994	51%	36828	
CGCRC339	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	7365646100	3020305300	50%	43162	5205
CGCRC340	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	7884655000	3336939252	42%	39587	
CGCRC341	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	7551674450	2968209912	51%	35490	
CGCRC342	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	693451600	3633145275	51%	41908	
CGCRC343	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	738227030	2379682897	30%	4858	
CGCRC344	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	6013175650	30467514994	51%	36127	
CGCRC345	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	6013454600	3020305300	50%	38513	
CGCRC346	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	7551674450	2968209912	51%	4141	
CGCRC347	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	7399611760	3425540889	40%	43379	
CGCRC348	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	7434818400	30467514994	51%	36828	
CGCRC349	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	7365646100	3020305300	50%	4259	
CGCRC350	Colorectal Cancer	Preoperative, Treatment naïve	N	N	100	80930	7884655000	3336939252	44%	37992	

Patient	Patient Type	Timepoint	Fragment Profile Analysis	Mutation Analysis	Read Length	Bases in Target Region	Bases Mapped to Genome	Percent Mapped to Target Regions	Total Coverage	Distinct Coverage
CGCRC359	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	7818667760	425110101	5040	2566
CGCRC367	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	8880243200	33630633597	51%	38844
CGCRC368	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	4101646000	4101646000	51%	48636
CGCRC370	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	6940330100	3189364121	46%	38153
CGCRC373	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	6587621730	3120088035	47%	37234
CGCRC376	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	6779783100	3162416807	47%	37735
CGCRC377	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	6716339200	3131415570	47%	37160
CGCRC378	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	6523963950	2411096720	37%	28728
CGCRC379	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	6986252100	3371981103	48%	39999
CGCRC380	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	7097496300	2710244446	38%	32020
CGCRC381	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	69851936100	3287050581	47%	9357
CGCRC382	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	6959048700	2562325859	37%	30040
CGCRC384	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	7012798600	3293884583	47%	38158
CGCRC385	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	7542017830	336570505	45%	38884
CGCRC386	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	6876059650	3064412286	45%	38431
CGCRC387	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	7398647470	3047254560	41%	36141
CGCRC388	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	6852692800	3137284885	48%	37285
CGCRC389	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	6851206500	3102100941	47%	36553
CGCRC390	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	7280616800	3376687585	47%	38686
CGCRC391	Colorectal Cancer	Preoperative, Treatment naïve	N	Y	100	80930	8883624600	3202877861	47%	37978
CGLU315	Lung Cancer	Pre-treatment, Day -53	Y	N	100	80930	7863415600	1991331171	25%	5029
CGLU316	Lung Cancer	Pre-treatment, Day -53	Y	N	100	80930	7502591600	3137284885	50%	38655
CGLU316	Lung Cancer	Pre-treatment, Day -53	Y	N	100	80930	6852692800	3137284885	48%	37285
CGLU316	Lung Cancer	Pre-treatment, Day -53	Y	N	100	80930	6851206500	3102100941	47%	36553
CGLU316	Lung Cancer	Pre-treatment, Day -53	Y	N	100	80930	7280616800	3376687585	47%	38686
CGLU344	Lung Cancer	Pre-treatment, Day -21	Y	N	100	80930	7489836503	3202877861	45%	37978
CGLU344	Lung Cancer	Pre-treatment, Day -21	Y	N	100	80930	7842919800	1147703178	25%	23601
CGLU344	Lung Cancer	Pre-treatment, Day -21	Y	N	100	80930	5638083100	2291108925	48%	44262
CGLU344	Lung Cancer	Pre-treatment, Day -21	Y	N	100	80930	765823151600	3187059470	48%	37813
CGLU344	Lung Cancer	Pre-treatment, Day -21	Y	N	100	80930	6847281850	1947630979	30%	23094
CGLU344	Lung Cancer	Pre-treatment, Day -21	Y	N	100	80930	6151628500	1462448715	21%	32452
CGLU344	Lung Cancer	Pre-treatment, Day -21	Y	N	100	80930	6934701700	2124660124	31%	60633
CGLU344	Lung Cancer	Pre-treatment, Day -21	Y	N	100	80930	5638083100	31621956578	45%	37519
CGLU344	Lung Cancer	Pre-treatment, Day -21	Y	N	100	80930	765823151600	3187059470	48%	37813
CGLU369	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	15109
CGLU369	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU369	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU369	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7767146300	2124660124	31%	60633
CGLU369	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7193099100	2291108925	39%	27067
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900	1947630979	30%	23094
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	6934701700	1462448715	21%	32452
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7372274529	3722724529	48%	3471
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7680245300	1271467382	18%	2354
CGLU373	Lung Cancer	Pre-treatment, Day -2	Y	N	100	80930	7078131900			

Patient	Patient Type	Timepoint	Fragment Profile Analysis	Mutation Analysis	Read Length	Bases in Target Region	Bases Mapped to Genome	Percent Mapped to Target Regions	Total Coverage	Distinct Coverage
CGPLBR70	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7251160700	3641330706	43203	6884
CGPLBR71	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8515402600	44963956381	53340	6805
CGPLBR72	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	856946800	4369761697	51%	52081
CGPLBR73	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	795993200	4063933538	50%	47555
CGPLBR74	Breast Cancer	Preoperative, Treatment naïve	Y	N	100	80930	882463640	4063900599	48%	48252
CGPLBR75	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8265079100	3965098865	48%	46955
CGPLBR76	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7774735200	389352420	50%	46192
CGPLBR77	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	757279760	3265963429	43%	38568
CGPLBR80	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	68476500	3147476693	46%	37201
CGPLBR82	Breast Cancer	Preoperative, Treatment naïve	N	Y	100	80930	8236705200	4170465005	51%	49361
CGPLBR83	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7434668100	367685019	49%	43528
CGPLBR86	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7616262500	3644791327	48%	43490
CGPLBR87	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	61940121300	3004882010	49%	5306
CGPLBR88	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	6011567200	2847926237	47%	38945
CGPLBR91	Breast Cancer	Preoperative, Treatment naïve	N	Y	100	80930	7192457700	3480203404	48%	41570
CGPLBR92	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7673881600	3600279233	47%	42975
CGPLBR93	Breast Cancer	Preoperative, Treatment naïve	N	Y	100	80930	7656717800	3986713397	53%	47866
CGPLBR96	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	6297446700	2463064757	39%	7937
CGPLBR97	Breast Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7114921600	3557089027	50%	42488
CGPLH35	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	6919126300	2317878764	33%	1989
CGPLH36	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	6069823400	2038548115	33%	22719
CGPLH37	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	5657707000	1935301929	35%	21673
CGPLH42	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	57932045400	2388036949	41%	27197
CGPLH43	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	5658321700	2017813329	36%	23228
CGPLH45	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	8465953200	2770176078	33%	3114
CGPLH46	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	5083711600	1899395790	37%	21821
CGPLH47	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	6016388600	2062392156	34%	23459
CGPLH48	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	4958455000	1809825992	36%	20702
CGPLH49	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	7953812200	2511365904	32%	1650
CGPLH50	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	6989467600	2561288100	37%	32829
CGPLH51	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	7862091396	2525091320	32%	1678
CGPLH52	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	6939536500	2397922659	35%	2312
CGPLH54	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	19511934700	2290823134	22%	1698
CGPLH55	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	9912569200	2521962244	25%	27096
CGPLH56	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	5777591930	2023874863	35%	3114
CGPLH57	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	9234604600	14393265244	16%	16543
CGPLH59	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	972602400	2987875484	31%	2501
CGPLH63	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	19511934700	2290823134	22%	27175
CGPLH64	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	9912569200	2521962244	25%	3161
CGPLH75	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	3476444000	1505778480	44%	17805
CGPLH76	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	7499116100	3685762725	49%	4643
CGPLH77	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	6512468400	2537359346	39%	3131
CGPLH78	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	8629495000	2521574759	29%	46316
CGPLH79	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	5438652600	996196802	18%	11477
CGPLH80	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	3476444000	1505778480	44%	3016
CGPLH81	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	664268800	31123681850	47%	45507
CGPLH82	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	7741705900	394770591	51%	4414
CGPLH83	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	6516189500	1447503106	51%	46620
CGPLH84	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	6957686000	3946069580	21%	3078
CGPLH86	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	8326493200	3910639227	50%	6514
CGPLH90	Healthy	Preoperative, Treatment naïve	N	Y	100	80930	8664194700	4470145931	52%	3847
Lung Cancer	Pre-treatment, Day -2	Preoperative, Treatment naïve	N	Y	100	80930	5658646100	172161895	30%	6025
Lung Cancer	Pre-treatment, Day -2	Preoperative, Treatment naïve	N	Y	100	80930	6193049700	25336569840	41%	3078
Lung Cancer	Pre-treatment, Day -2	Preoperative, Treatment naïve	N	Y	100	80930	5854396500	1194237002	20%	14331
Lung Cancer	Pre-treatment, Day -2	Preoperative, Treatment naïve	N	Y	100	80930	5086197700	1373505586	27%	16480
Lung Cancer	Pre-treatment, Day -38	Preoperative, Treatment naïve	N	Y	100	80930	8668655700	3960731089	46%	3146

Patient	Patient Type	Timepoint	Fragment Profile Analysis	Mutation Analysis	Read Length	Bases in Target Region	Bases Mapped to Genome	Percent Mapped to Target Regions	Total Coverage	Distinct Coverage
CGPLLU14	Lung Cancer	Pre-treatment, Day -16	N	Y	100	80930	82710435830	4105092736	50%	50152 4497
CGPLLU14	Lung Cancer	Pre-treatment, Day -3	N	Y	100	80930	7149069200	3405754720	48%	40382 6170
CGPLLU14	Lung Cancer	Post-treatment, Day 0	N	Y	100	80930	32855332200	3285504484	50%	39044 4061
CGPLLU14	Lung Cancer	Post-treatment, Day 33	N	Y	100	80930	7140378800	3464286558	47%	41108 4259
CGPLLU14	Lung Cancer	Post-treatment, Day 7	N	Y	100	80930	7530190730	3752054349	50%	48839 2469
CGPLLU14	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8176827400	4216576524	48%	49370 16771
CGPLLU14	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8568444200	41950393049	49%	49084 6968
CGPLLU14	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7416360600	3530746046	48%	47302 4691
CGPLLU14	Lung Cancer	Preoperative, Treatment naïve	N	Y	100	80930	7789487000	3260139772	42%	36568 12229
CGPLLU161	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7625462000	3470147687	46%	49118 10099
CGPLLU162	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8019293200	3946533983	49%	46471 12108
CGPLLU163	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8110303800	3592746235	44%	42161 6947
CGPLLU164	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	838514650	4147501817	49%	48770 6996
CGPLLU165	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	765683000	3868237773	50%	45625 9711
CGPLLU168	Lung Cancer	Preoperative, Treatment naïve	N	Y	100	80930	9376353000	4800407624	51%	56547 10261
CGPLLU169	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7446194460	3067532518	41%	36321 6137
CGPLLU174	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	852324200	4002541569	47%	47084 7862
CGPLLU175	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8143905000	4054958929	50%	47708 5588
CGPLLU176	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8421611300	4197108809	50%	49476 8780
CGPLLU177	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8483124700	41693571469	49%	46580 6445
CGPLLU178	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7774958738	3304945738	43%	38768 6862
CGPLLU179	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8192813800	3337532475	48%	46498 6568
CGPLLU180	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7936779200	3062397881	39%	36381 5388
CGPLLU197	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	717524730	3545719100	49%	42008 6817
CGPLLU198	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	884012860	3472820569	50%	40670 7951
CGPLLU202	Lung Cancer	Preoperative, Treatment naïve	N	Y	100	80930	7463874980	3736258011	50%	44500 9917
CGPLLU203	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7450261600	3703545153	50%	44317 6856
CGPLLU204	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	9225429100	4350573991	47%	51627 9810
CGPLLU205	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	7397314650	3635240205	49%	43016 7124
CGPLLU206	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	713043800	3736258011	52%	44291 8499
CGPLLU207	Lung Cancer	Preoperative, Treatment naïve	N	Y	100	80930	7346976450	3865814032	52%	45782 6940
CGPLLU208	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	6723337500	3362944595	50%	39631 11946
CGPLLU209	Lung Cancer	Preoperative, Treatment naïve	Y	Y	100	80930	8365680630	4182616104	50%	50851 7569
CGPLLU244	Lung Cancer	Pre-treatment, Day -7	N	Y	100	80930	7739561100	3768487116	49%	45925 8552
CGPLLU244	Lung Cancer	Pre-treatment, Day -1	N	Y	100	80930	8081928000	422532272	52%	51279 8646
CGPLLU244	Lung Cancer	Post-treatment, Day 6	N	Y	100	80930	8864936700	4371936263	50%	53852 7361
CGPLLU244	Lung Cancer	Post-treatment, Day 62	N	Y	100	80930	767935200	3935822054	51%	47768 7266
CGPLLU245	Lung Cancer	Post-treatment, Day 62	N	Y	100	80930	885252630	4824268339	54%	58338 16394
CGPLLU245	Lung Cancer	Post-treatment, Day 7	N	Y	100	80930	8518229300	4460236927	53%	54083 10125
CGPLLU245	Lung Cancer	Post-treatment, Day 21	N	Y	100	80930	9031131050	4824728475	53%	58313 16398
CGPLLU246	Lung Cancer	Post-treatment, Day -21	N	Y	100	80930	8520586000	35095690305	41%	42349 8086
CGPLLU246	Lung Cancer	Post-treatment, Day -32	N	Y	100	80930	5461467860	2826351657	52%	34243 82556
CGPLLU246	Lung Cancer	Post-treatment, Day 0	N	Y	100	80930	8137616600	4135036174	51%	50121 4273
CGPLLU246	Lung Cancer	Post-treatment, Day 42	N	Y	100	80930	6091276860	441332333	53%	53495 7303
CGPLLU246	Lung Cancer	Post-treatment, Day 21	N	Y	100	80930	6254777700	3016326208	48%	36164 12138
CGPLLU246	Lung Cancer	Post-treatment, Day -21	N	Y	100	80930	6185531030	3087883231	50%	33023 8388
CGPLLU246	Lung Cancer	Post-treatment, Day 0	N	Y	100	80930	627454630	28614566	46%	34508 6817
CGPLLU246	Lung Cancer	Post-treatment, Day -1	N	Y	100	80930	5701274030	124127038	22%	30877 9803
CGPLLU246	Lung Cancer	Post-treatment, Day 9	N	Y	100	80930	6091276860	4242585558	48%	28486 5793
CGPLLU246	Lung Cancer	Post-treatment, Day 42	N	Y	100	80930	6430197900	2945953499	46%	35004 7742
CGPLLU246	Lung Cancer	Post-treatment, Day 0	N	Y	100	80930	588951030	279220895	48%	33423 8423
CGPLLU246	Lung Cancer	Post-treatment, Day -1	N	Y	100	80930	585386030	286386038	44%	30977 4273
CGPLLU246	Lung Cancer	Post-treatment, Day 0	N	Y	100	80930	5867524930	2347651479	40%	28146 5793
CGPLLU246	Lung Cancer	Post-treatment, Day 42	N	Y	100	80930	6054263630	2066938782	34%	24594 6221
CGPLLU246	Lung Cancer	Post-treatment, Day 0	N	Y	100	80930	6178591380	349585805	51%	41432 7765
CGPLLU246	Lung Cancer	Post-treatment, Day 0	N	Y	100	80930	651302050	2096370387	32%	25142 6598

Patient	Patient Type	Timepoint	Fragment Profile		Mutation Analysis	Read Length	Bases in Target Region	Bases Mapped to Target Regions	Percent Mapped to Target Regions	Total Coverage	Distinct Coverage
			Preoperative	Treatment naïve							
CGPL050	Ovarian Cancer	Preoperative	Preoperative, Treatment naïve	N	Y	100	80930	823920450	4472380276	54.15%	3836
CGPLPA118	Bile Duct Cancer	Preoperative	Preoperative, Treatment naïve	N	Y	100	80930	90348327650	57021	4802	
CGPLPA122	Bile Duct Cancer	Preoperative	Treatment naïve	N	Y	100	80930	7363332130	39916160379	53%	7375
CGPLPA124	Bile Duct Cancer	Preoperative	Treatment naïve	N	Y	100	80930	7573428260	3865807442	52%	8658
CGPLPA126	Bile Duct Cancer	Preoperative	Treatment naïve	N	Y	100	80930	7946493560	4661463168	51%	10498
CGPLPA128	Bile Duct Cancer	Preoperative	Treatment naïve	N	Y	100	80930	724923830	2244188735	31%	26436
CGPLPA129	Bile Duct Cancer	Preoperative	Treatment naïve	N	Y	100	80930	7558938950	400332804	53%	3413
CGPLPA130	Bile Duct Cancer	Preoperative	Treatment naïve	N	Y	100	80930	697486650	1247144906	18%	47192
CGPLPA131	Bile Duct Cancer	Preoperative	Treatment naïve	N	Y	100	80930	7225237950	3370866342	47%	14653
CGPLPA134	Bile Duct Cancer	Preoperative	Treatment naïve	N	Y	100	80930	7268868160	3154945644	52%	30661
CGPLPA136	Bile Duct Cancer	Preoperative	Treatment naïve	N	Y	100	80930	747650750	4073978408	54%	44306
CGPLPA140	Bile Duct Cancer	Preoperative	Treatment naïve	N	Y	100	80930	7361768340	33771768342	51%	7023
CGST012	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	57115563460	2944020854	46%	5244
CGST110	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	9179291560	4298262266	47%	4839
CGST114	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	7151517230	3254567293	46%	5731
CGST113	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	644971560	3198549984	50%	40752
CGST141	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	6781001380	3440527391	46%	5404
CGST116	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	5336041060	2351386289	46%	35354
CGST118	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	6647324030	3138657777	47%	4992
CGST226	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	6288468160	2884979993	46%	2586
CGST30	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	614121310	3039994564	51%	37194
CGST32	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	6958139350	3999120469	44%	36726
CGST33	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	5563039400	3168371917	48%	3112
CGST36	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	7043791490	2992801875	42%	6737
CGST41	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	6975053190	3240466562	48%	4016
CGST45	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	6130812290	2944524276	48%	36264
CGST47	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	5961490090	3083523351	52%	37006
CGST48	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	641797230327	1497230570	23%	2410
CGST58	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	5818345630	1274708429	22%	15261
CGST80	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	638604650	3298497188	52%	5280
CGST81	Gastric Cancer	Preoperative	Treatment naïve	N	Y	100	80930	865561450	151912452	18%	6413

APPENDIX C: Table 3. Targeted cfDNA fragment analyses in cancer patients

Patient	Patient Type	Stage at Diagnosis	Alteration Type	Gene	Amino Acid (Protein)	Nucleotide	Mutation Type	Hotspot Alteration	Detected in Tissue	Mutant Allele Fraction	Distinct Coverage	Wild-type Fragments		
												Median cfDNA Fragment Size (bp)	Mode cfDNA Fragment Size (bp)	Fragment Size (bp)
OGCR291	Cancer	IV	Tumor-derived	STK11	399>G	chr19_1297027_1207627_C_T	Substitution	No	No	0.14%	11688	105	167	169
OGCR291	Cancer	IV	Tumor-derived	TP53	212>M	chr17_7571724_7571724_C_T	Substitution	Yes	Yes	0.10%	11779	105	155	171
OGCR291	Cancer	IV	Tumor-derived	KRAS	167Q>X	chr17_7578431_7578431_G_A	Substitution	Yes	Yes	22.85%	11026	193	156	169
OGCR291	Cancer	IV	Tumor-derived	AFC	126G>A	chr12_25386284_25386284_C_G	Substitution	Yes	Yes	14.63%	7632	97	152	167
OGCR291	Cancer	IV	Tumor-derived	APC	145R>X	chr5_1121756389_1121756389_C_T	Substitution	No	Yes	11.23%	7216	181	155	167
OGCR291	Cancer	IV	Tumor-derived	PIK3CA	542E>K	chr3_17853622_17853622_G_A	Substitution	Yes	Yes	11.05%	10757	86	154	166
OGCR291	Cancer	IV	Tumor-derived	KRAS	164A>Y	chr12_25378561_25378561_G_A	Substitution	Yes	Yes	18.11%	5429	180	151	171
OGCR292	Cancer	IV	Tumor-derived	CTNNB1	41T>A	chr3_41266124_41266124_G_A	Substitution	Yes	Yes	1.44%	6120	101	157	167
OGCR292	Cancer	IV	Germinal	EGFR	22844C>G	chr17_55248982_55248982_C_G	Substitution	NA	Yes	0.13%	10993	180	155	169
OGCR292	Cancer	IV	Tumor-derived	TP53	176Z>S	chr17_7578404_7578404_G_A	Substitution	No	Yes	31.93%	7587	97	158	171
OGCR293	Cancer	II	Tumor-derived	APC	213R>S	chr5_1121756389_1121756389_C_T	Substitution	Yes	Yes	0.35%	7872	95	159	168
OGCR294	Cancer	II	Tumor-derived	APC	1367Q>X	chr5_1121753946_1121753946_C_T	Substitution	Yes	Yes	0.14%	7339	84	155	167
OGCR295	Cancer	IV	Tumor-derived	PDGFRA	48>C>T	chr4_551249882_551249882_C_T	Substitution	No	Yes	0.13%	12054	89	159	167
OGCR295	Cancer	IV	Hematopoietic	IDH1	104G>Y	chr20_15136113131361131361_C_A	Substitution	No	Yes	0.45%	5802	160	157	168
OGCR296	Cancer	II	Hematopoietic	EGFR	932E>K	chr7_55266472_55266472_G_A	Substitution	NA	Yes	0.34%	6330	169	157	169
OGCR297	Cancer	III	Germinal	KIT	18L>F	chr4_556524235_556524235_C_T	Substitution	NA	Yes	30.48%	6375	89	161	166
OGCR298	Cancer	II	Hematopoietic	DNM73A	882R>H	chr2_254572457245724_G_C	Substitution	Yes	Yes	41.39%	3580	162	155	164
OGCR298	Cancer	II	Hematopoietic	DNM73A	714R>C	chr2_25465351_254653514_G_C	Substitution	No	Yes	0.08%	13032	160	155	168
OGCR298	Cancer	II	Hematopoietic	PDGFRA	414G>Y	chr3_178824787_178824787_G_T	Substitution	No	Yes	0.11%	13475	93	158	169
OGCR298	Cancer	II	Hematopoietic	DNM73A	735Y>C	chr2_25656289_25656289_C_T	Substitution	No	Yes	0.55%	5815	160	158	168
OGCR299	Cancer	III	Hematopoietic	DNM73A	710Z>S	chr2_25463229_25463229_G_C	Substitution	No	Yes	0.30%	11985	160	154	165
OGCR299	Cancer	II	Hematopoietic	DNM73A	720R>G	chr2_25463235_25463235_G_C	Substitution	No	Yes	0.12%	1563	96	162	172
OGCR301	Cancer	II	Tumor-derived	ATM	2387D>X	chr11_168198487_168198487_C_T	Substitution	No	Yes	0.15%	7487	160	182	173
OGCR302	Cancer	II	Tumor-derived	TP53	141C>Y	chr7_7578527_7578527_C_T	Substitution	Yes	Yes	0.05%	13475	93	158	169
OGCR302	Cancer	II	Tumor-derived	BRAF	163V>E	chr7_1404536_1404536_A_T	Substitution	No	Yes	0.12%	11763	95	154	165
OGCR303	Cancer	II	Hematopoietic	TP53	173V>L	chr17_7578413_7578413_C_A	Substitution	Yes	Yes	0.08%	13967	95	159	171
OGCR303	Cancer	II	Hematopoietic	DNM73A	755F>S	chr2_25463229_25463229_A_G	Substitution	No	Yes	0.21%	10467	81	160	169
OGCR303	Cancer	II	Hematopoietic	DNM73A	237R>H	chr2_25463235_25463235_G_T	Substitution	No	Yes	0.17%	10945	103	156	169
OGCR304	Cancer	II	Tumor-derived	EGFR	113R>P	chr7_5527308_5527308_A_T	Substitution	No	Yes	0.21%	5881	160	155	169
OGCR304	Cancer	II	Tumor-derived	ATM	30774>G>A	chr11_168142134_168142134_G_A	Substitution	No	Yes	0.055%	24784	84	153	164
OGCR304	Cancer	II	Hematopoietic	PDGFRA	163V>E	chr7_1404536_1404536_A_T	Substitution	Yes	Yes	0.12%	11763	95	154	165
OGCR305	Cancer	II	Tumor-derived	TP53	2017V>L	chr17_7578413_7578413_C_A	Substitution	No	Yes	0.08%	12987	95	159	171
OGCR305	Cancer	II	Tumor-derived	DNM73A	755F>S	chr2_25463229_25463229_A_G	Substitution	No	Yes	0.11%	12507	103	155	169
OGCR306	Cancer	II	Tumor-derived	EGFR	196R>X	chr7_7578635_7578635_G_A	Substitution	Yes	Yes	0.19%	10301	93	160	168
OGCR306	Cancer	II	Tumor-derived	CDKN2A	107R>C	chr9_21971038_21971038_G_A	Substitution	No	Yes	0.22%	16168	90	153	167
OGCR306	Cancer	II	Tumor-derived	DNMT3A	61Q>K	chr12_2583263_2583263_G_T	Substitution	Yes	Yes	0.27%	10502	103	152	165
OGCR306	Cancer	II	Tumor-derived	TP53	2007V>C	chr4_55130165_55130165_G_C	Substitution	No	Yes	0.45%	12987	101	154	165
OGCR307	Cancer	II	Tumor-derived	EGFR	618H>R	chr7_75787120_75787120_C_T	Substitution	No	Yes	0.08%	13789	95	159	171
OGCR307	Cancer	II	Tumor-derived	FBN1	178S>P	chr3_17883685_17883685_A_G	Substitution	No	Yes	0.19%	10845	103	157	165
OGCR308	Cancer	II	Tumor-derived	JAK2	805>V	chr9_21971038_21971038_G_A	Substitution	No	Yes	0.02%	9437	90	153	167
OGCR308	Cancer	II	Tumor-derived	SMN1	5012A>G	chr12_2583263_2583263_G_T	Substitution	Yes	Yes	0.27%	6390	100	152	166
OGCR308	Cancer	II	Tumor-derived	TP53	2007V>C	chr4_55130165_55130165_G_C	Substitution	No	Yes	0.43%	12987	101	154	165
OGCR308	Cancer	II	Tumor-derived	DNMT3A	755F>S	chr17_5523303_5523303_A_G	Substitution	No	Yes	0.11%	12507	93	155	169
OGCR308	Cancer	II	Tumor-derived	TP53	178S>P	chr7_7578729_7578729_G_T	Substitution	No	Yes	0.19%	10301	93	156	168
OGCR309	Cancer	II	Tumor-derived	ERBB4	115R>A	chr4_153248385_153248385_A_G	Substitution	Yes	Yes	0.98%	68594	101	157	165
OGCR309	Cancer	II	Tumor-derived	DNMT3A	895>L	chr9_5086852_5086852_G_G	Substitution	No	Yes	0.02%	3756	100	153	166
OGCR309	Cancer	II	Tumor-derived	PDGFRA	2414R>G	chr2_25457242_25457242_G_G	Substitution	No	Yes	7.30%	6860	100	158	168
OGCR309	Cancer	II	Tumor-derived	DNMT3A	2017V>C	chr2_5525385_5525385_G_C	Substitution	Yes	Yes	0.24%	12987	101	154	165
OGCR309	Cancer	II	Tumor-derived	TP53	178S>P	chr7_5523303_5523303_A_G	Substitution	No	Yes	6.32%	7395	81	160	166
OGCR309	Cancer	II	Tumor-derived	DNMT3A	178S>P	chr3_17883685_17883685_A_G	Substitution	Yes	Yes	0.98%	4885	103	155	168
OGCR309	Cancer	II	Tumor-derived	TP53	178S>P	chr7_140453136_140453136_A_G	Substitution	No	Yes	38.75%	3756	100	153	166
OGCR309	Cancer	II	Tumor-derived	DNMT3A	178S>P	chr2_25457242_25457242_G_G	Substitution	Yes	Yes	0.56%	6860	100	158	168
OGCR309	Cancer	II	Tumor-derived	TP53	178S>P	chr2_5525385_5525385_G_C	Substitution	No	Yes	0.34%	12987	101	154	165
OGCR309	Cancer	II	Tumor-derived	DNMT3A	178S>P	chr7_5523303_5523303_A_G	Substitution	Yes	Yes	27.69%	7729	103	156	166
OGCR309	Cancer	II	Tumor-derived	TP53	178S>P	chr5_112175729_112175729_G_T	Substitution	No	Yes	0.11%	14067	92	157	170
OGCR309	Cancer	II	Tumor-derived	DNMT3A	178S>P	chr2_25457242_25457242_G_T	Substitution	Yes	Yes	0.29%	8623	76	157	169
OGCR309	Cancer	II	Tumor-derived	TP53	178S>P	chr7_140453136_140453136_A_G	Substitution	No	Yes	0.70%	10606	103	155	167
OGCR309	Cancer	II	Tumor-derived	DNMT3A	178S>P	chr2_25457242_25457242_G_G	Substitution	Yes	Yes	3.00%	9084	101	157	168
OGCR309	Cancer	II	Tumor-derived	TP53	178S>P	chr1_1525385_1525385_G_G	Substitution	Yes	Yes	0.47%	3391	90	153	166
OGCR309	Cancer	II	Tumor-derived	DNMT3A	178S>P	chr2_25398295_25398295_G_G	Substitution	Yes	Yes	0.17%	5913	103	153	166
OGCR309	Cancer	II	Tumor-derived	TP53	178S>P	chr5_112175729_112175729_G_T	Substitution	Yes	Yes	0.07%	6150	72	161	174
OGCR309	Cancer	II	Tumor-derived	DNMT3A	178S>P	chr2_25398294_25398294_G_G	Substitution	Yes	Yes	0.30%	4684	103	156	168
OGCR309	Cancer	II	Tumor-derived	TP53	178S>P	chr2_25457242_25457242_G_G	Substitution	No	Yes	2.50%	6902	85	158	170
OGCR309	Cancer	II	Tumor-derived	DNMT3A	178S>P	chr5_112175729_112175729_G_G	Substitution	Yes	Yes	0.36%	7229	102	155	167
OGCR309	Cancer	II	Tumor-derived	TP53	178S>P	chr1_1525385_1525385_G_G	Substitution	Yes	Yes	0.27%	6739	94	155	169
OGCR309	Cancer	II	Tumor-derived	DNMT3A	178S>P	chr4_153247289_153247289_G_A	Substitution	Yes	Yes	0.25%	5623	101	155	166

Patient	Patient Type	Diagnosis	Stage at Diagnosis	Alteration Type	Gene	Amino Acid (Protein)	Nucleotide	Type	Alteration	Fraction	Distinct Coverage	Tissue	Hotspot	Detected in Tissue	Alteration	Hotspot	Median cfDNA Fragment Size (bp)	Minimum cfDNA Fragment Size (bp)	Median cfDNA Fragment Size (bp)	Wild-type Fragments
OGCR0316	Cerebral Cancer	III	Tumor-derived	TFS3	CDKN2A	245G>S	chr17_7577548_7577546_C_T	Substitution	Yes	Yes	6.52%	12860	103	150	157	7479	93	166	163	168
OGCR0316	Cerebral Cancer	III	Tumor-derived	CDTNB1	1M>R	37S>C	chr9_21974825_21974825 A,C	Substitution	No	Yes	5.74%	13682	103	148	165	165	165	165	162	162
OGCR0316	Cerebral Cancer	III	Tumor-derived	EGFR	2702S>T	35S>C	chr3_41265113_41265113 C,G	Substitution	No	Yes	5.47%	13682	103	148	153	16116	85	166	166	166
OGCR0316	Cerebral Cancer	III	Tumor-derived	ATM	3508R>P	220T>C	chr1_18923087_18923087 G,C	Substitution	No	Yes	0.11%	17060	103	150	153	166	85	166	166	163
OGCR0316	Cerebral Cancer	III	Tumor-derived	TFS3	ATM	1028W>R	chr11_183142132_188142132 T,C	Substitution	Yes	Yes	0.13%	14587	103	150	152	166	84	166	166	163
OGCR0317	Cerebral Cancer	III	Tumor-derived	ATM	1216R>X	115>Y	chr5_112142132_112142132 C,T	Substitution	No	Yes	0.23%	10483	103	150	152	164	152	164	165	165
OGCR0317	Cerebral Cancer	III	Tumor-derived	DNM73A	6589N>S	5522S>T	chr2_25463389_25463389 C,T	Substitution	No	Yes	0.28%	3497	103	150	149	166	98	158	158	163
OGCR0318	Cerebral Cancer	I	Hematopoietic	KIT	18L>F	78R>W	chr4_5562253_5562253 C,T	Substitution	No	Yes	34.76%	6521	103	163	170	175	163	170	175	163
OGCR0322	Cerebral Cancer	I	Tumor-derived	ERBB4	78R>W	78R>P	chr2_212889479_212889479 G,A	Substitution	No	Yes	0.12%	11633	103	180	182	174	167	167	174	174
OGCR0322	Cerebral Cancer	I	Tumor-derived	CDKN2A	12S>I	882R>H	chr2_25457242_25457242 C,T	Substitution	No	Yes	0.20%	6318	88	161	159	174	167	174	170	170
OGCR0321	Cerebral Cancer	I	Hematopoietic	EGFR	51S>Y	5522S>T	chr7_5522S25_5522S25 C,A	Substitution	No	Yes	0.08%	9559	94	159	171	158	159	171	172	172
OGCR0321	Cerebral Cancer	I	Germinal	TFS3	125T>R	125T>G	chr17_75775313_75775313 G,C	Substitution	No	Yes	41.86%	5545	103	150	158	170	98	158	158	170
OGCR0322	Cerebral Cancer	IV	Tumor-derived	TFS3	125T>G	125T>G	chr17_75775313_75775313 G,C	Substitution	No	Yes	19.98%	605	104	164	165	170	163	170	175	175
OGCR0322	Cerebral Cancer	IV	Tumor-derived	ERBB4	8R>F	600V>E	chr1_140453136_140453136 A,T	Substitution	No	Yes	43.03%	1265	89	158	159	171	165	171	171	171
OGCR0333	Cerebral Cancer	IV	Tumor-derived	TFS3	8R>F	8R>P	chr2_212498194_212498194 T,G	Substitution	No	Yes	22.26%	3338	102	153	153	168	165	165	168	168
OGCR0333	Cerebral Cancer	IV	Tumor-derived	ERBB4	681E>A	12G>V	chr2_25457242_25457242 C,A	Substitution	No	Yes	1.00%	3038	102	172	175	170	170	174	174	174
OGCR0334	Cerebral Cancer	IV	Tumor-derived	TFS3	125T>S	125T>S	chr7_5522S25_5522S25 C,A	Substitution	No	Yes	13.44%	1725	105	160	160	170	175	175	175	175
OGCR0334	Cerebral Cancer	IV	Tumor-derived	EGFR	682R>Y	682R>Y	chr3_178816890_178816890 C,T	Substitution	No	Yes	35.28%	1188	103	159	159	164	164	174	174	174
OGCR0334	Cerebral Cancer	IV	Tumor-derived	PK3CA	104P>R	104P>R	chr3_178816894_178816894 C,T	Substitution	No	Yes	3.85%	1798	103	159	159	173	166	166	166	173
OGCR0334	Cerebral Cancer	IV	Tumor-derived	ERBB4	600V>E	600V>E	chr7_140453136_140453136 A,T	Substitution	No	Yes	0.32%	2411	99	155	155	167	167	171	171	171
OGCR0335	Cerebral Cancer	IV	Tumor-derived	TFS3	125T>G	125T>H	chr17_75775313_75775313 G,C	Substitution	No	Yes	75.26%	737	104	156	156	170	170	170	170	170
OGCR0335	Cerebral Cancer	IV	Tumor-derived	KRAS	12G>D	12G>V	chr12_25398281_25398281 C,A	Substitution	No	Yes	42.87%	1080	102	153	153	166	166	168	168	168
OGCR0336	Cerebral Cancer	IV	Tumor-derived	KRAS	12G>V	12G>V	chr5_1121751347_1121751347 G,C	Substitution	No	Yes	81.61%	381	102	161	161	171	171	171	171	
OGCR0336	Cerebral Cancer	IV	Tumor-derived	EGFR	125T>S	125T>S	chr13_178816894_178816894 C,T	Substitution	No	Yes	3.12%	6497	72	153	153	177	177	177	177	177
OGCR0337	Cerebral Cancer	IV	Tumor-derived	EGFR	485M>I	485M>I	chr1_112168261_112168261 G,A	Substitution	No	Yes	46.26%	1686	103	159	159	173	163	173	173	173
OGCR0337	Cerebral Cancer	IV	Tumor-derived	KRAS	12G>D	12G>D	chr12_25398281_25398281 C,A	Substitution	No	Yes	27.03%	1498	105	153	153	164	164	166	166	166
OGCR0338	Cerebral Cancer	IV	Tumor-derived	EGFR	876R>P	876R>P	chr17_75775313_75775313 G,C	Substitution	No	Yes	1.94%	1255	105	158	158	171	171	171	171	171
OGCR0338	Cerebral Cancer	IV	Tumor-derived	APC	125T>S	125T>S	chr5_1121751347_1121751347 G,C	Substitution	No	Yes	2.36%	1639	101	158	158	172	172	172	172	172
OGCR0338	Cerebral Cancer	IV	Tumor-derived	PIK3CA	407C>F	407C>F	chr3_178852085_178852085 A,T	Substitution	No	Yes	3.14%	1143	102	154	154	170	170	170	170	170
OGCR0338	Cerebral Cancer	IV	Tumor-derived	PIK3CA	104T>L	104T>L	chr3_178852085_178852085 A,T	Substitution	No	Yes	1.71%	1584	103	159	159	171	171	171	171	171
OGCR0338	Cerebral Cancer	IV	Tumor-derived	ERBB4	78R>P	78R>P	chr1_112168261_112168261 G,A	Substitution	No	Yes	18.26%	876	101	162	162	175	175	175	175	175
OGCR0340	Cerebral Cancer	IV	Tumor-derived	KRAS	12G>D	12G>D	chr5_112168261_112168261 G,A	Substitution	No	Yes	22.57%	796	105	158	158	164	164	168	168	168
OGCR0340	Cerebral Cancer	IV	Tumor-derived	APC	125T>S	125T>S	chr17_75775313_75775313 G,C	Substitution	No	Yes	0.53%	5631	105	158	158	166	166	168	168	168
OGPLB40	Breast Cancer	II	Germinal	AR	362P>R	362P>R	chr8_66766163_66766163 C,G	Substitution	No	Yes	28.93%	10277	78	162	162	168	168	173	173	173
OGPLB44	Breast Cancer	II	Hematopoietic	DNM73A	882R>H	882R>H	chr2_25457242_25457242 C,T	Substitution	Yes	Yes	0.82%	1715	99	161	161	171	171	171	171	171
OGPLB44	Breast Cancer	II	Tumor-derived	CDTNB1	705R>P	705R>P	chr2_25457242_25457242 G,A	Substitution	No	Yes	0.41%	10337	103	159	159	171	171	171	171	171
OGPLB44	Breast Cancer	II	Tumor-derived	PDGFRA	869Y>M	869Y>M	chr4_55155369_55155369 G,A	Substitution	No	Yes	0.13%	12640	103	159	159	164	164	168	168	168
OGPLB44	Breast Cancer	II	Germinal	ALK	1231R>Q	1231R>Q	chr2_25457242_25457242 C,T	Substitution	No	Yes	34.61%	5631	105	158	158	170	170	179	179	179
OGPLB44	Breast Cancer	II	Tumor-derived	EGFR	689R>Q	689R>Q	chr7_55246740_55246740 G,A	Substitution	No	Yes	0.19%	12467	101	167	167	174	174	174	174	174
OGPLB45	Breast Cancer	II	Hematopoietic	DNM73A	743P>S	743P>S	chr2_25457242_25457242 G,A	Substitution	No	Yes	0.18%	10277	78	162	162	171	171	171	171	171
OGPLB45	Breast Cancer	II	Tumor-derived	GIMAS	201R>H	201R>H	chr2_25457242_25457242 G,A	Substitution	Yes	Yes	0.68%	6011	101	153	153	166	166	167	167	167
OGPLB45	Breast Cancer	II	Tumor-derived	CDTNB1	301>S	301>S	chr3_41266032_41266032 A,C	Substitution	No	Yes	0.42%	3873	101	161	161	173	173	173	173	173
OGPLB45	Breast Cancer	II	Germinal	FGFR3	403R>E	403R>E	chr2_209103186_209103186 A,G	Substitution	No	Yes	34.82%	3405	97	165	165	170	170	170	170	170
OGPLB45	Breast Cancer	II	Tumor-derived	EGFR	283R>H	283R>H	chr2_25457242_25457242 C,T	Substitution	No	Yes	0.11%	10259	87	157	157	168	168	168	168	168
OGPLB45	Breast Cancer	II	Tumor-derived	PIK3CA	545E>K	157T>H	chr3_178852085_178852085 G,A	Substitution	No	Yes	0.18%	5163	101	158	158	167	167	167	167	167
OGPLB45	Breast Cancer	II	Tumor-derived	TFS3	103D>V	103D>V	chr2_25457242_25457242 G,A	Substitution	No	Yes	0.28%	62550	103	159	159	168	168	168	168	168
OGPLB45	Breast Cancer	II	Germinal	CDTNB1	774E>V	774E>V	chr2_25457242_25457242 G,A	Substitution	No	Yes	0.28%	7558	103	159	159	170	170	170	170	170
OGPLB45	Breast Cancer	II	Tumor-derived	ALK	708S>P	708S>P	chr2_25457242_25457242 G,A	Substitution	No	Yes	44.03%	2389	103	159	159	174	174	174	174	174
OGPLB45	Breast Cancer	II	Germinal	ERBB4	158A>E	158A>E	chr2_25457242_25457242 G,A	Substitution	No	Yes	0.27%	11348	103	159	159	173	173	173	173	173
OGPLB45	Breast Cancer	II	Tumor-derived	PIK3CA	20+G>T	20+G>T	chr2_25457242_25457242 G,A	Substitution	No	Yes	35.58%	3422	102	158	158	174	174	174	174	174
OGPLB45	Breast Cancer	II	Germinal	PIK3CA	104T>S	104T>S	chr3_41266032_41266032 A,C	Substitution	No	Yes	0.14%	7297	103	159	159	173	173	173	173	173
OGPLB45	Breast Cancer	II	Tumor-derived	CDTNB1	1121753122_1121753122 G,A	Substitution	NA	NA	NA	Yes	35.57%	4342	103	159	159	173	173	173	173	173
OGPLB45	Breast Cancer	II	Germinal	PTEN	170S>I	170S>I	chr3_178852085_178852085 A,G	Substitution	Yes	Yes	0.12%	11785	103	159	159	173	173	173	173	173
OGPLB45	Breast Cancer	II	Tumor-derived	CDTNB1	171275322_171275322 A,C	Substitution	NA	NA	NA	Yes	35.58%	6161	103	159	159	173	173	173	173	1

Patient	Patient Type	Stage at Diagnosis	Alteration Type	Gene	Amino Acid (Protein)	Nucleotide	Type	Alteration	Hotspot	Detected in Tissue	Mutant Allele Fraction	Distinct Coverage	Fragment Size (bp)	Minimum cfDNA Fragment Size (bp)	Median cfDNA Fragment Size (bp)	Wild-type Fragments
CGPLB86	Breast Cancer	I	Germinal	SMARCB1	T96>G	chr22_241593728-241591265_A_G	Substitution	NA	Yes	43.38%	3098	68	160	162	168	174
CGPLB87	Breast Cancer	I	Tumor-derived	JAK2	215R>X	chr9_50565891_55354591_C_T	Substitution	No	No	0.35%	3080	101	163	162	168	175
CGPLB87	Breast Cancer	I	Hematopoietic	DNM173A	862R>H	chr2_25457232_25457242_C_T	Substitution	No	No	0.31%	6180	101	163	160	167	175
CGPLB87	Breast Cancer	I	Tumor-derived	SMAD4	486R>C	chr18_48864684_48604664_C_T	Substitution	NA	Yes	0.40%	7746	86	160	166	166	172
CGPLB87	Breast Cancer	I	Germinal	AR	651S>N	chrX_68933310_68931310_G_A	Substitution	NA	Yes	42.94%	2266	106	160	166	166	172
CGPLB88	Breast Cancer	I	Tumor-derived	CDK6	51E>K	chr7_92463487_92462487_C_T	Substitution	No	No	0.15%	17537	89	185	200	223	223
CGPLB88	Breast Cancer	I	Germinal	APC	1128V>A	chr5_112174695_112174685_C_T	Substitution	NA	Yes	34.15%	5819	101	162	172	173	173
CGPLB88	Breast Cancer	I	Tumor-derived	TBP3	257>P	chr7_757757751_A_G	Substitution	No	Yes	0.29%	15530	77	150	164	162	162
CGPLB88	Breast Cancer	I	Hematopoietic	TBP3	213R>A	chr7_757757751_75775721_G_A	Substitution	Yes	No	0.10%	9893	159	159	164	171	171
CGPLB88	Breast Cancer	I	Tumor-derived	DNM173A	551D>G	chr2_25457232_25457245_C_A	Substitution	No	Yes	5.81%	8620	95	162	167	173	173
CGPLB88	Breast Cancer	I	Hematopoietic	AR	139>Q	chr9_6876826_68768226_G_A	Substitution	No	No	0.60%	8236	85	162	169	168	170
CGPLB88	Breast Cancer	I	Tumor-derived	DNM173A	862R>H	chr2_25457232_25457242_C_T	Substitution	Yes	Yes	0.11%	14056	93	160	168	168	170
CGPLB88	Breast Cancer	I	Germinal	PDGFRα	401A>D	chr4_55136860_55136880_C_A	Substitution	NA	Yes	34.12%	5329	120	161	165	171	171
CGPLB88	Breast Cancer	I	Tumor-derived	GNA3	201R>H	chr20_57484421_57484421_G_A	Substitution	Yes	Yes	0.13%	7010	97	158	169	170	170
CGPLB88	Breast Cancer	I	Hematopoietic	TBP3	241S>F	chr17_5073770_5073770_G_A	Substitution	Yes	Yes	1.95%	11371	103	156	165	167	167
CGPLB88	Lung Cancer	I	Tumor-derived	KIF4A	122>C	chr25_53939235_53939235_C_A	Substitution	Yes	Yes	5.10%	7641	103	155	166	166	166
CGPLB88	Lung Cancer	I	Tumor-derived	EGFR	373>S	chr7_55224336_53224336_C_T	Substitution	No	Yes	0.16%	5996	103	156	168	168	168
CGPLB88	Lung Cancer	I	Tumor-derived	ATM	282P>L	chr11_1981156727_1981156727_C_T	Substitution	No	No	0.22%	4956	101	159	166	168	168
CGPLB88	Lung Cancer	I	Germinal	PK3CA	545E>K	chr3_17835691_17835691_G_A	Substitution	Yes	Yes	2.94%	6540	103	153	166	170	170
CGPLB88	Lung Cancer	I	Tumor-derived	FREB4	426R>K	chr2_21256841_21256841_C_A	Substitution	No	No	0.18%	7648	101	156	164	168	168
CGPLB88	Lung Cancer	I	Hematopoietic	JAK2	617V>F	chr9_5073770_5073770_G_A	Substitution	Yes	No	0.23%	5920	100	155	164	166	166
CGPLB88	Lung Cancer	I	Tumor-derived	TBP3	282R>P	chr7_55224336_53224336_C_G	Substitution	No	Yes	1.30%	5356	103	155	166	168	168
CGPLB88	Lung Cancer	I	Hematopoietic	DNM173A	737>H	chr25_5363283_5363283_A_C	Substitution	No	Yes	0.84%	7284	101	158	165	170	170
CGPLB88	Lung Cancer	I	Tumor-derived	RB1	861>T>C	chr13_4883305_4883305_C_T	Substitution	No	Yes	0.87%	4183	103	160	166	170	170
CGPLB88	Lung Cancer	I	Tumor-derived	ATM	581L>F	chr11_10812695_10812699_A_T	Substitution	No	No	0.20%	6776	100	157	166	168	168
CGPLB88	Lung Cancer	I	Hematopoietic	TBP3	201R>Q	chr17_5073770_5073770_G_A	Substitution	No	Yes	0.15%	4807	100	155	165	167	167
CGPLB88	Lung Cancer	I	Tumor-derived	ALK	153T>E	chr17_55224336_53224336_C_T	Substitution	No	No	0.55%	5282	103	156	167	171	171
CGPLB88	Lung Cancer	I	Germinal	PDGFRα	200T>S	chr13_55136865_55136865_C_G	Substitution	No	Yes	0.94%	7122	103	158	166	173	173
CGPLB88	Lung Cancer	I	Tumor-derived	CDKN2A	126L>P	chr9_17977521_17977521_G_A	Substitution	No	Yes	43.47%	2825	101	156	165	174	174
CGPLB88	Lung Cancer	I	Tumor-derived	EGR3	888L>R	chr7_55255515_55255915_T_G	Substitution	No	No	0.22%	5940	95	157	166	174	174
CGPLB88	Lung Cancer	I	Tumor-derived	BRAF	354R>Q	chr7_140494187_140494187_C_T	Substitution	No	No	0.14%	11251	103	153	167	167	167
CGPLB88	Lung Cancer	I	Tumor-derived	TBP3	201L>Q	chr9_21977521_21977521_G_A	Substitution	No	No	0.21%	10805	85	155	165	173	173
CGPLB88	Lung Cancer	I	Hematopoietic	DNM173A	129Y>D	chr2_25467394_25467394_A_C	Substitution	No	Yes	0.15%	6795	103	156	166	173	173
CGPLB88	Lung Cancer	I	Tumor-derived	STK11	216S>Y	chr19_12236239_12236239_C_A	Substitution	No	Yes	1.25%	6795	91	156	161	164	164
CGPLB88	Lung Cancer	I	Tumor-derived	EGFR	881R>P	chr7_55255515_55255915_T_G	Substitution	No	Yes	42.52%	4581	92	157	164	173	173
CGPLB88	Lung Cancer	I	Tumor-derived	BRCA1	606>S>T	chr19_12231525_12231525_C_G	Substitution	No	No	0.22%	13656	87	160	174	174	174
CGPLB88	Lung Cancer	I	Tumor-derived	TBP3	278P>S	chr17_757757751_757757751_G_A	Substitution	Yes	No	0.10%	5247	103	155	165	167	167
CGPLB88	Lung Cancer	I	Hematopoietic	DNM173A	161A>S	chr2_25467394_25467394_A_C	Substitution	No	Yes	0.21%	10805	85	155	166	168	168
CGPLB88	Lung Cancer	I	Tumor-derived	STK11	126W>I	chr19_12236239_12236239_C_A	Substitution	No	Yes	0.15%	6795	83	156	161	166	166
CGPLB88	Lung Cancer	I	Tumor-derived	ERBB4	234N>S	chr19_12231525_12231525_C_G	Substitution	No	Yes	0.22%	5142	91	156	161	166	166
CGPLB88	Lung Cancer	I	Tumor-derived	GNA11	606>S>T	chr19_12231525_12231525_C_G	Substitution	No	No	0.20%	8657	103	155	166	167	167
CGPLB88	Lung Cancer	I	Tumor-derived	TBP3	160D>Y	chr17_757757751_757757751_G_A	Substitution	Yes	No	0.10%	5241	103	155	165	167	167
CGPLB88	Lung Cancer	I	Tumor-derived	DNM173A	136Q>W	chr17_757757751_757757751_G_A	Substitution	Yes	Yes	1.78%	10806	103	157	168	169	169
CGPLB88	Lung Cancer	I	Tumor-derived	ERBB4	129B>P	chr2_21256371_21256371_G_A	Substitution	No	Yes	0.98%	10919	103	156	161	169	169
CGPLB88	Lung Cancer	I	Tumor-derived	STK11	888L>R	chr2_212587243_212587243_T_C	Substitution	No	No	0.22%	5151	101	160	166	170	170
CGPLB88	Lung Cancer	I	Tumor-derived	TBP3	278P>S	chr17_757757751_757757751_G_A	Substitution	No	Yes	0.20%	8657	103	155	166	167	167
CGPLB88	Lung Cancer	I	Tumor-derived	DNM173A	175J>M	chr2_25467394_25467394_G_A	Substitution	No	Yes	0.08%	11070	103	156	165	168	168
CGPLB88	Lung Cancer	I	Tumor-derived	ERBB4	719H>R	chr7_757757751_757757751_G_A	Substitution	Yes	Yes	0.07%	11063	83	157	166	168	168
CGPLB88	Lung Cancer	I	Tumor-derived	DNM173A	759F>Y	chr19_12231525_12231525_G_A	Substitution	No	No	0.35%	5861	88	162	166	166	166
CGPLB88	Lung Cancer	I	Tumor-derived	DNM173A	877P>L	chr12_25393285_25393285_A_G	Substitution	Yes	Yes	0.15%	5858	103	157	162	168	168
CGPLB88	Lung Cancer	I	Tumor-derived	DNM173A	891Q>X	chr11_1981157758_1981157758_C_T	Substitution	NA	Yes	43.84%	2740	101	157	165	167	167
CGPLB88	Lung Cancer	I	Tumor-derived	DNM173A	882R>C	chr2_25467394_25467394_G_A	Substitution	No	Yes	0.28%	6565	103	158	166	168	168
CGPLB88	Lung Cancer	I	Tumor-derived	CDH11	251T>M	chr2_25467394_25467394_G_A	Substitution	No	Yes	0.92%	6513	101	161	164	175	175
CGPLB88	Lung Cancer	I	Tumor-derived	KRAS	122>V	chr12_25393285_25393285_A_G	Substitution	Yes	Yes	0.21%	5939	103	157	162	168	168
CGPLB88	Lung Cancer	I	Tumor-derived	DNM173A	897P>G	chr2_25457197_25457197_A_C	Substitution	No	Yes	1.53%	6944	102	157	165	171	171
CGPLB88	Lung Cancer	I	Tumor-derived	DNM173A	882R>C	chr2_25467394_25467394_G_A	Substitution	No	Yes	0.29%	11233	103	160	166	171	171
CGPLB88	Lung Cancer	I	Tumor-derived	CDH11	251T>M	chr2_25467394_25467394_G_A	Substitution	No	No	0.13%	10966	103	157	162	172	172
CGPLB88	Lung Cancer	I	Tumor-derived	PIK3CA	861Q>X	chr3_1788441745_1788441745_G_A	Substitution	No	No	0.11%	7235	101	155	167	170	170
CGPLB88	Lung Cancer	I	Hematopoietic	DNM173A	879R>D	chr2_25457252_25457252_T_C	Substitution	No	Yes	0.36%	8369	103	153	161	169	169
CGPLB88	Lung Cancer	I	Germinal	APC	261T>I	chr5_112174123_112174123_G_T	Substitution	NA	Yes	39.91%	2639	103	157	162	171	171

Patient	Patient Type	Stage at Diagnosis	Alteration Type	Gene	Amino Acid (Protein)	Nucleotide	Mutation Type	Hotspot	Detected in Tissue	Alteration Fraction	Distinct Coverage	Mutant Allele %	Minimum cDNA	25th Percentile cDNA Fragment Size (bp)	Median cDNA Fragment Size (bp)	Wild-type Fragments Median cDNA Fragment Size (bp)
CGPLU182	Lung Cancer	I	Tumor derived	STK11	267D>Y	chr19_1220691_1220691_G_C	Substitution	No	Yes	2.45%	6085	91	156	165	168	176
CGPLU183	Lung Cancer	I	Tumor derived	TP53	293G>A	chr17_7577058_7577058_C_A	Substitution	No	Yes	2.07%	6880	92	156	164	168	168
CGPLU183	Lung Cancer	I	Tumor derived	TP53	262R>P	chr17_7577058_7577058_C_A	Substitution	No	Yes	1.94%	7780	92	156	167	168	168
CGPLU183	Lung Cancer	I	Tumor derived	TP53	177P>L	chr17_7577058_7577058_C_A	Substitution	Yes	No	0.08%	5036	101	160	160	169	171
CGPLU183	Lung Cancer	I	Tumor derived	TP53	565S>Y	chr13_48955578_48955578_C_G	Substitution	No	Yes	1.01%	4678	109	157	157	168	168
CGPLU183	Lung Cancer	I	Hemizygous	DNM73A	682R>C	chr2_25457243_25457243_G_A	Substitution	Yes	No	0.16%	7196	102	162	162	166	172
CGPLU197	Lung Cancer	I	Hematozoic	DNM73A	879N>D	chr2_25457252_25457252_T_C	Substitution	No	No	0.38%	7147	103	161	161	166	166
CGPLU198	Lung Cancer	I	Tumor derived	TP53	162D>N	chr17_7578445_7578445_A_G	Substitution	No	Yes	0.87%	5322	97	157	157	168	168
CGPLU198	Lung Cancer	I	Tumor derived	EGFR	258I>R	chr7_5525515_5525515_T_G	Substitution	Yes	Yes	0.52%	8303	100	160	160	173	172
CGPLU202	Lung Cancer	II	Tumor derived	TP53	790T>M	chr7_745249077_75549371_C_T	Substitution	Yes	Yes	0.05%	14197	90	151	165	166	166
CGPLU202	Lung Cancer	II	Tumor derived	EGFR	688E>X	chr7_5525514_5525514_G_T	Substitution	No	No	0.13%	9279	51	150	150	167	167
CGPLU204	Lung Cancer	II	Tumor derived	KIT	956R>Q	chr4_55604656_55604656_G_A	Substitution	No	No	0.28%	7185	103	157	165	168	168
CGPLU205	Lung Cancer	II	Hematozoic	DNM73A	736R>C	chr2_25463287_25463287_G_A	Substitution	No	Yes	0.70%	10739	96	156	165	166	166
CGPLU205	Lung Cancer	II	Hematozoic	DNM73A	696Q>X	chr2_2546356_2546356_G_A	Substitution	No	Yes	0.47%	12065	100	154	165	165	165
CGPLU206	Lung Cancer	II	Tumor derived	TP53	672>IG>A	chr17_7578176_C_T	Substitution	Yes	Yes	26.13%	6746	94	145	165	164	164
CGPLU206	Lung Cancer	II	Tumor derived	TP53	131N>S	chr17_7578538_7578538_L_C	Substitution	No	No	0.21%	11225	103	147	164	164	164
CGPLU207	Lung Cancer	II	Tumor derived	TP53	376>G>A	chr17_7578535_7578535_C_T	Substitution	Yes	Yes	0.32%	11224	103	159	165	170	170
CGPLU207	Lung Cancer	II	Germine	ALK	419F>L	chr2_29806265_29806265_A_G	Substitution	NA	Yes	34.58%	4980	101	160	166	170	170
CGPLU207	Lung Cancer	II	Tumor derived	EGFR	790T>M	chr7_745249077_75549371_C_T	Substitution	Yes	No	0.05%	13216	85	161	165	172	172
CGPLU208	Lung Cancer	II	Tumor derived	TP53	260P>L	chr17_7577052_7577052_G_A	Substitution	Yes	Yes	1.35%	9211	101	156	168	168	168
CGPLU208	Lung Cancer	II	Germine	EGFR	224F>H	chr7_552520261_552520261_G_A	Substitution	NA	Yes	39.34%	5253	100	159	164	170	170
CGPLU208	Lung Cancer	II	Tumor derived	EGFR	858I>R	chr7_5525515_5525515_T_G	Substitution	Yes	Yes	0.86%	10233	103	160	170	171	171
CGPLU208	Lung Cancer	II	Tumor derived	MYC	98R>W	chr8_128756735_128756735_C_T	Substitution	No	No	0.17%	11421	100	158	165	171	171
CGPLU208	Lung Cancer	II	Germine	STK11	354F>L	chr19_1223125_1223125_G_C	Substitution	NA	Yes	26.84%	11695	96	153	166	169	169
CGPLU208	Lung Cancer	II	Tumor derived	TP53	100Q>Y	chr17_7579389_7579389_G_A	Substitution	No	Yes	9.91%	12771	94	155	163	168	168
CGPLU208	Lung Cancer	II	Tumor derived	CDKN2A	88E>X	chr9_12911036_21971036_C_A	Substitution	Yes	Yes	9.13%	16557	92	157	169	170	170
CGPLU208	Lung Cancer	II	Tumor derived	PDGFRA	521A>S	chr7_55156502_55156502_G_A	Substitution	No	Yes	0.82%	13057	97	158	167	171	171
CGPLU208	Lung Cancer	II	Tumor derived	EGFR	567M>V	chr7_55231525_55231525_A_G	Substitution	NA	Yes	30.41%	8521	103	155	167	169	169
CGPLU208	Lung Cancer	II	Tumor derived	TP53	342R>X	chr17_7574003_7574003_G_A	Substitution	Yes	Yes	3.14%	4421	101	161	165	172	172
CGPLU208	Lung Cancer	II	Tumor derived	TP53	248R>Q	chr17_7574938_7574938_C_T	Substitution	Yes	Yes	0.87%	7987	103	157	164	169	169
CGPLU208	Lung Cancer	II	Tumor derived	TP53	634>V	chr17_7574949_7574949_G_A	Substitution	NA	Yes	37.77%	3732	97	160	166	171	171
CGPLU209	Ovarian Cancer	II	Tumor derived	TP53	551A>S	chr4_55156502_55156502_G_A	Substitution	No	Yes	0.12%	12072	88	157	165	168	168
CGPLU209	Ovarian Cancer	II	Tumor derived	TP53	567M>V	chr7_55231525_55231525_A_G	Substitution	NA	Yes	0.82%	3057	97	158	167	171	171
CGPLU209	Ovarian Cancer	II	Tumor derived	TP53	342R>X	chr17_7574003_7574003_G_A	Substitution	NA	Yes	30.41%	8521	103	155	167	169	169
CGPLO1010	Ovarian Cancer	II	Tumor derived	TP53	248R>Q	chr17_7574938_7574938_C_T	Substitution	Yes	Yes	3.14%	4421	101	161	165	172	172
CGPLO1011	Ovarian Cancer	II	Germine	ALK	100Q>Y	chr17_7579389_7579389_G_A	Substitution	No	No	0.14%	7987	103	157	164	169	169
CGPLO1013	Ovarian Cancer	II	Tumor derived	PDGFRA	444W>C	chr2_29851298_29851298_C_A	Substitution	No	Yes	37.77%	3732	97	160	166	171	171
CGPLO1013	Ovarian Cancer	II	Tumor derived	TP53	410A>D	chr4_55156502_55156502_G_A	Substitution	Yes	Yes	0.12%	12072	88	157	165	168	168
CGPLO1013	Ovarian Cancer	II	Tumor derived	TP53	435R>H	chr4_55156502_55156502_G_A	Substitution	No	Yes	37.88%	4107	103	159	166	171	171
CGPLO1014	Ovarian Cancer	II	Tumor derived	TP53	230E>K	chr12_121431484_121431484_G_A	Substitution	No	No	0.35%	6427	103	161	165	170	170
CGPLO1015	Ovarian Cancer	II	Tumor derived	TP53	278P>S	chr17_7577058_7577058_G_A	Substitution	Yes	Yes	0.14%	11418	92	154	167	171	171
CGPLO1015	Ovarian Cancer	II	Tumor derived	EGFR	428H>D	chr7_55225445_55225445_G_C	Substitution	Yes	Yes	3.56%	7688	102	158	164	169	169
CGPLO1015	Ovarian Cancer	II	Tumor derived	TP53	248R>Q	chr17_55156380_55156380_C_A	Substitution	No	Yes	0.19%	7617	101	159	167	171	171
CGPLO1017	Ovarian Cancer	II	Tumor derived	TP53	107D>N	chr4_55161380_55161380_G_A	Substitution	NA	Yes	0.32%	4463	96	158	165	169	169
CGPLO1017	Ovarian Cancer	II	Germine	APC	112S>T	chr5_112174625_112174625_T_C	Substitution	NA	Yes	44.10%	2884	110	157	170	170	170
CGPLO1018	Ovarian Cancer	II	Germine	FGRFR	403R>S	chr4_1806168_1806168_A_G	Substitution	NA	Yes	40.81%	2945	101	159	164	169	169
CGPLO1019	Ovarian Cancer	II	Tumor derived	TP53	273R>H	chr17_757720_757720_C_T	Substitution	Yes	Yes	23.80%	9727	95	158	167	172	172
CGPLO1019	Ovarian Cancer	II	Germine	AR	176S>R	chrX_66765516_66765516_C_A	Substitution	NA	Yes	36.83%	4387	100	158	165	169	169
CGPLO1019	Ovarian Cancer	II	Tumor derived	TP53	193S>T	chr17_757723_23_12175423_C_T	Substitution	Yes	Yes	65.29%	2775	93	161	171	171	171
CGPLO1020	Ovarian Cancer	II	Tumor derived	TP53	253R>K	chr7_55221714_55221714_G_C	Substitution	Yes	Yes	45.35%	3816	102	156	170	170	170
CGPLO1020	Ovarian Cancer	II	Germine	STK11	354F>L	chr19_1223125_1223125_G_G	Substitution	NA	Yes	0.24%	5404	94	158	165	170	170
CGPLO1021	Ovarian Cancer	II	Germine	TP53	275C>Y	chr17_757714_757714_C_G	Substitution	No	Yes	44.05%	3744	102	158	166	169	169
CGPLO1021	Ovarian Cancer	II	Tumor derived	ERBB4	692S>T	chr2_21253104_21253104_C_G	Substitution	No	No	7.68%	21823	81	158	166	170	170
CGPLO1022	Ovarian Cancer	II	Tumor derived	TP53	193H>P	chr17_7578271_7578271_G_G	Substitution	No	Yes	2.04%	18806	101	159	165	168	168
CGPLO1022	Ovarian Cancer	II	Tumor derived	CTNNB1	411T>A	chr3_41266124_41266124_A_G	Substitution	Yes	Yes	14.38%	10801	89	160	165	167	167
CGPLO1022	Ovarian Cancer	II	Tumor derived	CTNNB1	411T>A	chr3_41266124_41266124_A_G	Substitution	Yes	Yes	6.46%	11952	103	155	165	167	167
CGPLO1022	Ovarian Cancer	II	Tumor derived	CTNNB1	411T>A	chr3_41266124_41266124_A_G	Substitution	Yes	Yes	6.34%	12399	92	150	164	165	165

	Mutant Fragments										Adjusted P Value of Difference between Mutant and Wild-type cDNA Fragment Sizes (bp)														
	Mean cDNA Fragment Size (bp)	5th Percentile cDNA Fragment Size (bp)	Maximum cDNA Fragment Size (bp)	Distinct Coverage	Minimum cDNA Fragment Size (bp)	25th Percentile cDNA Fragment Size (bp)	Median cDNA Fragment Size (bp)	Mode cDNA Fragment Size (bp)	Mean cDNA Fragment Size (bp)	75th Percentile cDNA Fragment Size (bp)	Maximum cDNA Fragment Size (bp)	Fragment Size (bp)	Fragment Size (bp)	Mean cDNA Fragment Size (bp)	75th Percentile cDNA Fragment Size (bp)	Maximum cDNA Fragment Size (bp)	Fragment Size (bp)	Fragment Size (bp)	Mean cDNA Fragment Size (bp)	75th Percentile cDNA Fragment Size (bp)	Maximum cDNA Fragment Size (bp)	Fragment Size (bp)			
179	186	400	400	19	100	142	233	165	190	191	196	166	176	167	156	191	199	305	305	4.0	1.54	3.38	0.250		
182	186	400	400	21	132	166	182	152	167	169	166	148	166	166	177	177	183	399	399	0.0	0.000	5.88	0.000		
180	183	400	400	5411	92	5411	5411	92	166	166	166	166	166	166	166	177	177	183	383	383	-0.25	-0.874	5.37	0.009	
177	182	400	400	1903	100	1903	1903	100	148	148	148	148	148	148	148	155	155	191	398	398	-1.0	-0.009	3.80	0.025	
184	185	400	400	1344	138	1344	1344	138	155	167	170	168	168	168	168	189	189	191	398	398	1.0	0.009	3.80	0.025	
181	182	400	400	2106	100	1905	1905	100	153	153	153	153	153	153	153	166	166	187	366	366	1.0	0.009	2.95	0.025	
175	179	400	400	1951	101	1951	1951	101	149	149	149	149	149	149	149	175	175	182	397	397	0.0	0.052	2.95	0.025	
176	180	400	400	176	75	176	176	75	162	162	162	162	162	162	162	172	172	182	370	370	3.0	1.48	3.98	0.052	
177	182	400	400	177	28	171	170	28	130	130	139	139	139	139	139	164	165	175	345	345	2.0	-12.79	3.13	0.002	
163	188	399	399	6863	100	6863	6863	100	160	160	168	168	168	168	168	173	185	185	400	400	0.0	0.002	3.13	0.002	
188	186	400	34	188	77	154	154	77	171	171	170	170	170	170	170	177	177	192	335	335	-0.5	-11.46	2.54	0.475	
175	179	396	9	175	136	147	176	136	171	171	176	176	176	176	176	177	177	176	390	390	4.0	1.22	3.80	0.025	
184	185	400	21	184	115	185	185	21	115	145	155	155	155	155	155	159	176	176	368	368	-11.0	-7.99	3.49	0.052	
176	183	399	75	176	123	176	176	75	162	162	172	172	172	172	172	190	190	196	370	370	3.0	3.98	3.49	0.052	
179	182	397	44	179	162	397	397	44	125	155	155	155	155	155	155	169	169	194	338	338	0.0	0.000	5.78	0.623	
185	186	400	400	8167	101	8167	8167	101	160	160	166	166	166	166	166	171	184	187	400	400	-1.0	-1.27	2.12	0.114	
187	186	400	400	3662	102	3662	3662	102	158	158	168	168	168	168	168	170	185	185	399	399	0.0	-2.62	2.15	0.475	
184	187	399	15	184	93	187	187	15	137	137	127	127	127	127	127	174	173	193	261	261	3.0	-11.00	3.07	0.507	
183	185	400	26	183	163	185	185	26	137	149	181	181	181	181	181	162	182	182	364	364	-3.0	-4.34	4.34	0.430	
181	182	397	35	181	118	182	182	35	118	147	176	176	176	176	176	163	172	172	336	336	-6.0	-9.35	6.68	0.166	
172	175	400	71	172	133	186	186	71	133	152	176	176	176	176	176	165	189	189	301	301	0.0	-3.57	2.12	0.475	
169	174	400	55	169	130	186	186	55	130	153	175	175	175	175	175	164	186	186	325	325	0.0	-6.68	6.68	0.114	
189	187	399	17	189	148	187	187	17	148	155	176	176	176	176	176	166	180	180	301	301	-3.0	-2.15	2.15	0.475	
176	183	400	18	176	156	185	185	18	170	170	174	174	174	174	174	174	174	174	216	216	-0.19	3.07	3.07	0.368	
169	175	397	51	169	128	175	175	51	143	143	268	268	268	268	268	152	164	164	268	268	-12.0	-5.12	5.12	0.000	
166	173	397	28	166	118	173	173	28	147	147	153	153	153	153	153	166	174	174	327	327	9.5	-8.37	8.37	0.036	
184	186	400	45	184	116	186	186	45	116	151	168	168	168	168	168	166	183	183	302	302	-3.0	-6.74	6.74	0.064	
165	186	400	25	165	157	186	186	25	130	153	185	185	185	185	185	191	175	175	346	346	-0.37	-8.84	8.84	0.057	
182	182	399	42	182	138	185	185	42	124	155	166	166	166	166	166	164	185	185	305	305	3.0	-2.15	2.15	0.475	
185	187	399	25	185	124	185	185	25	126	147	162	162	162	162	162	166	184	184	338	338	8.0	-4.51	4.51	0.171	
167	175	394	68	167	121	175	175	68	121	155	166	166	166	166	166	166	175	175	305	305	-0.19	0.234	0.234	0.445	
167	173	397	45	167	124	173	173	45	124	143	197	197	197	197	197	162	182	182	377	377	-1.0	-0.91	0.482	0.445	
170	175	398	103	170	126	186	186	103	126	147	162	162	162	162	162	166	184	184	302	302	-3.0	-6.74	6.74	0.064	
181	185	400	23	181	116	186	186	23	131	148	145	145	145	145	145	166	181	181	305	305	3.0	-6.68	6.68	0.057	
190	189	399	625	190	138	186	186	625	100	155	167	167	167	167	167	174	174	187	343	343	5.5	-4.51	4.51	0.171	
182	182	399	42	182	138	186	186	42	124	143	142	142	142	142	142	166	186	186	321	321	-1.0	0.06	0.06	0.445	
185	187	399	25	185	126	186	186	25	126	153	176	176	176	176	176	176	188	188	305	305	-0.19	0.234	0.234	0.445	
181	187	399	977	181	161	187	187	977	101	149	189	189	189	189	189	169	188	188	305	305	-1.0	-9.76	9.76	0.000	
173	179	391	111	173	160	184	184	111	101	146	166	166	166	166	166	166	181	181	305	305	-3.0	-5.57	5.57	0.032	
181	185	399	401	181	100	186	186	401	100	158	166	166	166	166	166	166	181	181	305	305	3.0	-6.74	6.74	0.064	
178	179	398	31	178	116	184	184	31	85	146	137	137	137	137	137	137	166	181	181	305	305	-0.37	0.770	7.70	0.057
184	184	400	24	184	116	184	184	24	116	146	146	146	146	146	146	166	181	181	305	305	-0.37	0.770	7.70	0.057	
185	186	399	309	185	117	184	184	309	116	147	122	122	122	122	122	161	181	181	305	305	-3.0	-2.15	2.15	0.475	
171	173	400	27	171	117	184	184	27	116	147	161	161	161	161	161	161	181	181	305	305	-0.19	0.009	0.009	0.475	
176	178	399	31	176	116	184	184	31	85	146	137	137	137	137	137	137	166	181	181	305	305	-0.37	0.770	7.70	0.057
182	182	399	395	182	100	184	184	395	100	146	135	135	135	135	135	138	166	181	181	305	305	-3.0	-2.15	2.15	0.475
181	181	399	395	181	100	184	184	395	100	146	135	135	135	135	135	138	166	181	181	305	305	-0.19	0.009	0.009	0.475
181	181	399	395	181	100	184	184	395	100	146	135	135	135	135	135	138	166	181	181	305	305	-0.37	0.770	7.70	0.057
181	181	399</td																							

Fragment Size (bp)	Mean cfDNA	75th Percentile	Maximum cfDNA	Mutant Fragments			Wild-type cfDNA			Mutant and Wild-type cfDNA			Difference between Mean Fragment Sizes (bp)		Adjusted P Value of Difference between Median Fragment Sizes (bp)		Difference between Mean Fragment Sizes (bp)	Difference between Median Fragment Sizes (bp)	
				Distinct Coverage	Median cfDNA	Mode cfDNA	Minimum cfDNA	25th Percentile	Median cfDNA	Mode cfDNA	75th Percentile	Maximum cfDNA	Mean cfDNA	Median cfDNA	Mode cfDNA	75th Percentile	Maximum cfDNA		
156	172	396	1616	100	146	164	159	163	170	164	173	176	354	354	354	35	-3.57	0.000	0.054
175	180	400	806	96	158	169	159	173	164	154	164	170	398	398	398	40	-8.36	0.816	0.000
165	172	399	1410	102	140	149	143	154	182	154	164	170	333	333	333	16.0	36.25	0.000	0.104
170	177	397	49	99	153	143	143	155	154	170	155	160	284	284	284	14.38	-6.66	0.000	0.479
166	173	396	33	140	155	154	140	140	140	155	173	176	296	296	296	7.0	7.0	0.000	0.411
180	178	400	73	95	140	140	140	140	140	164	167	182	324	324	324	15	10.93	0.000	0.714
172	177	400	38	115	160	164	164	164	164	175	178	181	329	329	329	15	-18.96	0.000	0.714
171	174	386	6	124	137	137	124	124	124	156	153	168	178	178	178	7.5	7.5	0.000	0.714
180	183	400	70	124	151	151	151	151	151	164	183	183	385	385	385	6.0	1.71	0.064	0.714
194	199	389	6556	96	162	168	175	193	196	196	196	196	399	399	399	0.0	-1.79	0.166	0.000
184	188	400	41	112	172	176	177	175	175	175	195	195	373	373	373	3.0	11.62	0.397	0.000
194	196	399	35	145	168	168	168	168	168	175	181	186	312	312	312	1.0	-13.40	0.587	0.000
182	184	399	20	166	180	185	191	205	205	205	219	219	357	357	357	23.48	23.48	0.013	0.984
163	186	397	5338	102	159	175	171	183	185	185	185	185	394	394	394	1.0	0.03	0.000	0.984
202	203	393	178	101	150	168	171	198	198	198	240	240	357	357	357	5.0	4.34	0.051	0.479
195	196	397	1380	104	153	163	171	201	201	201	258	258	400	400	400	0.0	5.94	0.036	0.000
185	189	400	1257	100	153	168	170	188	188	188	202	202	392	392	392	1.0	4.37	0.064	0.000
185	189	396	36	117	163	164	172	175	175	175	179	179	372	372	372	3.0	-10.29	0.463	0.000
203	210	391	336	105	153	141	171	200	200	200	240	240	399	399	399	4.0	-3.10	0.080	0.000
188	194	399	744	101	161	169	176	190	194	194	200	200	400	400	400	2.0	1.96	0.061	0.571
193	199	396	89	100	145	171	171	197	197	197	228	228	393	393	393	2.0	3.42	0.084	0.000
172	179	396	12	129	143	143	153	163	163	163	186	186	275	275	275	14.0	-8.89	0.000	0.000
186	188	387	3659	91	156	164	173	195	195	195	211	211	398	398	398	8.92	0.001	0.000	0.000
177	183	392	873	102	149	163	163	164	164	164	175	175	181	181	181	3.0	-0.39	0.039	0.000
194	200	377	1909	100	158	167	176	202	202	202	242	242	398	398	398	5.0	7.98	0.061	0.559
202	202	27	122	157	164	174	179	198	198	198	231	231	356	356	356	2.0	-3.82	0.685	0.000
171	178	385	1818	103	147	169	169	182	182	182	212	212	360	360	360	1.0	1.92	0.372	0.000
178	182	374	546	102	151	166	166	186	186	186	204	204	381	381	381	0.0	2.87	0.416	0.000
179	184	387	26	132	142	138	171	183	183	183	201	201	351	351	351	1.5	3.26	0.572	0.000
195	194	400	53	117	157	166	169	192	192	192	206	206	396	396	396	3.0	-2.06	0.451	0.000
178	179	397	40	124	150	166	169	186	186	186	216	216	384	384	384	1.0	4.53	0.539	0.000
188	191	390	38	107	153	180	180	194	194	194	210	210	326	326	326	0.5	-2.58	0.576	0.000
205	207	399	217	102	146	144	144	163	163	163	212	212	360	360	360	-12.0	-17.11	0.004	0.000
196	195	397	266	111	147	150	150	166	166	166	204	204	379	379	379	8.0	-7.53	0.208	0.000
186	188	400	76	123	157	171	171	190	190	190	204	204	346	346	346	1.0	-3.64	0.479	0.000
186	188	398	93	161	166	166	166	172	172	172	198	198	399	399	399	1.0	0.155	0.000	0.000
191	191	390	38	107	153	180	180	194	194	194	216	216	384	384	384	3.0	9.95	0.061	0.000
187	189	397	266	111	147	150	150	166	166	166	211	211	387	387	387	1.0	7.98	0.560	0.000
202	202	400	102	138	161	161	161	179	179	179	201	201	400	400	400	2.0	14.14	0.341	0.000
196	196	391	932	93	161	166	166	172	172	172	198	198	372	372	372	1.5	2.90	0.587	0.000
181	182	387	30	138	158	168	168	185	185	185	191	191	311	311	311	1.0	9.25	0.679	0.000
181	181	400	277	104	162	162	162	176	176	176	201	201	384	384	384	3.0	-2.85	0.679	0.000
189	191	400	65	123	185	185	185	198	198	198	211	211	387	387	387	1.0	10.89	0.314	0.000
187	189	400	31	136	163	171	171	187	187	187	201	201	398	398	398	0.5	-3.83	0.015	0.000
202	202	400	5286	102	166	168	168	181	181	181	201	201	400	400	400	2.0	2.00	0.571	0.000
196	196	400	102	138	161	161	161	179	179	179	199	199	372	372	372	1.0	8.77	0.000	0.000
181	181	387	30	138	158	168	168	185	185	185	191	191	311	311	311	1.0	9.25	0.679	0.000
181	181	400	64	113	158	163	163	176	176	176	197	197	344	344	344	9.0	3.52	0.564	0.000
176	176	398	27	121	149	160	160	176	176	176	197	197	392	392	392	5.0	10.77	0.314	0.000
191	192	398	2943	100	165	165	165	176	176	176	197	197	398	398	398	3.0	-0.01	0.263	0.000
176	176	385	25	136	153	153	153	166	166	166	187	187	373	373	373	1.0	0.610	0.588	0.000
176	176	400	28	110	136	136	136	147	147	147	161	161	340	340	340	0.0	1.78	0.314	0.000
182	184	400	321	131	160	168	168	177	177	177	197	197	388	388	388	-5.63	0.463	0.839	0.000
175	177	399	26	139	147	147	147	160	160	160	176	176	327	327	327	-19.0	8.77	0.000	0.000
172	172	399	15	121	146	146	146	153	153	153	161	161	344	344	344	9.0	3.52	0.564	0.000
186	184	398	35	121	149	162	162	176	176	176	197	197	392	392	392	1.0	1.12	0.588	0.000
189	191	400	86	121	165	165	165	177	177	177	197	197	398	398	398	-0.32	-0.32	0.839	0.000
176	176	389	3339	101	157	164	164	175	175	175	197	197	398	398	398	0.0	0.00	0.263	0.000
176	176	400	209	125	160	160	160	176	176	176	197	197	373	373	373	1.0	12.56	0.314	0.000
211	211	393	41	156	176	176	176	197	197	197	215	215	398						

Fragment Size (bp)	Mean cfDNA Fragment Size (bp)	75th Percentile cfDNA Fragment Size (bp)	Maximum cfDNA Fragment Size (bp)	Minimum cfDNA Fragment Size (bp)	Distinct Coverage	Median cfDNA Fragment Size (bp)	Mode cfDNA Fragment Size (bp)	25th Percentile cfDNA Fragment Size (bp)	Mean cfDNA Fragment Size (bp)	Median cfDNA Fragment Size (bp)	75th Percentile cfDNA Fragment Size (bp)	Maximum cfDNA Fragment Size (bp)	Adjusted P Value of Difference between Mean and Wild-type cfDNA Fragment Sizes	
195	400	3096	79	159	161	173	164	191	397	397	191	191	-2.45	0.251
202	203	73	142	178	176	184	237	338	186	186	186	186	-16.38	0.114
205	203	23	161	168	168	171	186	186	190	190	190	190	-6.17	0.435
196	400	170	125	158	173	173	168	168	190	190	190	190	-8.80	0.293
195	196	400	2089	101	162	169	176	203	203	203	203	203	4.45	0.000
195	192	400	125	84	192	194	207	243	243	243	243	243	16.0	0.574
238	280	400	125	84	192	194	207	243	243	243	243	243	5.51	0.065
197	194	400	5715	108	163	164	174	206	206	206	206	206	1.0	0.190
172	173	398	103	78	146	149	158	166	173	173	173	173	-5.94	0.880
196	191	329	35	119	161	172	171	191	186	186	186	186	-4.34	0.697
189	190	400	826	162	162	166	171	187	187	187	187	187	-1.94	0.475
194	195	400	95	135	160	161	170	182	184	184	184	184	-5.0	0.155
184	184	400	27	128	150	150	169	174	185	185	185	185	-9.68	0.571
179	184	399	4771	103	161	168	171	178	183	183	183	183	0.0	0.0
167	185	399	7	147	154	154	167	164	174	174	174	174	-1.37	0.155
179	179	395	330	106	152	166	166	178	178	178	178	178	-2.90	0.252
172	177	399	536	106	151	167	163	172	175	175	175	175	-1.35	0.685
179	183	400	45	136	163	175	172	185	191	191	191	191	-6.34	0.880
182	182	397	18	136	146	146	155	162	170	170	170	170	-14.0	0.007
172	177	397	293	101	152	169	164	170	174	174	174	174	-19.82	0.646
171	177	399	23	130	152	162	162	163	177	177	177	177	-3.0	0.0
180	183	399	54	104	161	154	176	195	206	206	206	206	-7.62	0.064
184	184	400	154	96	148	157	163	176	185	185	185	185	-5.5	0.154
186	187	399	79	102	163	177	174	185	200	200	200	200	-2.70	0.270
183	185	400	44	118	148	148	163	163	188	188	188	188	-1.98	0.039
182	184	400	35	136	164	204	181	194	203	203	203	203	-11.80	0.039
191	191	400	13	138	164	169	169	198	198	198	198	198	-1.0	0.610
192	191	400	50	128	155	161	171	216	301	360	360	360	-17.02	0.623
191	193	400	81	108	150	108	173	198	224	385	385	385	-0.0	0.624
190	191	389	2587	101	159	165	172	185	187	187	187	187	-5.17	0.005
192	197	400	58	92	173	192	192	202	202	202	202	202	-9.79	0.007
183	188	400	74	90	147	142	167	176	182	182	182	182	-6.5	0.061
175	191	400	37	144	163	185	172	192	192	192	192	192	-17.15	0.005
194	194	400	93	164	181	181	181	197	211	370	370	370	-3.34	0.169
184	186	400	66	104	158	194	174	183	194	194	194	194	-4.60	0.270
191	190	396	101	126	155	176	176	194	213	331	331	331	-2.50	0.718
192	197	400	58	92	156	164	168	187	193	193	193	193	-2.54	0.113
188	188	400	74	90	147	142	167	176	182	182	182	182	-6.78	0.302
175	191	400	37	144	163	185	172	192	192	192	192	192	-16.65	0.407
194	202	400	34	139	163	165	170	178	175	349	349	349	-3.0	0.876
184	184	400	61	108	150	152	165	181	186	393	393	393	-4.0	0.926
191	191	396	101	126	155	176	176	194	213	331	331	331	-6.15	0.234
192	197	400	55	121	158	161	167	186	186	186	186	186	-9.98	0.130
188	188	400	16	144	172	179	179	187	187	187	187	187	-17.73	0.154
186	186	399	30	134	161	175	175	190	208	339	339	339	-1.0	0.213
180	180	397	34	139	163	165	170	178	175	349	349	349	-4.0	0.179
182	182	400	262	101	150	152	165	181	186	393	393	393	-4.47	0.427
182	182	400	277	101	150	147	166	182	185	393	393	393	-3.0	0.415
180	182	395	65	121	158	161	167	186	186	338	338	338	-4.0	0.252
177	182	400	16	144	172	179	179	187	187	187	187	187	-17.67	0.000
185	184	399	36	131	147	147	143	151	175	181	181	181	-1.73	0.054
184	185	400	46	109	154	164	164	188	188	188	188	188	-5.19	0.252
181	181	394	21	108	144	144	144	173	200	357	357	357	-0.5	0.119
182	180	400	18	111	127	127	127	158	188	352	352	352	-4.47	0.427
179	181	400	72	121	156	173	173	186	186	396	396	396	-2.0	0.415
177	182	400	30	106	160	174	174	180	180	282	282	282	-2.0	0.252
175	180	399	36	131	147	147	143	151	175	181	181	181	-4.24	0.479
184	185	392	21	108	144	144	144	173	206	367	367	367	-0.5	0.000
182	184	395	16	147	156	156	164	186	186	186	186	186	-6.42	0.874
176	180	392	2742	102	154	164	148	150	152	162	162	162	-22.25	0.000
174	181	399	298	103	146	146	148	164	173	173	173	173	-9.89	0.425
186	186	396	5	116	182	182	185	185	187	187	187	187	-27.67	0.000
186	183	399	1073	100	142	164	164	165	165	165	165	165	-5.52	0.054
179	184	400	46	109	151	143	143	175	174	183	183	183	-14.58	0.252
181	181	394	21	108	146	146	146	173	205	367	367	367	-0.5	0.000
182	182	400	30	146	146	146	146	173	205	367	367	367	-6.42	0.874
176	183	392	2742	102	154	164	148	150	152	162	162	162	-22.25	0.000
174	184	397	34	109	156	156	164	186	186	186	186	186	-9.89	0.425
186	186	395	50	130	161	175	175	188	188	188	188	188	-27.67	0.000
186	187	398	28	139	150	150	150	173	205	367	367	367	-0.5	0.054
179	184	400	24	136	153	153	153	176	207	367	367	367	-14.58	0.252
185	185	394	48	111	154	154	154	176	207	367	367	367	-6.42	0.874
181	187	397	103	130	161	175	175	188	188	188	188	188	-22.25	0.000
186	187	398	28	139	150	150	150	173	205	367	367	367	-0.5	0.054
179	184	400	24	136	153	153	153	176	207	367	367	367	-14.58	0.252
185	185	394	48	111	154	154	154	176	207	367	367	367	-6.42	0.874
181	187	397	103	130	161	175	175	188	188	188	188	188	-22.25	0.000
186	187	398	28	139	150	150	150	173	205	367	367	367	-0.5	0.054
179	184	400	24	136	153	153	153	176	207	367	367	367	-14.58	0.252
1														

APPENDIX - D: Table 4. Summary of whole genome cfDNA analyses

Patient	Timepoint	Analysis type	Patient Type	Read Length	Total Bases Sequenced	High Quality Bases Analyzed	Coverage
CGCRC291	Preoperative treatment naïve	WGS	Colorectal Cancer	100	7232125000	4695396600	1.86
CGCRC292	Preoperative treatment naïve	WGS	Colorectal Cancer	100	6794092800	4471065400	1.77
CGCRC293	Preoperative treatment naïve	WGS	Colorectal Cancer	100	8373899600	5686176000	2.26
CGCRC294	Preoperative treatment naïve	WGS	Colorectal Cancer	100	8081312000	5347045800	2.12
CGCRC296	Preoperative treatment naïve	WGS	Colorectal Cancer	100	10072029200	6770998200	2.69
CGCRC299	Preoperative treatment naïve	WGS	Colorectal Cancer	100	10971591600	7632723200	3.03
CGCRC300	Preoperative treatment naïve	WGS	Colorectal Cancer	100	9894332600	6699951000	2.66
CGCRC301	Preoperative treatment naïve	WGS	Colorectal Cancer	100	7857346200	5021002000	1.99
CGCRC302	Preoperative treatment naïve	WGS	Colorectal Cancer	100	11671913000	8335275800	3.31
CGCRC304	Preoperative treatment naïve	WGS	Colorectal Cancer	100	19011739200	12957614200	5.14
CGCRC305	Preoperative treatment naïve	WGS	Colorectal Cancer	100	7177341400	4809957200	1.91
CGCRC306	Preoperative treatment naïve	WGS	Colorectal Cancer	100	8302233200	5608043600	2.23
CGCRC307	Preoperative treatment naïve	WGS	Colorectal Cancer	100	8034729400	5342620000	2.12
CGCRC308	Preoperative treatment naïve	WGS	Colorectal Cancer	100	8670084800	5934037200	2.35
CGCRC311	Preoperative treatment naïve	WGS	Colorectal Cancer	100	6947634400	4704501800	1.87
CGCRC315	Preoperative treatment naïve	WGS	Colorectal Cancer	100	5205544000	3419565400	1.36
CGCRC316	Preoperative treatment naïve	WGS	Colorectal Cancer	100	6405368600	4447534800	1.76
CGCRC317	Preoperative treatment naïve	WGS	Colorectal Cancer	100	6060390400	4104616800	1.63
CGCRC318	Preoperative treatment naïve	WGS	Colorectal Cancer	100	6848768600	4439404800	1.76
CGCRC319	Preoperative treatment naïve	WGS	Colorectal Cancer	100	10545294400	7355181600	2.92
CGCRC320	Preoperative treatment naïve	WGS	Colorectal Cancer	100	5961999200	3945054000	1.57
CGCRC321	Preoperative treatment naïve	WGS	Colorectal Cancer	100	8248054000	5614355000	2.23
CGCRC333	Preoperative treatment naïve	WGS	Colorectal Cancer	100	10540267600	6915490600	2.74
CGCRC336	Preoperative treatment naïve	WGS	Colorectal Cancer	100	10675581800	7087691800	2.81
CGCRC338	Preoperative treatment naïve	WGS	Colorectal Cancer	100	13788172600	8970308600	3.56
CGCRC341	Preoperative treatment naïve	WGS	Colorectal Cancer	100	10753467600	7311539200	2.90
CGCRC342	Preoperative treatment naïve	WGS	Colorectal Cancer	100	11836966000	7552793200	3.00
CGH14	Human adult elutriated lymphocytes	WGS	Healthy	100	36525427600	24950300200	9.90
CGH15	Human adult elutriated lymphocytes	WGS	Healthy	100	29930855000	23754049400	9.43
CGLU316	Pre-treatment, Day -53	WGS	Lung Cancer	100	10354123200	6896471400	2.74
CGLU316	Pre-treatment, Day -4	WGS	Lung Cancer	100	7870039200	5254938800	2.09
CGLU316	Post-treatment, Day 18	WGS	Lung Cancer	100	8155322000	5416262400	2.15
CGLU316	Post-treatment, Day 87	WGS	Lung Cancer	100	9442310400	6087893400	2.42
CGLU344	Pre-treatment, Day -21	WGS	Lung Cancer	100	8728318600	5769097200	2.29
CGLU344	Pre-treatment, Day 0	WGS	Lung Cancer	100	11710249400	7826902600	3.11
CGLU344	Post-treatment, Day 0,1875	WGS	Lung Cancer	100	11569683000	7654701600	3.04
CGLU344	Post-treatment, Day 59	WGS	Lung Cancer	100	11042459200	6320138800	2.51
CGLU369	Pre-treatment, Day -2	WGS	Lung Cancer	100	8636932800	5779595800	2.29
CGLU369	Post-treatment, Day 12	WGS	Lung Cancer	100	9227709600	6136755200	2.44
CGLU369	Post-treatment, Day 68	WGS	Lung Cancer	100	7995282600	5239077200	2.08
CGLU369	Post-treatment, Day 110	WGS	Lung Cancer	100	8750541000	5626139000	2.23
CGLU373	Pre-treatment, Day -2	WGS	Lung Cancer	100	11746059600	7547485800	3.00
CGLU373	Post-treatment, Day 0,125	WGS	Lung Cancer	100	13801136800	9255579400	3.67
CGLU373	Post-treatment, Day 7	WGS	Lung Cancer	100	11537896800	7654111200	3.04
CGLU373	Post-treatment, Day 47	WGS	Lung Cancer	100	8046326400	5397702400	2.14
CGPLBR100	Preoperative treatment naïve	WGS	Breast Cancer	100	8440532400	5729474800	2.27
CGPLBR101	Preoperative treatment naïve	WGS	Breast Cancer	100	9786253600	6673495200	2.65
CGPLBR102	Preoperative treatment naïve	WGS	Breast Cancer	100	8664980400	5669781800	2.25
CGPLBR103	Preoperative treatment naïve	WGS	Breast Cancer	100	9846936200	6662883400	2.64
CGPLBR104	Preoperative treatment naïve	WGS	Breast Cancer	100	9443375400	6497061000	2.58
CGPLBR12	Preoperative treatment naïve	WGS	Breast Cancer	100	7017577800	4823327400	1.91
CGPLBR18	Preoperative treatment naïve	WGS	Breast Cancer	100	10309652800	7130386000	2.83
CGPLBR23	Preoperative treatment naïve	WGS	Breast Cancer	100	9034484800	6219625800	2.47
CGPLBR24	Preoperative treatment naïve	WGS	Breast Cancer	100	9891454200	6601857400	2.62
CGPLBR28	Preoperative treatment naïve	WGS	Breast Cancer	100	7997607200	5400803200	2.14
CGPLBR30	Preoperative treatment naïve	WGS	Breast Cancer	100	8502597200	5885822400	2.34
CGPLBR31	Preoperative treatment naïve	WGS	Breast Cancer	100	12660085600	8551995600	3.39
CGPLBR32	Preoperative treatment naïve	WGS	Breast Cancer	100	8773498600	5839034600	2.32
CGPLBR33	Preoperative treatment naïve	WGS	Breast Cancer	100	10931742800	6967030600	2.76
CGPLBR34	Preoperative treatment naïve	WGS	Breast Cancer	100	10861398600	7453225800	2.96
CGPLBR35	Preoperative treatment naïve	WGS	Breast Cancer	100	9180193600	6158440200	2.44
CGPLBR36	Preoperative treatment naïve	WGS	Breast Cancer	100	9159948400	6091817800	2.42
CGPLBR37	Preoperative treatment naïve	WGS	Breast Cancer	100	10307505800	6929530600	2.75
CGPLBR38	Preoperative treatment naïve	WGS	Breast Cancer	100	9983824000	6841725400	2.71
CGPLBR40	Preoperative treatment naïve	WGS	Breast Cancer	100	10148823800	7024345400	2.79
CGPLBR41	Preoperative treatment naïve	WGS	Breast Cancer	100	11168192000	7562945800	3.00
CGPLBR45	Preoperative treatment naïve	WGS	Breast Cancer	100	8793780600	6011109400	2.39
CGPLBR46	Preoperative treatment naïve	WGS	Breast Cancer	100	7228607600	4706130000	1.87
CGPLBR47	Preoperative treatment naïve	WGS	Breast Cancer	100	7906911400	5341655000	2.12
CGPLBR48	Preoperative treatment naïve	WGS	Breast Cancer	100	6992032000	4428636200	1.76
CGPLBR49	Preoperative treatment naïve	WGS	Breast Cancer	100	7311195000	4559460200	1.81
CGPLBR50	Preoperative treatment naïve	WGS	Breast Cancer	100	11107960600	7582776600	3.01

Patient	Timepoint	Analysis type	Patient Type	Read Length	Total Bases Sequenced	High Quality Bases Analyzed	Coverage
CGPLBR51	Preoperative treatment naïve	WGS	Breast Cancer	100	8393547400	5102369000	2.02
CGPLBR52	Preoperative treatment naïve	WGS	Breast Cancer	100	9491894800	6141729000	2.44
CGPLBR55	Preoperative treatment naïve	WGS	Breast Cancer	100	9380109800	6518855200	2.59
CGPLBR56	Preoperative treatment naïve	WGS	Breast Cancer	100	12191816800	8293011200	3.29
CGPLBR57	Preoperative treatment naïve	WGS	Breast Cancer	100	9847584400	6713638000	2.66
CGPLBR59	Preoperative treatment naïve	WGS	Breast Cancer	100	7476477000	5059878200	2.01
CGPLBR60	Preoperative treatment naïve	WGS	Breast Cancer	100	6531354600	4331253800	1.72
CGPLBR61	Preoperative treatment naïve	WGS	Breast Cancer	100	9311029200	6430520800	2.55
CGPLBR63	Preoperative treatment naïve	WGS	Breast Cancer	100	8971949000	6044009600	2.40
CGPLBR65	Preoperative treatment naïve	WGS	Breast Cancer	100	7197301400	4835015200	1.92
CGPLBR68	Preoperative treatment naïve	WGS	Breast Cancer	100	10003774000	6974918800	2.77
CGPLBR69	Preoperative treatment naïve	WGS	Breast Cancer	100	10080881800	6903459200	2.74
CGPLBR70	Preoperative treatment naïve	WGS	Breast Cancer	100	8824002800	6002533800	2.38
CGPLBR71	Preoperative treatment naïve	WGS	Breast Cancer	100	10164136800	6994668600	2.78
CGPLBR72	Preoperative treatment naïve	WGS	Breast Cancer	100	18416841400	12328783000	4.89
CGPLBR73	Preoperative treatment naïve	WGS	Breast Cancer	100	10281460200	7078613200	2.81
CGPLBR76	Preoperative treatment naïve	WGS	Breast Cancer	100	10105270400	6800705000	2.70
CGPLBR81	Preoperative treatment naïve	WGS	Breast Cancer	100	5087126000	3273367200	1.30
CGPLBR82	Preoperative treatment naïve	WGS	Breast Cancer	100	10576496600	7186662600	2.85
CGPLBR83	Preoperative treatment naïve	WGS	Breast Cancer	100	8977124400	5947525000	2.36
CGPLBR84	Preoperative treatment naïve	WGS	Breast Cancer	100	6272538600	4066870600	1.61
CGPLBR87	Preoperative treatment naïve	WGS	Breast Cancer	100	8460954800	5375710200	2.13
CGPLBR88	Preoperative treatment naïve	WGS	Breast Cancer	100	8665810400	5499898200	2.18
CGPLBR90	Preoperative treatment naïve	WGS	Breast Cancer	100	6663469200	4392442400	1.74
CGPLBR91	Preoperative treatment naïve	WGS	Breast Cancer	100	10933002400	7647842000	3.03
CGPLBR92	Preoperative treatment naïve	WGS	Breast Cancer	100	10392674000	6493598000	2.58
CGPLBR93	Preoperative treatment naïve	WGS	Breast Cancer	100	5659836000	3931106800	1.56
CGPLH189	Preoperative treatment naïve	WGS	Healthy	100	11400610400	7655568800	3.04
CGPLH190	Preoperative treatment naïve	WGS	Healthy	100	11444671600	7581175200	3.01
CGPLH192	Preoperative treatment naïve	WGS	Healthy	100	12199010800	8126804800	3.22
CGPLH193	Preoperative treatment naïve	WGS	Healthy	100	10201897600	6635285400	2.63
CGPLH194	Preoperative treatment naïve	WGS	Healthy	100	11005087400	7081852600	2.81
CGPLH196	Preoperative treatment naïve	WGS	Healthy	100	12891462600	8646881800	3.43
CGPLH197	Preoperative treatment naïve	WGS	Healthy	100	11961841600	8052855200	3.20
CGPLH198	Preoperative treatment naïve	WGS	Healthy	100	13605489000	8885716000	3.53
CGPLH199	Preoperative treatment naïve	WGS	Healthy	100	1818090200	5615316000	2.23
CGPLH200	Preoperative treatment naïve	WGS	Healthy	100	14400027600	9310342000	3.69
CGPLH201	Preoperative treatment naïve	WGS	Healthy	100	6208766800	4171848400	1.66
CGPLH202	Preoperative treatment naïve	WGS	Healthy	100	11282922800	7363530600	2.92
CGPLH203	Preoperative treatment naïve	WGS	Healthy	100	13540689600	9068747600	3.60
CGPLH205	Preoperative treatment naïve	WGS	Healthy	100	10343537800	6696988600	2.66
CGPLH208	Preoperative treatment naïve	WGS	Healthy	100	12796300000	8272073400	3.28
CGPLH209	Preoperative treatment naïve	WGS	Healthy	100	13123035400	8531813600	3.39
CGPLH210	Preoperative treatment naïve	WGS	Healthy	100	10184218800	6832204600	2.71
CGPLH211	Preoperative treatment naïve	WGS	Healthy	100	14655260200	8887067600	3.53
CGPLH300	Preoperative treatment naïve	WGS	Healthy	100	7062083400	4553351200	1.81
CGPLH307	Preoperative treatment naïve	WGS	Healthy	100	7239128200	4547697200	1.80
CGPLH308	Preoperative treatment naïve	WGS	Healthy	100	8512551400	5526653600	2.19
CGPLH309	Preoperative treatment naïve	WGS	Healthy	100	11664474200	7431836600	2.95
CGPLH310	Preoperative treatment naïve	WGS	Healthy	100	11045691000	7451506200	2.96
CGPLH311	Preoperative treatment naïve	WGS	Healthy	100	10406803200	6786479600	2.69
CGPLH314	Preoperative treatment naïve	WGS	Healthy	100	10371343800	6925866600	2.75
CGPLH315	Preoperative treatment naïve	WGS	Healthy	100	9508538400	6208744600	2.46
CGPLH316	Preoperative treatment naïve	WGS	Healthy	100	10131063600	6891181000	2.73
CGPLH317	Preoperative treatment naïve	WGS	Healthy	100	8364314400	5302232600	2.10
CGPLH319	Preoperative treatment naïve	WGS	Healthy	100	8780528200	5585897000	2.22
CGPLH320	Preoperative treatment naïve	WGS	Healthy	100	8956232600	5784619200	2.30
CGPLH322	Preoperative treatment naïve	WGS	Healthy	100	9563837800	6445517800	2.56
CGPLH324	Preoperative treatment naïve	WGS	Healthy	100	6765038600	4469201600	1.77
CGPLH325	Preoperative treatment naïve	WGS	Healthy	100	8008213400	5099262800	2.02
CGPLH326	Preoperative treatment naïve	WGS	Healthy	100	9554226200	6112544800	2.43
CGPLH327	Preoperative treatment naïve	WGS	Healthy	100	8239168800	5351280200	2.12
CGPLH328	Preoperative treatment naïve	WGS	Healthy	100	7197086800	4516894800	1.79
CGPLH329	Preoperative treatment naïve	WGS	Healthy	100	8921554800	5493709800	2.18
CGPLH330	Preoperative treatment naïve	WGS	Healthy	100	10693603400	7077793600	2.81
CGPLH331	Preoperative treatment naïve	WGS	Healthy	100	8982792000	5538096200	2.20
CGPLH333	Preoperative treatment naïve	WGS	Healthy	100	7856985400	5178829600	2.06
CGPLH335	Preoperative treatment naïve	WGS	Healthy	100	9370663400	6035739400	2.40
CGPLH336	Preoperative treatment naïve	WGS	Healthy	100	8002498200	5340331400	2.12
CGPLH337	Preoperative treatment naïve	WGS	Healthy	100	7399022000	4954467600	1.97
CGPLH338	Preoperative treatment naïve	WGS	Healthy	100	8917121600	6170927200	2.45
CGPLH339	Preoperative treatment naïve	WGS	Healthy	100	8591130800	5866411400	2.33
CGPLH340	Preoperative treatment naïve	WGS	Healthy	100	8046351000	5368062000	2.13
CGPLH341	Preoperative treatment naïve	WGS	Healthy	100	7914788600	5200304800	2.06

Patient	Timepoint	Analysis type	Patient Type	Read Length	Total Bases Sequenced	High Quality Bases Analyzed	Coverage
CGPLH342	Preoperative treatment naïve	WGS	Healthy	100	8633473000	5701972400	2.26
CGPLH343	Preoperative treatment naïve	WGS	Healthy	100	6694769800	4410670800	1.75
CGPLH344	Preoperative treatment naïve	WGS	Healthy	100	7628192400	4961476600	1.97
CGPLH345	Preoperative treatment naïve	WGS	Healthy	100	7121569400	4747223000	1.88
CGPLH346	Preoperative treatment naïve	WGS	Healthy	100	7707924600	4873321600	1.93
CGPLH35	Preoperative treatment naïve	WGS	Healthy	100	47305985200	4774186200	12.63
CGPLH350	Preoperative treatment naïve	WGS	Healthy	100	9745839800	6054055200	2.40
CGPLH351	Preoperative treatment naïve	WGS	Healthy	100	13317435800	8714465000	3.46
CGPLH352	Preoperative treatment naïve	WGS	Healthy	100	7659351600	4752309400	1.89
CGPLH353	Preoperative treatment naïve	WGS	Healthy	100	8435782400	5275098200	2.09
CGPLH354	Preoperative treatment naïve	WGS	Healthy	100	8018644000	4857577600	1.93
CGPLH355	Preoperative treatment naïve	WGS	Healthy	100	8624675800	5709726400	2.27
CGPLH356	Preoperative treatment naïve	WGS	Healthy	100	8817952800	5729595200	2.27
CGPLH357	Preoperative treatment naïve	WGS	Healthy	100	11931596200	7690004400	3.05
CGPLH358	Preoperative treatment naïve	WGS	Healthy	100	12802561200	8451274800	3.35
CGPLH36	Preoperative treatment naïve	WGS	Healthy	100	40173545600	3974810400	10.52
CGPLH360	Preoperative treatment naïve	WGS	Healthy	100	7280078400	4918566200	1.95
CGPLH361	Preoperative treatment naïve	WGS	Healthy	100	7493498400	4966813800	1.97
CGPLH362	Preoperative treatment naïve	WGS	Healthy	100	11345644200	7532133600	2.99
CGPLH363	Preoperative treatment naïve	WGS	Healthy	100	6117382800	3965952400	1.57
CGPLH364	Preoperative treatment naïve	WGS	Healthy	100	10823498400	7195557000	2.86
CGPLH365	Preoperative treatment naïve	WGS	Healthy	100	5938367400	3954556200	1.57
CGPLH366	Preoperative treatment naïve	WGS	Healthy	100	7063168600	4731853000	1.88
CGPLH367	Preoperative treatment naïve	WGS	Healthy	100	7119631800	4627882200	1.84
CGPLH368	Preoperative treatment naïve	WGS	Healthy	100	7726718400	4975233400	1.97
CGPLH369	Preoperative treatment naïve	WGS	Healthy	100	10967584200	7130956800	2.83
CGPLH37	Preoperative treatment naïve	WGS	Healthy	100	45570545400	4591328800	12.15
CGPLH370	Preoperative treatment naïve	WGS	Healthy	100	9237170600	6106373800	2.42
CGPLH371	Preoperative treatment naïve	WGS	Healthy	100	8077798800	5237070600	2.08
CGPLH380	Preoperative treatment naïve	WGS	Healthy	100	14049589200	8614241200	3.42
CGPLH381	Preoperative treatment naïve	WGS	Healthy	100	16743792000	10767882800	4.27
CGPLH382	Preoperative treatment naïve	WGS	Healthy	100	18474025200	12276437200	4.87
CGPLH383	Preoperative treatment naïve	WGS	Healthy	100	13215954000	8430420600	3.35
CGPLH384	Preoperative treatment naïve	WGS	Healthy	100	8481814000	546336200	2.17
CGPLH385	Preoperative treatment naïve	WGS	Healthy	100	9596118800	6445445600	2.56
CGPLH386	Preoperative treatment naïve	WGS	Healthy	100	7399540400	4915484800	1.95
CGPLH387	Preoperative treatment naïve	WGS	Healthy	100	6860332600	4339724400	1.72
CGPLH388	Preoperative treatment naïve	WGS	Healthy	100	8679705600	5463945400	2.17
CGPLH389	Preoperative treatment naïve	WGS	Healthy	100	7266863600	4702386000	1.87
CGPLH390	Preoperative treatment naïve	WGS	Healthy	100	7509035600	4913901800	1.95
CGPLH391	Preoperative treatment naïve	WGS	Healthy	100	7252286000	4702404800	1.87
CGPLH392	Preoperative treatment naïve	WGS	Healthy	100	7302618200	4722407000	1.87
CGPLH393	Preoperative treatment naïve	WGS	Healthy	100	8879138000	5947871800	2.36
CGPLH394	Preoperative treatment naïve	WGS	Healthy	100	8737031000	5597774400	2.22
CGPLH395	Preoperative treatment naïve	WGS	Healthy	100	7783904800	4907146000	1.95
CGPLH396	Preoperative treatment naïve	WGS	Healthy	100	7585567200	5076638200	2.01
CGPLH398	Preoperative treatment naïve	WGS	Healthy	100	13001418200	8607025000	3.42
CGPLH399	Preoperative treatment naïve	WGS	Healthy	100	9867699200	5526546000	2.19
CGPLH400	Preoperative treatment naïve	WGS	Healthy	100	10573939000	6290438200	2.50
CGPLH401	Preoperative treatment naïve	WGS	Healthy	100	9415150000	6139638000	2.44
CGPLH402	Preoperative treatment naïve	WGS	Healthy	100	5541458000	2972927800	1.18
CGPLH403	Preoperative treatment naïve	WGS	Healthy	100	6470913200	3549772600	1.41
CGPLH404	Preoperative treatment naïve	WGS	Healthy	100	7369651800	4120205000	1.64
CGPLH405	Preoperative treatment naïve	WGS	Healthy	100	7360239000	4293522600	1.70
CGPLH406	Preoperative treatment naïve	WGS	Healthy	100	6026125400	3426007400	1.36
CGPLH407	Preoperative treatment naïve	WGS	Healthy	100	7073375200	4079286800	1.62
CGPLH408	Preoperative treatment naïve	WGS	Healthy	100	8006103200	5121285600	2.03
CGPLH409	Preoperative treatment naïve	WGS	Healthy	100	7343124600	4432335600	1.76
CGPLH410	Preoperative treatment naïve	WGS	Healthy	100	7551842000	4818779600	1.91
CGPLH411	Preoperative treatment naïve	WGS	Healthy	100	6119678400	3636478400	1.44
CGPLH412	Preoperative treatment naïve	WGS	Healthy	100	7960821200	4935752200	1.96
CGPLH413	Preoperative treatment naïve	WGS	Healthy	100	7623405400	4827888400	1.92
CGPLH414	Preoperative treatment naïve	WGS	Healthy	100	7381312400	4743337200	1.88
CGPLH415	Preoperative treatment naïve	WGS	Healthy	100	7240754200	4162208800	1.65
CGPLH416	Preoperative treatment naïve	WGS	Healthy	100	7745658600	4670226000	1.85
CGPLH417	Preoperative treatment naïve	WGS	Healthy	100	7627498600	4403085600	1.75
CGPLH418	Preoperative treatment naïve	WGS	Healthy	100	9090285000	5094814000	2.02
CGPLH419	Preoperative treatment naïve	WGS	Healthy	100	7914120200	5078589800	2.02
CGPLH42	Preoperative treatment naïve	WGS	Healthy	100	39492040600	3901039400	10.32
CGPLH420	Preoperative treatment naïve	WGS	Healthy	100	7014307800	4711393600	1.87
CGPLH422	Preoperative treatment naïve	WGS	Healthy	100	9103972900	6053559800	2.40
CGPLH423	Preoperative treatment naïve	WGS	Healthy	100	10154714200	6128800200	2.43
CGPLH424	Preoperative treatment naïve	WGS	Healthy	100	11002394000	6573756000	2.61
CGPLH425	Preoperative treatment naïve	WGS	Healthy	100	14681352600	9272557000	3.68

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CGPLH426	Preoperative treatment naïve	WGS	Healthy	100	8336731000	5177430800	2.05
CGPLH427	Preoperative treatment naïve	WGS	Healthy	100	8242924400	5632991800	2.24
CGPLH428	Preoperative treatment naïve	WGS	Healthy	100	8512550400	5604756600	2.22
CGPLH429	Preoperative treatment naïve	WGS	Healthy	100	8363802900	5477121400	2.17
CGPLH43	Preoperative treatment naïve	WGS	Healthy	100	38513193400	3815698400	10.10
CGPLH430	Preoperative treatment naïve	WGS	Healthy	100	10357365400	6841611000	2.71
CGPLH431	Preoperative treatment naïve	WGS	Healthy	100	7599875800	5006909000	1.99
CGPLH432	Preoperative treatment naïve	WGS	Healthy	100	7932532400	4932304200	1.96
CGPLH434	Preoperative treatment naïve	WGS	Healthy	100	10417028600	6965998800	2.76
CGPLH435	Preoperative treatment naïve	WGS	Healthy	100	8747793800	5677115200	2.25
CGPLH436	Preoperative treatment naïve	WGS	Healthy	100	7990589400	5228737800	2.07
CGPLH437	Preoperative treatment naïve	WGS	Healthy	100	10156991200	6935537200	2.75
CGPLH438	Preoperative treatment naïve	WGS	Healthy	100	9473604000	6445456000	2.56
CGPLH439	Preoperative treatment naïve	WGS	Healthy	100	8303723400	5439877200	2.16
CGPLH440	Preoperative treatment naïve	WGS	Healthy	100	9055233800	6018631400	2.39
CGPLH441	Preoperative treatment naïve	WGS	Healthy	100	10290682000	6896415200	2.74
CGPLH442	Preoperative treatment naïve	WGS	Healthy	100	9878551600	6591249800	2.62
CGPLH443	Preoperative treatment naïve	WGS	Healthy	100	9837225800	6360740800	2.52
CGPLH444	Preoperative treatment naïve	WGS	Healthy	100	9199271400	5755941600	2.28
CGPLH445	Preoperative treatment naïve	WGS	Healthy	100	8089236400	5218259800	2.07
CGPLH446	Preoperative treatment naïve	WGS	Healthy	100	78906664200	5181606000	2.06
CGPLH447	Preoperative treatment naïve	WGS	Healthy	100	7775775000	5120239800	2.03
CGPLH448	Preoperative treatment naïve	WGS	Healthy	100	8686964800	5605079200	2.22
CGPLH449	Preoperative treatment naïve	WGS	Healthy	100	8604545400	5527726600	2.19
CGPLH45	Preoperative treatment naïve	WGS	Healthy	100	39029653000	3771601200	9.98
CGPLH450	Preoperative treatment naïve	WGS	Healthy	100	8428264800	5439950000	2.16
CGPLH451	Preoperative treatment naïve	WGS	Healthy	100	8128977600	5136265600	2.06
CGPLH452	Preoperative treatment naïve	WGS	Healthy	100	6474313400	4216316400	1.67
CGPLH453	Preoperative treatment naïve	WGS	Healthy	100	9831832800	6224917600	2.47
CGPLH455	Preoperative treatment naïve	WGS	Healthy	100	7373753000	4593473600	1.82
CGPLH456	Preoperative treatment naïve	WGS	Healthy	100	8455416200	5457148200	2.17
CGPLH457	Preoperative treatment naïve	WGS	Healthy	100	8647618000	5534503800	2.20
CGPLH458	Preoperative treatment naïve	WGS	Healthy	100	6633156400	4415186000	1.75
CGPLH459	Preoperative treatment naïve	WGS	Healthy	100	8361049200	5497193800	2.18
CGPLH46	Preoperative treatment naïve	WGS	Healthy	100	35361484600	3516232800	9.30
CGPLH460	Preoperative treatment naïve	WGS	Healthy	100	6788835400	4472282800	1.77
CGPLH463	Preoperative treatment naïve	WGS	Healthy	100	8534880800	5481759200	2.18
CGPLH464	Preoperative treatment naïve	WGS	Healthy	100	6692520000	4184463400	1.66
CGPLH465	Preoperative treatment naïve	WGS	Healthy	100	7772884600	4878430800	1.94
CGPLH466	Preoperative treatment naïve	WGS	Healthy	100	9056275000	5830877400	2.31
CGPLH467	Preoperative treatment naïve	WGS	Healthy	100	6931419200	4585861000	1.82
CGPLH468	Preoperative treatment naïve	WGS	Healthy	100	9334067400	6314830400	2.51
CGPLH469	Preoperative treatment naïve	WGS	Healthy	100	7376691000	4545246600	1.80
CGPLH47	Preoperative treatment naïve	WGS	Healthy	100	38485647600	3534983600	9.35
CGPLH470	Preoperative treatment naïve	WGS	Healthy	100	7899727600	5221650600	2.07
CGPLH471	Preoperative treatment naïve	WGS	Healthy	100	9200430600	6162371000	2.42
CGPLH472	Preoperative treatment naïve	WGS	Healthy	100	8143742400	5399946600	2.14
CGPLH473	Preoperative treatment naïve	WGS	Healthy	100	8123924600	5419825400	2.15
CGPLH474	Preoperative treatment naïve	WGS	Healthy	100	8853071400	6084059400	2.41
CGPLH475	Preoperative treatment naïve	WGS	Healthy	100	8115374000	5291718000	2.10
CGPLH476	Preoperative treatment naïve	WGS	Healthy	100	8163162600	5096369600	2.02
CGPLH477	Preoperative treatment naïve	WGS	Healthy	100	8350093200	5465468600	2.17
CGPLH478	Preoperative treatment naïve	WGS	Healthy	100	8259642200	5406516200	2.15
CGPLH479	Preoperative treatment naïve	WGS	Healthy	100	8027598600	5417376800	2.15
CGPLH48	Preoperative treatment naïve	WGS	Healthy	100	42232410000	4165893400	11.02
CGPLH480	Preoperative treatment naïve	WGS	Healthy	100	7832983200	5020127000	1.99
CGPLH481	Preoperative treatment naïve	WGS	Healthy	100	7578518800	4883280800	1.94
CGPLH482	Preoperative treatment naïve	WGS	Healthy	100	8279364800	5652263600	2.24
CGPLH483	Preoperative treatment naïve	WGS	Healthy	100	8660338800	5823859200	2.31
CGPLH484	Preoperative treatment naïve	WGS	Healthy	100	8445420000	5794328000	2.30
CGPLH485	Preoperative treatment naïve	WGS	Healthy	100	8371255400	5490207800	2.18
CGPLH486	Preoperative treatment naïve	WGS	Healthy	100	8216712200	5506871000	2.19
CGPLH487	Preoperative treatment naïve	WGS	Healthy	100	7936294200	5309250200	2.11
CGPLH488	Preoperative treatment naïve	WGS	Healthy	100	8355603600	5453160000	2.16
CGPLH49	Preoperative treatment naïve	WGS	Healthy	100	33912191800	3310056000	8.76
CGPLH490	Preoperative treatment naïve	WGS	Healthy	100	7768712400	5175567800	2.05
CGPLH491	Preoperative treatment naïve	WGS	Healthy	100	9070904000	6011275000	2.39
CGPLH492	Preoperative treatment naïve	WGS	Healthy	100	7208727200	4753213800	1.89
CGPLH493	Preoperative treatment naïve	WGS	Healthy	100	10542882600	7225870800	2.87
CGPLH494	Preoperative treatment naïve	WGS	Healthy	100	10908197600	7046645000	2.80
CGPLH495	Preoperative treatment naïve	WGS	Healthy	100	8945040400	5891597800	2.34
CGPLH496	Preoperative treatment naïve	WGS	Healthy	100	10859729400	7549608000	3.00
CGPLH497	Preoperative treatment naïve	WGS	Healthy	100	9630507400	6473162800	2.57
CGPLH498	Preoperative treatment naïve	WGS	Healthy	100	10060232600	6744622800	2.68

Patient	Timepoint	Analysis type	Patient Type	Read Length	Total Bases Sequenced	High Quality Bases Analyzed	Coverage
CGPLH499	Preoperative treatment naïve	WGS	Healthy	100	10221293600	6951282800	2.76
CGPLH50	Preoperative treatment naïve	WGS	Healthy	100	41248860600	4073272800	10.78
CGPLH500	Preoperative treatment naïve	WGS	Healthy	100	9703168200	6239893800	2.48
CGPLH501	Preoperative treatment naïve	WGS	Healthy	100	9104779800	6161502800	2.45
CGPLH502	Preoperative treatment naïve	WGS	Healthy	100	8514467400	5290881400	2.10
CGPLH503	Preoperative treatment naïve	WGS	Healthy	100	9019992200	6100383400	2.42
CGPLH504	Preoperative treatment naïve	WGS	Healthy	100	9320330200	6199750200	2.46
CGPLH505	Preoperative treatment naïve	WGS	Healthy	100	7495497400	4914559000	1.95
CGPLH506	Preoperative treatment naïve	WGS	Healthy	100	10526142000	6963312600	2.76
CGPLH507	Preoperative treatment naïve	WGS	Healthy	100	9091018400	6146678600	2.44
CGPLH508	Preoperative treatment naïve	WGS	Healthy	100	10889315600	7360201400	2.92
CGPLH509	Preoperative treatment naïve	WGS	Healthy	100	9729084600	6702691600	2.66
CGPLH51	Preoperative treatment naïve	WGS	Healthy	100	35967451400	349233200	9.24
CGPLH510	Preoperative treatment naïve	WGS	Healthy	100	11162691600	7626795400	3.03
CGPLH511	Preoperative treatment naïve	WGS	Healthy	100	11888619600	8110427600	3.22
CGPLH512	Preoperative treatment naïve	WGS	Healthy	100	10726438400	7110078000	2.82
CGPLH513	Preoperative treatment naïve	WGS	Healthy	100	10701564200	7155271400	2.84
CGPLH514	Preoperative treatment naïve	WGS	Healthy	100	8822067000	5958773800	2.36
CGPLH515	Preoperative treatment naïve	WGS	Healthy	100	7792074800	5317464600	2.11
CGPLH516	Preoperative treatment naïve	WGS	Healthy	100	8642620000	5846439400	2.32
CGPLH517	Preoperative treatment naïve	WGS	Healthy	100	11915929600	8013937000	3.18
CGPLH518	Preoperative treatment naïve	WGS	Healthy	100	12804517400	8606661600	3.42
CGPLH519	Preoperative treatment naïve	WGS	Healthy	100	11513222200	7922798400	3.14
CGPLH52	Preoperative treatment naïve	WGS	Healthy	100	49247304200	4849531400	12.83
CGPLH520	Preoperative treatment naïve	WGS	Healthy	100	8942102400	6030683400	2.39
CGPLH54	Preoperative treatment naïve	WGS	Healthy	100	45399346400	4466164600	11.82
CGPLH55	Preoperative treatment naïve	WGS	Healthy	100	42547725000	423337600	11.33
CGPLH56	Preoperative treatment naïve	WGS	Healthy	100	33460308000	3226338000	8.53
CGPLH57	Preoperative treatment naïve	WGS	Healthy	100	36504735200	3509125000	9.28
CGPLH59	Preoperative treatment naïve	WGS	Healthy	100	39642810600	3820011000	10.11
CGPLH625	Preoperative treatment naïve	WGS	Healthy	100	6408225000	4115487600	1.63
CGPLH626	Preoperative treatment naïve	WGS	Healthy	100	9915193600	6391857000	2.54
CGPLH63	Preoperative treatment naïve	WGS	Healthy	100	37447047600	3506737000	9.28
CGPLH639	Preoperative treatment naïve	WGS	Healthy	100	8158965800	5216049600	2.07
CGPLH64	Preoperative treatment naïve	WGS	Healthy	100	34275506800	3264508000	8.63
CGPLH640	Preoperative treatment naïve	WGS	Healthy	100	8058876800	5333551800	2.12
CGPLH642	Preoperative treatment naïve	WGS	Healthy	100	7545555600	4909732800	1.95
CGPLH643	Preoperative treatment naïve	WGS	Healthy	100	7865776800	5254772000	2.09
CGPLH644	Preoperative treatment naïve	WGS	Healthy	100	6890139000	4599387400	1.83
CGPLH646	Preoperative treatment naïve	WGS	Healthy	100	7757219400	5077408200	2.01
CGPLH75	Preoperative treatment naïve	WGS	Healthy	100	23882926000	2250344400	5.95
CGPLH76	Preoperative treatment naïve	WGS	Healthy	100	30631483600	3086042200	8.16
CGPLH77	Preoperative treatment naïve	WGS	Healthy	100	31651741400	3041290200	8.04
CGPLH78	Preoperative treatment naïve	WGS	Healthy	100	31165831200	3130079800	8.28
CGPLH79	Preoperative treatment naïve	WGS	Healthy	100	31935043000	3128488200	8.27
CGPLH80	Preoperative treatment naïve	WGS	Healthy	100	32965093000	3311371800	8.76
CGPLH81	Preoperative treatment naïve	WGS	Healthy	100	27035311200	2455084400	6.49
CGPLH82	Preoperative treatment naïve	WGS	Healthy	100	28447051200	2893368200	7.65
CGPLH83	Preoperative treatment naïve	WGS	Healthy	100	26702240200	2459494000	6.50
CGPLH84	Preoperative treatment naïve	WGS	Healthy	100	25176861400	2524467400	6.68
CGPLLU13	Pre-treatment, Day -2	WGS	Lung Cancer	100	9126585600	5915061800	2.35
CGPLLU13	Post-treatment, Day 5	WGS	Lung Cancer	100	7739120200	5071745800	2.01
CGPLLU13	Post-treatment, Day 28	WGS	Lung Cancer	100	9081585400	5764371600	2.29
CGPLLU13	Post-treatment, Day 91	WGS	Lung Cancer	100	9576557000	6160760200	2.44
CGPLLU14	Pre-treatment, Day -38	WGS	Lung Cancer	100	13659198400	9033455800	3.58
CGPLLU14	Pre-treatment, Day -16	WGS	Lung Cancer	100	7178855800	4856648600	1.93
CGPLLU14	Pre-treatment, Day -3	WGS	Lung Cancer	100	7653473000	4816193600	1.91
CGPLLU14	Pre-treatment, Day 0	WGS	Lung Cancer	100	785197400	5193256600	2.06
CGPLLU14	Post-treatment, Day 0.33	WGS	Lung Cancer	100	7193040800	4869701600	1.93
CGPLLU14	Post-treatment, Day 7	WGS	Lung Cancer	100	710205000	4741432600	1.88
CGPLLU144	Preoperative treatment naïve	WGS	Lung Cancer	100	4934813600	3415936400	1.36
CGPLLU147	Preoperative treatment naïve	WGS	Lung Cancer	100	24409561000	2118672800	5.61
CGPLLU161	Preoperative treatment naïve	WGS	Lung Cancer	100	8998813400	6016145000	2.39
CGPLLU162	Preoperative treatment naïve	WGS	Lung Cancer	100	9703792400	6407866400	2.54
CGPLLU163	Preoperative treatment naïve	WGS	Lung Cancer	100	9150620200	5083569800	2.41
CGPLLU165	Preoperative treatment naïve	WGS	Lung Cancer	100	28374436400	2651138600	7.01
CGPLLU168	Preoperative treatment naïve	WGS	Lung Cancer	100	5692739400	3695191000	1.47
CGPLLU169	Preoperative treatment naïve	WGS	Lung Cancer	100	9093975600	5805320800	2.30
CGPLLU175	Preoperative treatment naïve	WGS	Lung Cancer	100	33794816800	3418750400	9.04
CGPLLU176	Preoperative treatment naïve	WGS	Lung Cancer	100	8778553800	5794950200	2.30
CGPLLU177	Preoperative treatment naïve	WGS	Lung Cancer	100	3734614900	2578596200	1.02
CGPLLU180	Preoperative treatment naïve	WGS	Lung Cancer	100	28305936600	2756034200	7.29
CGPLLU198	Preoperative treatment naïve	WGS	Lung Cancer	100	23244959200	2218577200	5.86
CGPLLU202	Preoperative treatment naïve	WGS	Lung Cancer	100	21110128200	1831279400	4.84

Patient	Timepoint	Analysis type	Patient Type	Read Length	Total Bases Sequenced	High Quality Bases Analyzed	Coverage
CGPLLU203	Preoperative treatment naïve	WGS	Lung Cancer	100	4304235600	28936429000	1.15
CGPLLU205	Preoperative treatment naïve	WGS	Lung Cancer	100	10502467000	7386984800	2.93
CGPLLU206	Preoperative treatment naïve	WGS	Lung Cancer	100	21888248200	2026666000	5.36
CGPLLU207	Preoperative treatment naïve	WGS	Lung Cancer	100	10806230600	7363649000	2.92
CGPLLU208	Preoperative treatment naïve	WGS	Lung Cancer	100	7795426800	5199545800	2.06
CGPLLU209	Preoperative treatment naïve	WGS	Lung Cancer	100	26174542000	2621961800	6.93
CGPLLU244	Pre-treatment, Day -7	WGS	Lung Cancer	100	9967531400	6704365800	2.66
CGPLLU244	Pre-treatment, Day -1	WGS	Lung Cancer	100	9547119200	5785172600	2.30
CGPLLU244	Post-treatment, Day 6	WGS	Lung Cancer	100	9535898600	6452174000	2.56
CGPLLU244	Post-treatment, Day 62	WGS	Lung Cancer	100	8783628600	5914149000	2.35
CGPLLU245	Pre-treatment, Day -32	WGS	Lung Cancer	100	10025823200	6313303800	2.51
CGPLLU245	Pre-treatment, Day 0	WGS	Lung Cancer	100	9462480400	6612867800	2.62
CGPLLU245	Post-treatment, Day 7	WGS	Lung Cancer	100	9143925000	6431013200	2.55
CGPLLU245	Post-treatment, Day 21	WGS	Lung Cancer	100	9072713800	6368533000	2.53
CGPLLU246	Pre-treatment, Day -21	WGS	Lung Cancer	100	9579787000	6458003400	2.56
CGPLLU246	Pre-treatment, Day 0	WGS	Lung Cancer	100	9512703600	6440535600	2.56
CGPLLU246	Post-treatment, Day 9	WGS	Lung Cancer	100	9512646000	6300939200	2.50
CGPLLU246	Post-treatment, Day 42	WGS	Lung Cancer	100	11136103000	7358747400	2.92
CGPLLU264	Pre-treatment, Day -1	WGS	Lung Cancer	100	9196305000	6239803600	2.48
CGPLLU264	Post-treatment, Day 6	WGS	Lung Cancer	100	8247416600	5600454200	2.22
CGPLLU264	Post-treatment, Day 27	WGS	Lung Cancer	100	8681022200	5856109000	2.32
CGPLLU264	Post-treatment, Day 69	WGS	Lung Cancer	100	8931976400	5974246000	2.37
CGPLLU265	Pre-treatment, Day 0	WGS	Lung Cancer	100	9460534000	6111185200	2.43
CGPLLU265	Post-treatment, Day 3	WGS	Lung Cancer	100	8051601200	4984166600	1.98
CGPLLU265	Post-treatment, Day 7	WGS	Lung Cancer	100	8082224600	5110092600	2.03
CGPLLU265	Post-treatment, Day 84	WGS	Lung Cancer	100	8368637400	5369526400	2.13
CGPLLU266	Pre-treatment, Day 0	WGS	Lung Cancer	100	8583766400	5846473600	2.32
CGPLLU266	Post-treatment, Day 16	WGS	Lung Cancer	100	8795793600	5984531400	2.37
CGPLLU266	Post-treatment, Day 83	WGS	Lung Cancer	100	9157947600	6227735000	2.47
CGPLLU266	Post-treatment, Day 328	WGS	Lung Cancer	100	7299455400	5049379000	2.00
CGPLLU267	Pre-treatment, Day -1	WGS	Lung Cancer	100	10658657800	6892667000	2.73
CGPLLU267	Post-treatment, Day 34	WGS	Lung Cancer	100	8492833400	5101097800	2.02
CGPLLU267	Post-treatment, Day 90	WGS	Lung Cancer	100	12030314800	7757930400	3.08
CGPLLU269	Pre-treatment, Day 0	WGS	Lung Cancer	100	9170168000	5830454400	2.31
CGPLLU269	Post-treatment, Day 9	WGS	Lung Cancer	100	8905640400	5298461400	2.10
CGPLLU269	Post-treatment, Day 28	WGS	Lung Cancer	100	8455306600	5387927400	2.14
CGPLLU271	Post-treatment, Day 259	WGS	Lung Cancer	100	8112060400	5404379000	2.14
CGPLLU271	Pre-treatment, Day 0	WGS	Lung Cancer	100	13150818200	8570453400	3.40
CGPLLU271	Post-treatment, Day 6	WGS	Lung Cancer	100	9008880600	5854051400	2.32
CGPLLU271	Post-treatment, Day 20	WGS	Lung Cancer	100	8670913000	5461577000	2.17
CGPLLU271	Post-treatment, Day 104	WGS	Lung Cancer	100	8867441400	5609039000	2.23
CGPLLU43	Pre-treatment, Day -1	WGS	Lung Cancer	100	8407811200	5203486400	2.06
CGPLLU43	Post-treatment, Day 6	WGS	Lung Cancer	100	9264335200	5626714400	2.23
CGPLLU43	Post-treatment, Day 27	WGS	Lung Cancer	100	8902283000	5485556200	2.18
CGPLLU43	Post-treatment, Day 83	WGS	Lung Cancer	100	9201509200	5875084200	2.33
CGPLLU86	Pre-treatment, Day 0	WGS	Lung Cancer	100	9152729200	6248173200	2.48
CGPLLU86	Post-treatment, Day 0.5	WGS	Lung Cancer	100	6703253000	4663026800	1.85
CGPLLU86	Post-treatment, Day 7	WGS	Lung Cancer	100	6590121400	4559562400	1.81
CGPLLU86	Post-treatment, Day 17	WGS	Lung Cancer	100	8653551800	5900136000	2.34
CGPLLU88	Pre-treatment, Day 0	WGS	Lung Cancer	100	8096528000	5505475400	2.18
CGPLLU88	Post-treatment, Day 7	WGS	Lung Cancer	100	8283192200	5784217600	2.30
CGPLLU88	Post-treatment, Day 297	WGS	Lung Cancer	100	9297110800	6407258000	2.54
CGPLLU89	Pre-treatment, Day 0	WGS	Lung Cancer	100	7842145200	5356095400	2.13
CGPLLU89	Post-treatment, Day 7	WGS	Lung Cancer	100	7234220200	4930375200	1.96
CGPLLU89	Post-treatment, Day 22	WGS	Lung Cancer	100	6242889800	4057361000	1.61
CGPLOV11	Preoperative treatment naïve	WGS	Ovarian Cancer	100	8985130400	5871959600	2.33
CGPLOV12	Preoperative treatment naïve	WGS	Ovarian Cancer	100	9705820000	6430505400	2.55
CGPLOV13	Preoperative treatment naïve	WGS	Ovarian Cancer	100	10307949400	7029712000	2.79
CGPLOV15	Preoperative treatment naïve	WGS	Ovarian Cancer	100	8472829400	5562142400	2.21
CGPLOV16	Preoperative treatment naïve	WGS	Ovarian Cancer	100	10977781000	7538581600	2.99
CGPLOV19	Preoperative treatment naïve	WGS	Ovarian Cancer	100	8800876200	5855304000	2.32
CGPLOV20	Preoperative treatment naïve	WGS	Ovarian Cancer	100	8714443600	5695165800	2.26
CGPLOV21	Preoperative treatment naïve	WGS	Ovarian Cancer	100	10180394800	7120260400	2.83
CGPLOV22	Preoperative treatment naïve	WGS	Ovarian Cancer	100	10107760000	6821916800	2.71
CGPLOV23	Preoperative treatment naïve	WGS	Ovarian Cancer	100	10643398000	7206330800	2.86
CGPLOV24	Preoperative treatment naïve	WGS	Ovarian Cancer	100	6780929000	4623300400	1.83
CGPLOV25	Preoperative treatment naïve	WGS	Ovarian Cancer	100	7817548600	5359975200	2.13
CGPLOV26	Preoperative treatment naïve	WGS	Ovarian Cancer	100	11763101400	8178024400	3.25
CGPLOV28	Preoperative treatment naïve	WGS	Ovarian Cancer	100	9522546400	6259423400	2.48
CGPLOV31	Preoperative treatment naïve	WGS	Ovarian Cancer	100	9104831200	6109358400	2.42
CGPLOV32	Preoperative treatment naïve	WGS	Ovarian Cancer	100	9222073600	6035150000	2.39
CGPLOV37	Preoperative treatment naïve	WGS	Ovarian Cancer	100	8898328600	5971018200	2.37
CGPLOV38	Preoperative treatment naïve	WGS	Ovarian Cancer	100	8756825200	5861536600	2.33
CGPLOV40	Preoperative treatment naïve	WGS	Ovarian Cancer	100	9709391600	6654707200	2.64

Patient	Timepoint	Analysis type	Patient Type	Read Length	Total Bases Sequenced	High Quality Bases Analyzed	Coverage
CGPLOV41	Preoperative treatment naïve	WGS	Ovarian Cancer	100	8923625000	5973070400	2.37
CGPLOV42	Preoperative treatment naïve	WGS	Ovarian Cancer	100	10719380400	7353214200	2.92
CGPLOV43	Preoperative treatment naïve	WGS	Ovarian Cancer	100	10272189000	6423288600	2.55
CGPLOV44	Preoperative treatment naïve	WGS	Ovarian Cancer	100	98611862600	6789185800	2.69
CGPLOV46	Preoperative treatment naïve	WGS	Ovarian Cancer	100	8788956400	5789863400	2.30
CGPLOV47	Preoperative treatment naïve	WGS	Ovarian Cancer	100	9380561800	6480763600	2.57
CGPLOV48	Preoperative treatment naïve	WGS	Ovarian Cancer	100	9258552600	6380106400	2.53
CGPLOV49	Preoperative treatment naïve	WGS	Ovarian Cancer	100	8787025400	6134503600	2.43
CGPLOV50	Preoperative treatment naïve	WGS	Ovarian Cancer	100	10144154400	6984721400	2.77
CGPLPA112	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	12740651400	9045622000	3.59
CGPLPA113	Preoperative treatment naïve	WGS	Duodenal Cancer	100	8802479000	5909308000	2.34
CGPLPA114	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	8792313600	6019061000	2.39
CGPLPA115	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	8636551400	5958809000	2.36
CGPLPA117	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	9128885200	6288833200	2.50
CGPLPA118	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	7931485800	5407532800	2.15
CGPLPA122	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	10886985000	7530118800	2.99
CGPLPA124	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	8562012400	5860171000	2.33
CGPLPA125	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	9715576600	6390321000	2.54
CGPLPA126	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	8056768800	5651600800	2.24
CGPLPA127	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	8000301000	5382987600	2.14
CGPLPA128	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	6165751600	4256521400	1.69
CGPLPA129	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	7143147400	4917370400	1.95
CGPLPA130	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	5664335000	3603919400	1.43
CGPLPA131	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	8292962000	5844942000	2.32
CGPLPA134	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	7088917000	5048887600	2.00
CGPLPA135	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	8759665600	5800618200	2.30
CGPLPA136	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	7535715800	5248227600	2.08
CGPLPA137	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	8391815400	5901273800	2.34
CGPLPA139	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	8992280200	6328314400	2.51
CGPLPA14	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	8787706200	5731317600	2.27
CGPLPA140	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	16365641800	11216732000	4.45
CGPLPA141	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	15086298000	10114790200	4.01
CGPLPA15	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	8255566800	5531577600	2.20
CGPLPA155	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	9457155800	6621881800	2.63
CGPLPA156	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	9845385800	6728653000	2.67
CGPLPA165	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	8356604600	5829895800	2.31
CGPLPA168	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	10365661600	7048115600	2.80
CGPLPA17	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	8073547400	4667808000	1.86
CGPLPA184	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	9014218400	6230922200	2.47
CGPLPA187	Preoperative treatment naïve	WGS	Bile Duct Cancer	100	8883536200	6140874400	2.44
CGPLPA23	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	9353452000	6246525400	2.48
CGPLPA25	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	10077515400	6103322200	2.42
CGPLPA26	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	8354272400	5725781000	2.27
CGPLPA28	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	8477461600	5688846800	2.26
CGPLPA33	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	7287615600	4596723800	1.82
CGPLPA34	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	6122902400	4094828000	1.62
CGPLPA37	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	12714886200	8527779200	3.38
CGPLPA38	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	8525500600	5501341400	2.18
CGPLPA39	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	10502663600	6812333000	2.70
CGPLPA40	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	9083670000	5394717800	2.14
CGPLPA42	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	5972126000	3890395200	1.54
CGPLPA46	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	4720090200	2626298800	1.04
CGPLPA47	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	7317385800	4543833000	1.80
CGPLPA48	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	7553856200	5022695600	1.99
CGPLPA52	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	5655875000	3551861600	1.41
CGPLPA53	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	9504749000	6323344800	2.51
CGPLPA58	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	8088090200	5118138200	2.03
CGPLPA59	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	14547364600	9617778600	3.82
CGPLPA67	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	8222177400	5351172600	2.12
CGPLPA69	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	7899181400	5006114800	1.99
CGPLPA71	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	7349620400	4955417400	1.97
CGPLPA74	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	6666371400	4571394200	1.81
CGPLPA76	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	9755658600	6412606800	2.54
CGPLPA85	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	10856223000	7339498600	2.90
CGPLPA86	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	8744365400	5514523200	2.19
CGPLPA92	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	8073791200	5390492800	2.14
CGPLPA93	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	10390273000	7136589400	2.85
CGPLPA94	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	11060347600	7641336400	3.03
CGPLPA95	Preoperative treatment naïve	WGS	Pancreatic Cancer	100	12416627200	7206503800	2.86
CGST102	Preoperative treatment naïve	WGS	Gastric cancer	100	6637004600	4545072800	1.80
CGST11	Preoperative treatment naïve	WGS	Gastric cancer	100	9718427800	6259579800	2.48
CGST110	Preoperative treatment naïve	WGS	Gastric cancer	100	9319661600	6359317400	2.52
CGST114	Preoperative treatment naïve	WGS	Gastric cancer	100	6865213000	4841171600	1.92
CGST13	Preoperative treatment naïve	WGS	Gastric cancer	100	928454800	6360843800	2.52

Patient	Timepoint	Analysis type	Patient Type	Read Length	Total Bases Sequenced	High Quality Bases Analyzed	Coverage
CGST131	Preoperative treatment naïve	WGS	Gastric cancer	100	5924382000	386057200	1.53
CGST141	Preoperative treatment naïve	WGS	Gastric cancer	100	8486380800	5860491000	2.33
CGST16	Preoperative treatment naïve	WGS	Gastric cancer	100	13820725800	9377828000	3.72
CGST18	Preoperative treatment naïve	WGS	Gastric cancer	100	7781288000	5278862400	2.09
CGST21	Preoperative treatment naïve	WGS	Gastric cancer	100	7171165400	4103970800	1.63
CGST26	Preoperative treatment naïve	WGS	Gastric cancer	100	8983961800	6053405600	2.40
CGST28	Preoperative treatment naïve	WGS	Gastric cancer	100	9683035400	6745116400	2.68
CGST30	Preoperative treatment naïve	WGS	Gastric cancer	100	8684086600	5741416000	2.28
CGST32	Preoperative treatment naïve	WGS	Gastric cancer	100	8568194600	5783369200	2.29
CGST33	Preoperative treatment naïve	WGS	Gastric cancer	100	9351699600	6448718400	2.56
CGST38	Preoperative treatment naïve	WGS	Gastric cancer	100	8403876400	5770589200	2.29
CGST39	Preoperative treatment naïve	WGS	Gastric cancer	100	10573763000	7597016000	3.01
CGST41	Preoperative treatment naïve	WGS	Gastric cancer	100	9434854200	6609415400	2.62
CGST45	Preoperative treatment naïve	WGS	Gastric cancer	100	8203858600	5625223000	2.23
CGST47	Preoperative treatment naïve	WGS	Gastric cancer	100	8938597600	6178990600	2.45
CGST48	Preoperative treatment naïve	WGS	Gastric cancer	100	9106628800	6517085200	2.59
CGST53	Preoperative treatment naïve	WGS	Gastric cancer	100	9005374200	5854996200	2.32
CGST58	Preoperative treatment naïve	WGS	Gastric cancer	100	10020368600	6133458400	2.43
CGST67	Preoperative treatment naïve	WGS	Gastric cancer	100	9198135600	5911071000	2.35
CGST77	Preoperative treatment naïve	WGS	Gastric cancer	100	8228789400	5119116800	2.03
CGST80	Preoperative treatment naïve	WGS	Gastric cancer	100	10596963400	7283152800	2.89
CGST81	Preoperative treatment naïve	WGS	Gastric cancer	100	8494881200	5838064000	2.32

APPENDIX E: Table 5. High coverage whole genome cfDNA analyses of healthy individuals and lung cancer patients

Patient	Patient Type	Analysis Type	Timepoint	Stage at Diagnosis	Median cfDNA Fragment Size (bp)	Correlation of Fragment Ratio Profile to Median Fragment Ratio Profile of Healthy Individuals		Correlation of GC Corrected Fragment Ratio Profile to Median Fragment Ratio Profile of Lymphocytes	
						Correlation of Fragment Ratio Profile to Median Fragment Ratio Profile of Healthy Individuals	Correlation of Fragment Ratio Profile to Median Fragment Ratio Profile of Lymphocytes	Correlation of GC Corrected Fragment Ratio Profile to Median Fragment Ratio Profile of Healthy Individuals	Correlation of Fragment Ratio Profile to Median Fragment Ratio Profile of Lymphocytes
CGPL-H75	Healthy	WGS	Preoperative treatment naïve	NA	168	0.977	0.952	0.920	-0.836
CGPL-H77	Healthy	WGS	Preoperative treatment naïve	NA	166	0.970	0.960	0.904	-0.912
CGPL-H80	Healthy	WGS	Preoperative treatment naïve	NA	168	0.955	0.949	0.960	-0.917
CGPL-H81	Healthy	WGS	Preoperative treatment naïve	NA	167	0.949	0.953	0.969	-0.833
CGPL-H82	Healthy	WGS	Preoperative treatment naïve	NA	166	0.969	0.949	0.954	-0.917
CGPL-H83	Healthy	WGS	Preoperative treatment naïve	NA	167	0.949	0.939	0.919	-0.904
CGPL-H84	Healthy	WGS	Preoperative treatment naïve	NA	168	0.967	0.948	0.951	-0.913
CGPL-H82	Healthy	WGS	Preoperative treatment naïve	NA	167	0.946	0.968	0.958	-0.924
CGPL-H85	Healthy	WGS	Preoperative treatment naïve	NA	166	0.981	0.973	0.945	-0.921
CGPL-H37	Healthy	WGS	Preoperative treatment naïve	NA	168	0.968	0.970	0.951	-0.922
CGPL-H64	Healthy	WGS	Preoperative treatment naïve	NA	167	0.968	0.976	0.948	-0.925
CGPL-H85	Healthy	WGS	Preoperative treatment naïve	NA	166	0.947	0.964	0.948	-0.917
CGPL-H48	Healthy	WGS	Preoperative treatment naïve	NA	168	0.959	0.965	0.960	-0.923
CGPL-H93	Healthy	WGS	Preoperative treatment naïve	NA	167	0.960	0.968	0.952	-0.921
CGPL-H38	Healthy	WGS	Preoperative treatment naïve	NA	168	0.955	0.954	0.935	-0.919
CGPL-H42	Healthy	WGS	Preoperative treatment naïve	NA	167	0.973	0.953	0.948	-0.918
CGPL-H43	Healthy	WGS	Preoperative treatment naïve	NA	166	0.952	0.958	0.953	-0.928
CGPL-H58	Healthy	WGS	Preoperative treatment naïve	NA	168	0.970	0.965	0.951	-0.925
CGPL-H45	Healthy	WGS	Preoperative treatment naïve	NA	168	0.965	0.950	0.949	-0.911
CGPL-H47	Healthy	WGS	Preoperative treatment naïve	NA	167	0.952	0.944	0.954	-0.924
CGPL-H46	Healthy	WGS	Preoperative treatment naïve	NA	168	0.966	0.965	0.953	-0.923
CGPL-H63	Healthy	WGS	Preoperative treatment naïve	NA	168	0.977	0.968	0.939	-0.920
CGPL-H51	Healthy	WGS	Preoperative treatment naïve	NA	168	0.935	0.955	0.957	-0.914
CGPL-H57	Healthy	WGS	Preoperative treatment naïve	NA	169	0.965	0.950	0.955	-0.917
CGPL-H46	Healthy	WGS	Preoperative treatment naïve	NA	168	0.958	0.951	0.950	-0.924
CGPL-H66	Healthy	WGS	Preoperative treatment naïve	NA	166	0.940	0.957	0.939	-0.911
CGPL-H64	Healthy	WGS	Preoperative treatment naïve	NA	169	0.960	0.940	0.949	-0.918
CGPL-H78	Healthy	WGS	Preoperative treatment naïve	NA	166	0.956	0.936	0.938	-0.911
CGPL-H79	Healthy	WGS	Preoperative treatment naïve	NA	168	0.960	0.957	0.953	-0.917
CGPL-H76	Healthy	WGS	Preoperative treatment naïve	NA	167	0.969	0.965	0.953	-0.917
CGPLU175	Lung Cancer	WGS	Preoperative treatment naïve	I	165	0.316	0.284	0.244	-0.282
CGPLU180	Lung Cancer	WGS	Preoperative treatment naïve	I	166	0.907	0.846	0.826	-0.819
CGPLU198	Lung Cancer	WGS	Preoperative treatment naïve	I	166	0.972	0.946	0.926	-0.911
CGPLU232	Lung Cancer	WGS	Preoperative treatment naïve	I	163	0.821	0.605	0.905	-0.843
CGPLU165	Lung Cancer	WGS	Preoperative treatment naïve	I	163	0.924	0.961	0.815	-0.851
CGPLU239	Lung Cancer	WGS	Preoperative treatment naïve	I	163	0.578	0.526	0.513	-0.534
CGPLU147	Lung Cancer	WGS	Preoperative treatment naïve	II	166	0.963	0.919	0.939	-0.912
CGPLU236	Lung Cancer	WGS	Preoperative treatment naïve	II	158	0.488	0.343	0.460	-0.481

APPENDIX F: Table 6. Monitoring response to therapy using whole genome analyses of cfDNA fragmentation profiles and targeted mutations analyses

Patient	Patient Type	Analysis Type	Timepoint	Stage	Progression-free Survival (months)	Correlation of Fragment Ratio Profile to Healthy Individuals	Correlation of Fragment Ratio Profile to Lymphocyte Nucleosome Distances	Targeted Mutation	Maximum Mutant Allele Fraction
CGLU114	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 38	IV	15.4	0.941	0.841	EGFR 851L>Q	0.85%
CGLU114	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day -16	IV	15.4	0.953	-0.653	EGFR 851L>Q	0.18%
CGLU114	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 3	IV	15.4	0.908	-0.814	EGFR 719S>S	0.49%
CGLU114	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 0	IV	15.4	0.983	-0.752	EGFR 851L>Q	1.39%
CGLU114	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 0.33	IV	15.4	0.820	-0.692	EGFR 719S>S	1.05%
CGLU114	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 7	IV	15.4	0.927	-0.887	EGFR 851L>Q	0.09%
CGLU114	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 0	IV	15.4	0.657	-0.584	EGFR 745KLE>A>T	9.06%
CGLU114	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 7	IV	18.0	0.938	-0.759	EGFR 851L>Q	0.15%
CGLU114	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 237	IV	18.0	0.946	-0.869	EGFR 745KLE>A>T	0.93%
CGLU1244	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 7	IV	1.2	0.850	-0.706	EGFR 851L>R	4.98%
CGLU1244	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day -1	IV	1.2	0.867	-0.764	EGFR 621-R	3.41%
CGLU1244	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 6	IV	1.2	0.703	-0.639	EGFR 851L>R	5.57%
CGLU1244	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 21	IV	1.2	0.655	-0.680	EGFR 851L>R	11.65%
CGLU1244	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 21	IV	1.2	0.655	-0.680	EGFR 745KLE>A>K	10.40%
CGLU1245	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day -32	IV	1.7	0.871	-0.724	EGFR 745KLE>A>K	14.10%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 0	IV	1.7	0.738	-0.608	EGFR 745KLE>A>K	0.93%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 7	IV	1.7	0.731	-0.559	EGFR 745KLE>A>K	8.56%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 21	IV	1.7	0.613	-0.426	EGFR 745KLE>A>K	10.65%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 21	IV	1.3	0.687	-0.757	EGFR 745KLE>A>K	0.49%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 0	IV	1.3	0.469	-0.376	EGFR 851L>R	6.47%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 9	IV	1.3	0.674	-0.746	EGFR 851L>R	1.72%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 42	IV	1.3	0.775	-0.685	EGFR 851L>R	5.28%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 0	IV	12.4	0.817	-0.630	EGFR 745KLE>A>D	0.03%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 7	IV	12.4	0.916	-0.811	EGFR 748EL>E>D	0.03%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 0.5	IV	12.4	0.658	-0.694	EGFR 748EL>E>D	0.08%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 7	IV	12.4	0.552	-0.848	EGFR 748EL>E>D	0.00%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 17	IV	12.4	0.684	-0.728	EGFR 747NL>A>S>D	0.42%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 0	IV	1.7	0.864	-0.728	EGFR 747NL>A>S>D	0.20%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 7	IV	6.7	0.908	-0.803	EGFR 747NL>A>S>D	0.00%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 22	IV	6.7	0.863	-0.681	EGFR 747NL>A>S>D	0.00%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 53	IV	1.4	0.331	-0.351	EGFR L861Q	15.72%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 4	IV	1.4	0.225	-0.253	EGFR L861Q	45.67%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 18	IV	1.4	0.336	-0.364	EGFR G719A	33.38%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 37	IV	1.4	0.340	-0.364	EGFR L861Q	66.01%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 21	IV	Ongoing	0.935	-0.818	EGFR E746 A>P>D	0.00%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 22	IV	Ongoing	0.916	-0.774	EGFR E746 A>P>D	0.22%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 53	IV	Ongoing	0.953	-0.860	EGFR E746 A>P>D	0.48%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 4	IV	Ongoing	0.944	-0.832	EGFR E746 A>P>D	0.00%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 18	IV	Ongoing	0.925	-0.826	EGFR L861R	20.61%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 37	IV	Ongoing	0.950	-0.903	EGFR E746 A>P>D	0.22%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 0	IV	Ongoing	0.945	-0.889	EGFR E746 A>P>D	0.16%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day C1875	IV	Ongoing	0.886	-0.883	EGFR E746 A>P>D	0.16%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 59	IV	Ongoing	0.944	-0.804	EGFR E746 A>P>D	0.82%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 2	IV	7.5	0.958	-0.853	EGFR E746 A>P>D	0.00%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 12	IV	7.5	0.967	-0.866	EGFR E746 A>P>D	0.15%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 38	IV	7.5	0.951	-0.890	EGFR E746 A>P>D	0.00%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 110	IV	7.5	0.925	-0.400	EGFR E746 A>P>D	7.65%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 5	IV	1.5	0.272	-0.257	EGFR E746 A>P>D	13.19%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day -2	IV	1.5	0.564	-0.536	EGFR D761N	6.03%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 0.125	IV	1.5	0.530	-0.513	EGFR D761N	9.28%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 7	IV	Ongoing	0.946	-0.824	EGFR D761N	0.00%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 47	IV	Ongoing	0.927	-0.788	EGFR D761N	0.15%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 2	IV	Ongoing	0.952	-0.856	EGFR D761N	0.00%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 27	IV	Ongoing	0.930	-0.894	EGFR D761N	0.00%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 55	IV	Ongoing	0.920	-0.855	EGFR L861R	NA
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 91	IV	Ongoing	0.946	-0.824	EGFR L861R	NA
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day -1	IV	Ongoing	0.946	-0.842	EGFR L861R	0.21%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 6	IV	Ongoing	0.955	-0.844	EGFR L861R	0.21%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 84	IV	Ongoing	0.946	-0.825	EGFR L861R	0.00%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Pre-treatment, Day 0	IV	9.6	0.961	-0.904	EGFR L861R	0.00%
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 18	IV	9.6	0.959	-0.886	EGFR L861R	NA
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 55	IV	9.6	0.961	-0.886	EGFR L861R	NA
CGLU1246	Lung Cancer	Targeted Mutation Analysis and WES	Post-treatment, Day 90	IV	9.6	0.958	-0.885	EGFR L861R	NA

Patient	Patient Type	Analysis Type	Timepoint	Stage	Progression-free Survival (months)	Correlation of Fragment Ratio Profile to Median Fragment Ratio Profile of Healthy Individuals	Correlation of Fragment Ratio Profile to Lymphocyte Nucleosome Distances	Targeted Mutation	Maximum Mutant Allele Fraction
CGPLU267	Lung Cancer	Targeted Mutation Analysis and WGS	Pre-treatment, Day -1	IV	3.9	0.919	0.863	EGFR.L858R	1.93%
CGPLU267	Lung Cancer	Targeted Mutation Analysis and WGS	Post-treatment, Day 34	IV	3.9	0.863	-0.889	EGFR.L858R	0.14%
CGPLU267	Lung Cancer	Targeted Mutation Analysis and WGS	Post-treatment, Day 90	IV	3.9	0.962	-0.876	EGFR.L858R	0.38%
CGPLU269	Lung Cancer	Targeted Mutation Analysis and WGS	Pre-treatment, Day 0	IV	Ongoing	0.951	-0.864	EGFR.L858R	0.19%
CGPLU269	Lung Cancer	Targeted Mutation Analysis and WGS	Post-treatment, Day 9	IV	Ongoing	0.941	-0.884	EGFR.L858R	0.052%
CGPLU269	Lung Cancer	Targeted Mutation Analysis and WGS	Post-treatment, Day 28	IV	Ongoing	0.957	-0.876	EGFR.L858R	0.00%
CGPLU271	Lung Cancer	Targeted Mutation Analysis and WGS	Pre-treatment, Day 0	IV	8.2	0.371	-0.284	EGFR.E746_A750del	3.36%
CGPLU271	Lung Cancer	Targeted Mutation Analysis and WGS	Post-treatment, Day 6	IV	8.2	0.947	-0.826	EGFR.E746_A750del	0.17%
CGPLU271	Lung Cancer	Targeted Mutation Analysis and WGS	Post-treatment, Day 26	IV	8.2	0.952	-0.839	EGFR.E746_A750del	0.03%
CGPLU271	Lung Cancer	Targeted Mutation Analysis and WGS	Post-treatment, Day 104	IV	8.2	0.944	-0.810	EGFR.E746_A750del	0.05%
CGPLU271	Lung Cancer	Targeted Mutation Analysis and WGS	Post-treatment, Day 259	IV	8.2	0.950	-0.831	EGFR.E746_A750del	0.44%
CGPLU143	Lung Cancer	Targeted Mutation Analysis and WGS	Pre-treatment, Day -1	IV	Ongoing	0.944	-0.903	NA	0.05%
CGPLU143	Lung Cancer	Targeted Mutation Analysis and WGS	Post-treatment, Day 6	IV	Ongoing	0.956	-0.889	NA	0.00%
CGPLU143	Lung Cancer	Targeted Mutation Analysis and WGS	Post-treatment, Day 27	IV	Ongoing	0.959	-0.901	NA	0.00%
CGPLU143	Lung Cancer	Targeted Mutation Analysis and WGS	Post-treatment, Day 83	IV	Ongoing	0.965	0.886	NA	0.00%

APPENDIX G: Table 7. Whole genome cfDNA analyses in healthy individuals and cancer patients

Patient	Patient Type	Analysis Type	Timepoint	Stage at Diagnosis	Median cfDNA Fragment Size (bp)	Correlation of SgC Correlated Fragment Profile to Median Fragment Ratio	Correlation of SgC Correlated Fragment Profile to Median Fragment Ratio	Fraction of Reads Mapped to Mitochondrial Genome	DELFt Score	DELFt Score (%) specificity	DELFt Detected using DELFt	Mutant Allele Fraction Detected using Targeted sequencing ^a
CGPLB75	Breast Cancer	Targeted Mutation Analysis and WGS	Preoperative treatment naïve	I	170	0.9110	0.9254	0.0775%	0.9334	0.9334	Y	0.12%
CGPLB181	Breast Cancer	Targeted Mutation Analysis and WGS	Preoperative treatment naïve	I	170	0.9542	0.9193	0.0241%	0.9895	0.9895	Y	Y
CGPLB182	Breast Cancer	Targeted Mutation Analysis and WGS	Preoperative treatment naïve	I	165	0.9254	0.9649	0.1640%	0.9834	0.9834	Y	0.28%
CGPLB183	Breast Cancer	Targeted Mutation Analysis and WGS	Preoperative treatment naïve	I	188	0.9451	0.9738	0.0419%	0.9810	0.9810	Y	-
CGPLB184	Breast Cancer	Targeted Mutation Analysis and WGS	Preoperative treatment naïve	I	169	0.9315	0.8855	0.0224%	0.9801	0.9801	Y	-
CGPLB187	Breast Cancer	Targeted Mutation Analysis and WGS	Preoperative treatment naïve	I	165	0.9554	0.9737	0.0224%	0.9898	0.9898	Y	0.45%
CGPLB188	Breast Cancer	Targeted Mutation Analysis and WGS	Preoperative treatment naïve	I	163	0.9570	0.9847	0.0181%	0.9898	0.9898	Y	0.36%
CGPLB189	Breast Cancer	Targeted Mutation Analysis and WGS	Preoperative treatment naïve	I	163	0.9539	0.9839	0.0181%	0.9857	0.9857	Y	-
CGPLB190	Breast Cancer	Targeted Mutation Analysis and WGS	Preoperative treatment naïve	I	164	0.7955	0.9403	0.0789%	0.8710	0.8710	Y	ND
CGPLB191	Breast Cancer	Targeted Mutation Analysis and WGS	Preoperative treatment naïve	I	162	0.6774	0.8935	0.1942%	0.9686	0.9686	Y	0.20%
CGPLB192	Breast Cancer	Targeted Mutation Analysis and WGS	Preoperative treatment naïve	I	164	0.8773	0.9076	0.0352%	0.7253	0.7253	Y	ND
CGPLB193	Breast Cancer	Targeted Mutation Analysis and WGS	Preoperative treatment naïve	I	168	0.9255	0.8847	0.0521%	0.9746	0.9746	N	-
CGPLH180	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9255	0.9268	0.1193%	0.5168	0.5168	N	-
CGPLH181	Healthy	WGS	Preoperative treatment naïve	NA	163	0.9546	0.9457	0.0276%	0.9173	0.9173	N	-
CGPLH184	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9223	0.9429	0.0420%	0.5734	0.5734	N	-
CGPLH185	Healthy	WGS	Preoperative treatment naïve	NA	158	0.9567	0.9293	0.0467%	0.9165	0.9165	N	-
CGPLH187	Healthy	WGS	Preoperative treatment naïve	NA	163	0.9238	0.9512	0.0286%	0.9388	0.9388	N	-
CGPLH189	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9635	0.9416	0.0344%	0.4639	0.4639	N	-
CGPLH190	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9238	0.9457	0.0382%	0.8571	0.8571	N	-
CGPLH192	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9618	0.9639	0.0101%	0.5584	0.5584	N	-
CGPLH193	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9182	0.9391	0.0382%	0.9835	0.9835	Y	Y
CGPLH194	Healthy	WGS	Preoperative treatment naïve	NA	158	0.9548	0.9180	0.0404%	0.9547	0.9547	N	-
CGPLH195	Healthy	WGS	Preoperative treatment naïve	NA	158	0.9471	0.9436	0.0501%	0.9085	0.9085	N	-
CGPLH197	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9534	0.9575	0.0485%	0.2485	0.2485	N	-
CGPLH198	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9575	0.9285	0.0409%	0.4431	0.4431	N	-
CGPLH199	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9422	0.9409	0.0313%	0.9236	0.9236	N	-
CGPLH200	Healthy	WGS	Preoperative treatment naïve	NA	169	0.9556	0.9267	0.0427%	0.2223	0.2223	N	-
CGPLH203	Healthy	WGS	Preoperative treatment naïve	NA	163	0.9447	0.9787	0.0323%	0.9835	0.9835	Y	Y
CGPLH204	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9535	0.9180	0.0404%	0.9385	0.9385	Y	Y
CGPLH203	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9471	0.9436	0.0317%	0.9085	0.9085	N	-
CGPLH205	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9534	0.9293	0.0387%	0.9388	0.9388	N	-
CGPLH206	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9575	0.9767	0.0226%	0.1788	0.1788	N	-
CGPLH208	Healthy	WGS	Preoperative treatment naïve	NA	158	0.9811	0.9265	0.0311%	0.9441	0.9441	N	-
CGPLH209	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9422	0.9455	0.0223%	0.9236	0.9236	N	-
CGPLH210	Healthy	WGS	Preoperative treatment naïve	NA	163	0.9556	0.9527	0.0323%	0.9390	0.9390	Y	Y
CGPLH211	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9535	0.9393	0.0322%	0.2589	0.2589	N	-
CGPLH211	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9119	0.9410	0.0317%	0.1752	0.1752	N	-
CGPLH213	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9776	0.9205	0.0387%	0.9388	0.9388	N	-
CGPLH217	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9765	0.9767	0.0226%	0.3459	0.3459	N	-
CGPLH218	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9811	0.9352	0.0311%	0.1788	0.1788	N	-
CGPLH219	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9556	0.9455	0.0223%	0.9441	0.9441	N	-
CGPLH219	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9547	0.9397	0.0317%	0.9735	0.9735	Y	Y
CGPLH221	Healthy	WGS	Preoperative treatment naïve	NA	163	0.9535	0.9346	0.0317%	0.9735	0.9735	Y	Y
CGPLH221	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9482	0.9491	0.0317%	0.9388	0.9388	N	-
CGPLH224	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9858	0.9491	0.0317%	0.1632	0.1632	N	-
CGPLH225	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9376	0.9427	0.0464%	0.3469	0.3469	N	-
CGPLH226	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9525	0.9352	0.0311%	0.4637	0.4637	N	-
CGPLH227	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH229	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9518	0.9787	0.0223%	0.2232	0.2232	N	-
CGPLH231	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9392	0.9527	0.0224%	0.2589	0.2589	N	-
CGPLH234	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9482	0.9491	0.0317%	0.9388	0.9388	N	-
CGPLH235	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9858	0.9451	0.0317%	0.3469	0.3469	N	-
CGPLH236	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9374	0.9352	0.0311%	0.1632	0.1632	N	-
CGPLH237	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH239	Healthy	WGS	Preoperative treatment naïve	NA	164	0.9518	0.9787	0.0223%	0.2232	0.2232	N	-
CGPLH241	Healthy	WGS	Preoperative treatment naïve	NA	164	0.9733	0.9165	0.0224%	0.9735	0.9735	Y	Y
CGPLH242	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9851	0.9451	0.0317%	0.9388	0.9388	N	-
CGPLH244	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9518	0.9451	0.0317%	0.9388	0.9388	N	-
CGPLH245	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH246	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH247	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9518	0.9787	0.0223%	0.2232	0.2232	N	-
CGPLH249	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH250	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH251	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH252	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH253	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH254	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH255	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH256	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH257	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH258	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH259	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH260	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH261	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH262	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH263	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH264	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH265	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH266	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH267	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH268	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH269	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH270	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH271	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH272	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH273	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH274	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH275	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	-
CGPLH276	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9512	0.9352	0.0222%	0.1310	0.1310	N	

Patient	Patient Type	Analysis Type	Timepoint	Stage at Diagnosis	Median cfDNA, Fragment Size (bp)	Correlation of 3G Corrected Fragment Profile to Median Fragment Ratio	Fraction of Reads Mapped to Mitochondrial Genome	DELFt Scores	Detected using DELFt	Mutant Allele Fraction Detected using Targeted sequencing ^a
						Profile of Healthy Individuals		(88% specificity)		
CGPL1360	Healthy	WGS	Preoperative treatment naïve	NA	168	0.8201	0.8575	0.0008	N	N
CGPL1361	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9521	0.9265%	0.1624	N	N
CGPL1362	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9236	0.9803	0.4832	N	N
CGPL1363	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9498	0.9520%	0.0339%	N	N
CGPL1364	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9311	0.9480	0.1628%	Y	Y
CGPL1365	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9321	0.9851	0.1747%	Y	Y
CGPL1366	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9536	0.9710	0.0344%	N	N
CGPL1367	Healthy	WGS	Preoperative treatment naïve	NA	165	0.8246	0.9181	0.1633%	N	N
CGPL1368	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9430	0.8076	0.1033%	N	N
CGPL1369	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9228	0.9541	0.0216%	N	N
CGPL1370	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9642	0.8423	0.0410%	N	N
CGPL1371	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9621	0.9414	0.0749%	N	N
CGPL1380	Healthy	WGS	Preoperative treatment naïve	NA	170	0.9652	0.9624	0.0653%	N	N
CGPL1381	Healthy	WGS	Preoperative treatment naïve	NA	165	0.8541	0.9040	0.0435%	N	N
CGPL1382	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9380	0.9407	0.0326%	N	N
CGPL1383	Healthy	WGS	Preoperative treatment naïve	NA	158	0.9730	0.9811	0.0325%	N	N
CGPL1384	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9861	0.9043	0.0287%	N	N
CGPL1385	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9856	0.9245	0.0734%	N	N
CGPL1386	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9820	0.8859	0.0565%	N	N
CGPL1387	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9893	0.9232	0.0529%	N	N
CGPL1388	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9246	0.9256	0.0325%	N	N
CGPL1389	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9409	0.9036	0.0340%	N	N
CGPL1390	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9216	0.9747	0.1389%	N	N
CGPL1391	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9334	0.9622	0.0287%	N	N
CGPL1392	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9165	0.9240	0.0746%	N	N
CGPL1393	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9014	0.9454%	0.1454%	N	N
CGPL1394	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9256	0.9407%	0.0407%	N	N
CGPL1395	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9252	0.9527%	0.0527%	N	N
CGPL1396	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9448	0.9235	0.0349%	N	N
CGPL1397	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9409	0.9764	0.1627%	N	N
CGPL1398	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9633	0.9767	0.1223%	N	N
CGPL1399	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9324	0.9622	0.0223%	N	N
CGPL1400	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9165	0.9562	0.0562%	N	N
CGPL1401	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9047	0.9352	0.0307%	N	N
CGPL1402	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9227	0.9256	0.0325%	N	N
CGPL1403	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9511	0.9246	0.0442%	N	N
CGPL1404	Healthy	WGS	Preoperative treatment naïve	NA	167	0.7884	0.8920	0.1657%	Y	Y
CGPL1405	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9633	0.9762	0.1223%	N	N
CGPL1406	Healthy	WGS	Preoperative treatment naïve	NA	165	0.8200	0.9878	0.0746%	N	N
CGPL1407	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9165	0.9795	0.0517%	N	N
CGPL1408	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9047	0.9047	0.0307%	N	N
CGPL1409	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9227	0.9040	0.0307%	N	N
CGPL1410	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9250	0.9890	0.0523%	N	N
CGPL1411	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9673	0.9204	0.0358%	N	N
CGPL1412	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9198	0.9592	0.0587%	N	N
CGPL1413	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9227	0.9093	0.0326%	N	N
CGPL1414	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9553	0.9327	0.0407%	N	N
CGPL1415	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9554	0.9145%	0.0229%	N	N
CGPL1416	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9554	0.9592	0.1616%	N	N
CGPL1417	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9229	0.8950	0.0515%	N	N
CGPL1418	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9376	0.9204	0.0589%	N	N
CGPL1419	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9551	0.9791	0.0517%	N	N
CGPL1420	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9145	0.9352	0.0307%	N	N
CGPL1421	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9554	0.9727	0.0415%	N	N
CGPL1422	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9227	0.8950	0.0323%	N	N
CGPL1423	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9572	0.9098	0.0453%	N	N
CGPL1424	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9312	0.9005	0.0327%	N	N
CGPL1425	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9857	0.9792	0.0241%	N	N
CGPL1426	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9376	0.9857	0.0517%	N	N
CGPL1427	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9551	0.9791	0.0407%	N	N
CGPL1428	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9227	0.9093	0.0307%	N	N
CGPL1429	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9554	0.9727	0.0429%	N	N
CGPL1430	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9553	0.9597	0.0345%	N	N
CGPL1431	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9222	0.9481	0.0722%	N	N
CGPL1432	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9560	0.9809	0.0548%	N	N
CGPL1433	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9154	0.9295%	0.0269%	N	N
CGPL1434	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9554	0.9727	0.0187%	N	N
CGPL1435	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9217	0.9095	0.0286%	N	N
CGPL1436	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9553	0.9597	0.0345%	N	N
CGPL1437	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9553	0.9597	0.0345%	N	N
CGPL1438	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9553	0.9597	0.0345%	N	N
CGPL1439	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9547	0.9597	0.0345%	N	N
CGPL1440	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9552	0.9597	0.0345%	N	N
CGPL1441	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9553	0.9597	0.0345%	N	N

Patient	Patient Type	Analysis Type	Timepoint	Stage at Diagnosis	Median cfDNA, Fragment Size (bp)	Correlation of GC Content/Fraction of Median Fragment Ratio to Median Fragment Profile of Healthy Individuals	Fraction of Reads Mapped to Mitochondrial Genome	DELFt Scores	DELFt Detected using DELFt (88% specificity)	Mutant Allele Fraction Detected using Targeted sequencing*
CGPL1443	Healthy	WGS	Preoperative treatment naïve	NA	170	0.3131	0.4801	0.0227%	N	N
CGPL1444	Healthy	WGS	Preoperative treatment naïve	NA	171	0.3048	0.3048	0.1644%	N	N
CGPL1445	Healthy	WGS	Preoperative treatment naïve	NA	171	0.3426	0.6750	0.0267%	0.1939	N
CGPL1446	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9032	0.9032	0.3221%	N	N
CGPL1447	Healthy	WGS	Preoperative treatment naïve	NA	169	0.9211	0.8893	0.0167%	0.03017	N
CGPL1448	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9553	0.9191	0.0461%	0.3588	N
CGPL1449	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9850	0.9245	0.0238%	0.0116	N
CGPL1450	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9572	0.9195	0.0331%	0.0597	N
CGPL1451	Healthy	WGS	Preoperative treatment naïve	NA	169	0.9658	0.9165	0.0282%	0.0194	N
CGPL1452	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9498	0.8948	0.0469%	0.4722	N
CGPL1453	Healthy	WGS	Preoperative treatment naïve	NA	166	0.9572	0.8239	0.0168%	0.3419	N
CGPL1455	Healthy	WGS	Preoperative treatment naïve	NA	166	0.9226	0.9222	0.0465%	0.4536	N
CGPL1456	Healthy	WGS	Preoperative treatment naïve	NA	166	0.9537	0.9096	0.0287%	0.0240	N
CGPL1457	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9423	0.9042	0.0286%	0.0384	N
CGPL1458	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9511	0.9275	0.0288%	0.1691	N
CGPL1459	Healthy	WGS	Preoperative treatment naïve	NA	158	0.9039	0.9209	0.0221%	0.3237	N
CGPL1460	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9331	0.8963	0.0227%	0.1157	N
CGPL1463	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9536	0.9242	0.0153%	0.3685	N
CGPL1464	Healthy	WGS	Preoperative treatment naïve	NA	170	0.9133	0.8811	0.0289%	0.2040	N
CGPL1465	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9551	0.9164	0.0235%	0.0124	N
CGPL1466	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9573	0.9403	0.0155%	0.1723	N
CGPL1467	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9273	0.8744	0.0239%	0.2338	N
CGPL1468	Healthy	WGS	Preoperative treatment naïve	NA	167	0.8853	0.8245	0.0247%	0.5427	N
CGPL1469	Healthy	WGS	Preoperative treatment naïve	NA	168	0.8825	0.6739	0.0261%	0.5551	N
CGPL1470	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9073	0.9228	0.0715%	0.3327	N
CGPL1471	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9354	0.9333	0.0159%	0.0406	N
CGPL1472	Healthy	WGS	Preoperative treatment naïve	NA	165	0.8808	0.8815	0.0461%	0.6152	N
CGPL1473	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9246	0.9728	0.0163%	0.2395	N
CGPL1474	Healthy	WGS	Preoperative treatment naïve	NA	168	0.8744	0.9445	0.0316%	0.5246	Y
CGPL1475	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9155	0.9233	0.0288%	0.0547	N
CGPL1476	Healthy	WGS	Preoperative treatment naïve	NA	168	0.8825	0.8093	0.0223%	0.0736	N
CGPL1477	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9027	0.9059	0.0130%	0.0103	N
CGPL1478	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9354	0.9344	0.0282%	0.1111	N
CGPL1479	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9207	0.9093	0.0221%	0.0546	N
CGPL1480	Healthy	WGS	Preoperative treatment naïve	NA	163	0.9522	0.9046	0.0257%	0.7473	Y
CGPL1481	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9568	0.9113	0.0318%	0.3282	N
CGPL1482	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9773	0.9336	0.0162%	0.0363	N
CGPL1483	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9516	0.9276	0.0221%	0.0495	N
CGPL1484	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9530	0.9356	0.0265%	0.0268	N
CGPL1485	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9547	0.9293	0.0221%	0.0548	N
CGPL1486	Healthy	WGS	Preoperative treatment naïve	NA	163	0.9522	0.9046	0.0257%	0.2338	N
CGPL1487	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9568	0.9113	0.0318%	0.3282	N
CGPL1488	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9773	0.9336	0.0162%	0.0363	N
CGPL1489	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9516	0.9276	0.0221%	0.0495	N
CGPL1490	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9530	0.9356	0.0265%	0.0268	N
CGPL1491	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9547	0.9293	0.0221%	0.0548	N
CGPL1492	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9198	0.8744	0.0220%	0.0268	N
CGPL1493	Healthy	WGS	Preoperative treatment naïve	NA	169	0.9575	0.9046	0.0254%	0.2154	N
CGPL1494	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9618	0.9293	0.0469%	0.3933	N
CGPL1495	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9550	0.9194	0.0432%	0.3424	N
CGPL1496	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9530	0.9356	0.0281%	0.0306	N
CGPL1497	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9547	0.9293	0.0221%	0.0548	N
CGPL1498	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9522	0.9046	0.0257%	0.2338	N
CGPL1499	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9575	0.9046	0.0222%	0.0548	N
CGPL1500	Healthy	WGS	Preoperative treatment naïve	NA	166	0.8777	0.8808	0.0285%	0.0465	N
CGPL1501	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9618	0.9293	0.0223%	0.0572	N
CGPL1502	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9550	0.9330	0.0359%	0.3434	N
CGPL1503	Healthy	WGS	Preoperative treatment naïve	NA	170	0.9530	0.9356	0.0323	0.3323	N
CGPL1504	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9547	0.9293	0.0221%	0.0548	N
CGPL1505	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9522	0.9046	0.0257%	0.2338	N
CGPL1506	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9575	0.9046	0.0222%	0.0548	N
CGPL1507	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9522	0.9046	0.0257%	0.0548	N
CGPL1508	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9577	0.9046	0.0222%	0.0548	N
CGPL1509	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9547	0.9356	0.0359%	0.3578	N
CGPL1510	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9551	0.9455	0.0403%	0.1231	N
CGPL1511	Healthy	WGS	Preoperative treatment naïve	NA	165	0.9554	0.9442	0.0418%	0.3146	N
CGPL1512	Healthy	WGS	Preoperative treatment naïve	NA	166	0.9523	0.9046	0.0257%	0.2338	N
CGPL1513	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9575	0.9046	0.0222%	0.0548	N
CGPL1514	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9522	0.9046	0.0257%	0.0548	N
CGPL1515	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9574	0.9046	0.0222%	0.0548	N
CGPL1516	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9547	0.9356	0.0359%	0.3578	N
CGPL1517	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9551	0.9455	0.0403%	0.1231	N
CGPL1518	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9554	0.9442	0.0418%	0.3146	N
CGPL1519	Healthy	WGS	Preoperative treatment naïve	NA	166	0.9523	0.9046	0.0257%	0.2338	N
CGPL1520	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9575	0.9046	0.0222%	0.0548	N
CGPL1521	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9522	0.9046	0.0257%	0.0548	N
CGPL1522	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9577	0.9046	0.0222%	0.0548	N
CGPL1523	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9547	0.9356	0.0359%	0.3578	N
CGPL1524	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9551	0.9455	0.0403%	0.1231	N
CGPL1525	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9554	0.9442	0.0418%	0.3146	N
CGPL1526	Healthy	WGS	Preoperative treatment naïve	NA	166	0.9523	0.9046	0.0257%	0.2338	N
CGPL1527	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9574	0.9046	0.0222%	0.0548	N
CGPL1528	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9547	0.9356	0.0359%	0.3578	N
CGPL1529	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9551	0.9455	0.0403%	0.1231	N
CGPL1530	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9554	0.9442	0.0418%	0.3146	N
CGPL1531	Healthy	WGS	Preoperative treatment naïve	NA	166	0.9523	0.9046	0.0257%	0.2338	N
CGPL1532	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9575	0.9046	0.0222%	0.0548	N
CGPL1533	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9547	0.9356	0.0359%	0.3578	N
CGPL1534	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9551	0.9455	0.0403%	0.1231	N
CGPL1535	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9554	0.9442	0.0418%	0.3146	N
CGPL1536	Healthy	WGS	Preoperative treatment naïve	NA	166	0.9523	0.9046	0.0257%	0.2338	N
CGPL1537	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9574	0.9046	0.0222%	0.0548	N
CGPL1538	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9547	0.9356	0.0359%	0.3578	N
CGPL1539	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9551	0.9455	0.0403%	0.1231	N
CGPL1540	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9554	0.9442	0.0418%	0.3146	N
CGPL1541	Healthy	WGS	Preoperative treatment naïve	NA	166	0.9523	0.9046	0.0257%	0.2338	N
CGPL1542	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9575	0.9046	0.0222%	0.0548	N
CGPL1543	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9547	0.9356	0.0359%	0.3578	N
CGPL1544	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9551	0.9455	0.0403%	0.1231	N
CGPL1545	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9554	0.9442	0.0418%	0.3146	N
CGPL1546	Healthy	WGS	Preoperative treatment naïve	NA	166	0.9523	0.9046	0.0257%	0.2338	N
CGPL1547	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9574	0.9046	0.0222%	0.0548	N
CGPL1548	Healthy	WGS	Preoperative treatment naïve	NA	167	0.9547	0.9356	0.0359%	0.3578	N
CGPL1549	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9551	0.9455	0.0403%	0.1231	N
CGPL1550	Healthy	WGS	Preoperative treatment naïve	NA	168	0.9554	0.9442	0.0418%		

Please see reference 10 for additional information on target gene sequencing analyses. DCL⁺ indicates not detected; DCL⁻ indicates detected.

WHAT IS CLAIMED IS:

1. A method of determining a cell free DNA (cfDNA) fragmentation profile of a mammal, the method comprising:
 - processing cfDNA fragments obtained from a sample obtained from the mammal into sequencing libraries;
 - subjecting the sequencing libraries to low-coverage whole genome sequencing to obtain sequenced fragments;
 - mapping the sequenced fragments to a genome to obtain windows of mapped sequences; and
 - analyzing the windows of mapped sequences to determine cfDNA fragment lengths.
2. The method of claim 1, wherein the mapped sequences comprise tens to thousands of windows.
3. The method of claims 1-2, wherein the windows are non-overlapping windows.
4. The method of any one of claims 1-3, wherein the windows each comprise about 5 million base pairs.
5. The method of any one of claims 1-4, wherein a cfDNA fragmentation profile is determined within each window.
6. The method of any one of claims 1-5, wherein cfDNA fragmentation profile comprises a median fragment size.
7. The method of any one of claims 1-5, wherein cfDNA fragmentation profile comprises a fragment size distribution.

8. The method of any one of claims 1-5, wherein the cfDNA fragmentation profile comprises a ratio of small cfDNA fragments to large cfDNA fragments in said windows of mapped sequences.
9. The method of any one of claims 1-5, wherein the cfDNA fragmentation profile comprises the sequence coverage of small cfDNA fragments in windows across the genome.
10. The method of any one of claims 1-5, wherein the cfDNA fragmentation profile comprises the sequence coverage of large cfDNA fragments in windows across the genome.
11. The method of any one of claims 1-5, wherein the cfDNA fragmentation profile comprises the sequence coverage of small and large cfDNA fragments in windows across the genome.
12. The method of any one of claims 1-11, wherein the cfDNA fragmentation profile is over the whole genome.
13. The method of any one of claims 1-11, wherein the cfDNA fragmentation profile is over a subgenomic interval.
14. A method of identifying a mammal as having cancer, the method comprising:
 - determining a cell free DNA (cfDNA) fragmentation profile in a sample obtained from the mammal;
 - comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile; and
 - identifying the mammal as having cancer when the cfDNA fragmentation profile obtained from the mammal is different from the reference cfDNA fragmentation profile.
15. The method of claim 14, wherein the reference cfDNA fragmentation profile is a cfDNA fragmentation profile of a healthy mammal.

16. The method of claim 15, wherein the reference cfDNA fragmentation profile is generated by determining a cfDNA fragmentation profile in a sample obtained from the healthy mammal.
17. The method of claim 14, wherein the reference DNA fragmentation pattern is a reference nucleosome cfDNA fragmentation profile.
18. The method of any one of claims 14-17, wherein the cfDNA fragmentation profile comprises a median fragment size, and wherein a median fragment size of the cfDNA fragmentation profile is shorter than a median fragment size of the reference cfDNA fragmentation profile.
19. The method of any one of claims 14-17, wherein the cfDNA fragmentation profile comprises a fragment size distribution, and wherein a fragment size distribution of the cfDNA fragmentation profile differs by at least 10 nucleotides as compared to a fragment size distribution of the reference cfDNA fragmentation profile.
20. The method of any one of claims 14-17, wherein the cfDNA fragmentation profile comprises a ratio of small cfDNA fragments to large cfDNA fragments in said windows of mapped sequences, wherein a small cfDNA fragment is 100 base pairs (bp) to 150 bp in length, wherein a large cfDNA fragments is 151 bp to 220 bp in length, and wherein a correlation of fragment ratios in the cfDNA fragmentation profile is lower than a correlation of fragment ratios of the reference cfDNA fragmentation profile.
21. The method of any one of claims 14-17, wherein the cfDNA fragmentation profile comprises the sequence coverage of small cfDNA fragments in windows across the genome.
22. The method of any one of claims 14-17, wherein the cfDNA fragmentation profile comprises the sequence coverage of large cfDNA fragments in windows across the genome.

23. The method of any one of claims 14-17, wherein the cfDNA fragmentation profile comprises the sequence coverage of small and large cfDNA fragments in windows across the genome.
24. The method of any one of claims 14-17, wherein the cancer is selected from the group consisting of: colorectal cancer, lung cancer, breast cancer, gastric cancers, pancreatic cancers, bile duct cancers, and ovarian cancer.
25. The method of claim 14, wherein the step of comparing comprises comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile over the whole genome.
26. The method of claim 14, wherein the step of comparing comprises comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile over a subgenomic interval.
27. The method of any one of claim 14-23, wherein the mammal has previously been administered a cancer treatment to treat the cancer.
28. The method of claim 27, wherein the cancer treatment is selected from the group consisting of: surgery, adjuvant chemotherapy, neoadjuvant chemotherapy, radiation therapy, hormone therapy, cytotoxic therapy, immunotherapy, adoptive T cell therapy, targeted therapy, and combinations thereof.
29. The method of any one of claims 14-28, further comprising administering to the mammal a cancer treatment selected from the group consisting of: surgery, adjuvant chemotherapy, neoadjuvant chemotherapy, radiation therapy, hormone therapy, cytotoxic therapy, immunotherapy, adoptive T cell therapy, targeted therapy, and combinations thereof.

30. The method of claim 29, wherein the mammal is monitored for the presence of cancer after administration of the cancer treatment.

31. The method of any one of claim 14 to claim 30, the method further comprising identifying one or more cancer-specific sequence alterations in the sample.

32. The method of any one of claim 14 to claim 30, the method further comprising identifying one or more chromosomal abnormalities in the sample.

33. The method of claim 32, wherein the one or more chromosomal abnormalities comprises a copy number change in one or more chromosome arms.

34. A method of identifying the tissue of origin of a cancer in a mammal identified as having a cancer, the method comprising:

determining a cell free DNA (cfDNA) fragmentation profile in a sample obtained from the mammal;

comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile; and

identifying the tissue of origin of the cancer in a mammal when the cfDNA fragmentation profile obtained from the mammal matches a reference cfDNA fragmentation profiles from a mammal identified as having a cancer with the same tissue of origin.

35. The method of claim 34, wherein the reference cfDNA fragmentation profile comprises reference cfDNA fragmentation profiles from mammals identified as having one or more of colorectal cancer, lung cancer, breast cancer, gastric cancers, pancreatic cancers, bile duct cancers, and ovarian cancer.

36. The method of claim 35, wherein the reference cfDNA fragmentation profile is generated by determining a cfDNA fragmentation profile in a sample obtained from the mammals

identified as having one or more or colorectal cancer, lung cancer, breast cancer, gastric cancers, pancreatic cancers, bile duct cancers, and ovarian cancer.

37. The method of claim 34, wherein the reference DNA fragmentation pattern is a reference nucleosome cfDNA fragmentation profile.

38. The method of any one of claims 34-37, wherein the cfDNA fragmentation profile comprises a median fragment size, and wherein a median fragment size of the cfDNA fragmentation profile is shorter than a median fragment size of the reference cfDNA fragmentation profile.

39. The method of any one of claims 34-37, wherein the cfDNA fragmentation profile comprises a fragment size distribution, and wherein a fragment size distribution of the cfDNA fragmentation profile differs by at least 10 nucleotides as compared to a fragment size distribution of the reference cfDNA fragmentation profile.

40. The method of any one of claims 34-37, wherein the cfDNA fragmentation profile comprises a ratio of small cfDNA fragments to large cfDNA fragments in said windows of mapped sequences, wherein a small cfDNA fragment is 100 base pairs (bp) to 150 bp in length, wherein a large cfDNA fragments is 151 bp to 220 bp in length, and wherein a correlation of fragment ratios in the cfDNA fragmentation profile is lower than a correlation of fragment ratios of the reference cfDNA fragmentation profile.

41. The method of any one of claims 34-37, wherein the cfDNA fragmentation profile comprises the sequence coverage of small cfDNA fragments in windows across the genome.

42. The method of any one of claims 34-37, wherein the cfDNA fragmentation profile comprises the sequence coverage of large cfDNA fragments in windows across the genome.

43. The method of any one of claims 34-37, wherein the cfDNA fragmentation profile comprises the sequence coverage of small and large cfDNA fragments in windows across the genome.
44. The method of any one of claims 34-37, wherein the cancer is selected from the group consisting of: colorectal cancer, lung cancer, breast cancer, gastric cancers, pancreatic cancers, bile duct cancers, and ovarian cancer.
45. The method of claim 34, wherein the step of comparing comprises comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile over the whole genome.
46. The method of claim 34, wherein the step of comparing comprises comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile over a subgenomic interval.
47. The method of any one of claims 34-46, the method further comprising identifying one or more cancer-specific sequence alterations in the sample.
48. The method of any one of claims 34-46, the method further comprising identifying one or more chromosomal abnormalities in the sample.
49. The method of claim 48, wherein the one or more chromosomal abnormalities comprises a copy number change in one or more chromosome arms.
50. A method treating a mammal having cancer, the method comprising:
identifying said mammal as having cancer, wherein said identifying comprises:
determining a cell free DNA (cfDNA) fragmentation profile in a sample obtained from the mammal;

comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile; and

identifying the mammal as having cancer when the cfDNA fragmentation profile obtained from the mammal is different from the reference cfDNA fragmentation profile; and

administering a cancer treatment to said mammal.

51. The method of claim 50, wherein said mammal is a human.

52. The method of any one of claims 50-51, wherein the cancer is selected from the group consisting of: colorectal cancer, lung cancer, breast cancer, gastric cancers, pancreatic cancers, bile duct cancers, and ovarian cancer.

53. The method of any one of claims 50-52, wherein said cancer treatment is selected from the group consisting of: surgery, adjuvant chemotherapy, neoadjuvant chemotherapy, radiation therapy, hormone therapy, cytotoxic therapy, immunotherapy, adoptive T cell therapy, targeted therapy, and combinations thereof.

54. The method of any one of claims 50-53, wherein the reference cfDNA fragmentation profile is a cfDNA fragmentation profile of a healthy mammal.

55. The method of claim 54, wherein the reference cfDNA fragmentation profile is generated by determining a cfDNA fragmentation profile in a sample obtained from the healthy mammal.

56. The method of any one of claims 50-53, wherein the reference DNA fragmentation pattern is a reference nucleosome cfDNA fragmentation profile.

57. The method of any one of claims 50-56, wherein the cfDNA fragmentation profile comprises a median fragment size, and wherein a median fragment size of the cfDNA

fragmentation profile is shorter than a median fragment size of the reference cfDNA fragmentation profile.

58. The method of any one of claims 50-56, wherein the cfDNA fragmentation profile comprises a fragment size distribution, and wherein a fragment size distribution of the cfDNA fragmentation profile differs by at least 10 nucleotides as compared to a fragment size distribution of the reference cfDNA fragmentation profile.

59. The method of any one of claims 50-56, wherein the cfDNA fragmentation profile comprises a ratio of small cfDNA fragments to large cfDNA fragments in said windows of mapped sequences, wherein a small cfDNA fragment is 100 base pairs (bp) to 150 bp in length, wherein a large cfDNA fragments is 151 bp to 220 bp in length, and wherein a correlation of fragment ratios in the cfDNA fragmentation profile is lower than a correlation of fragment ratios of the reference cfDNA fragmentation profile.

60. The method of any one of claims 50-56, wherein the cfDNA fragmentation profile comprises the sequence coverage of small cfDNA fragments in windows across the genome.

61. The method of any one of claims 50-56, wherein the cfDNA fragmentation profile comprises the sequence coverage of large cfDNA fragments in windows across the genome.

62. The method of any one of claims 50-56, wherein the cfDNA fragmentation profile comprises the sequence coverage of small and large cfDNA fragments in windows across the genome.

63. The method of any one of claims 50-62, wherein the step of comparing comprises comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile over the whole genome.

64. The method of any one of claims 50-62, wherein the step of comparing comprises comparing the cfDNA fragmentation profile to a reference cfDNA fragmentation profile over a subgenomic interval.
65. The method of any one of claims 50-64, wherein the mammal has previously been administered a cancer treatment to treat the cancer.
66. The method of claim 65, wherein the cancer treatment is selected from the group consisting of: surgery, adjuvant chemotherapy, neoadjuvant chemotherapy, radiation therapy, hormone therapy, cytotoxic therapy, immunotherapy, adoptive T cell therapy, targeted therapy, and combinations thereof.
67. The method of any one of claims 50-66, wherein the mammal is monitored for the presence of cancer after administration of the cancer treatment.

1/33

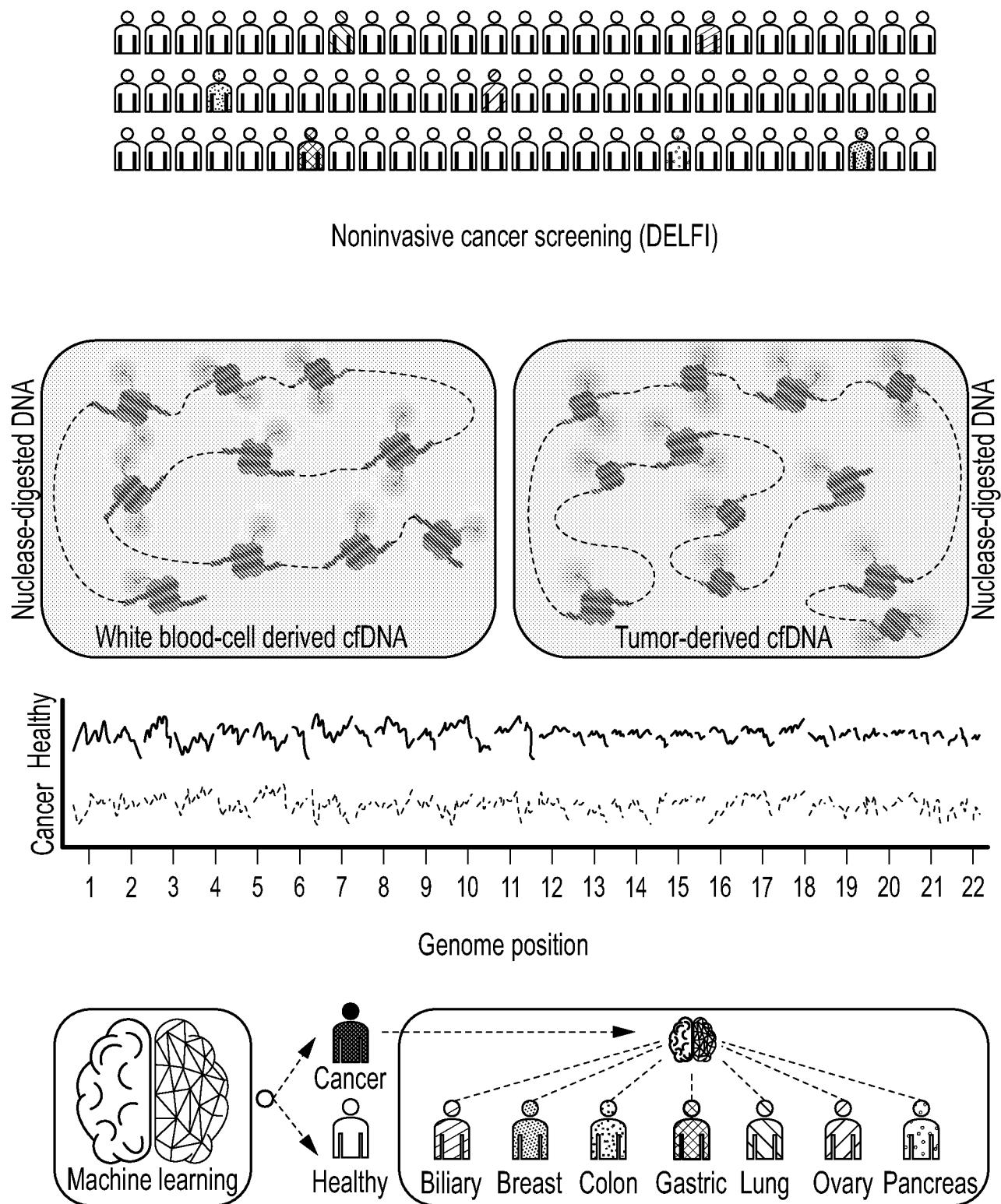


FIG. 1

2/33

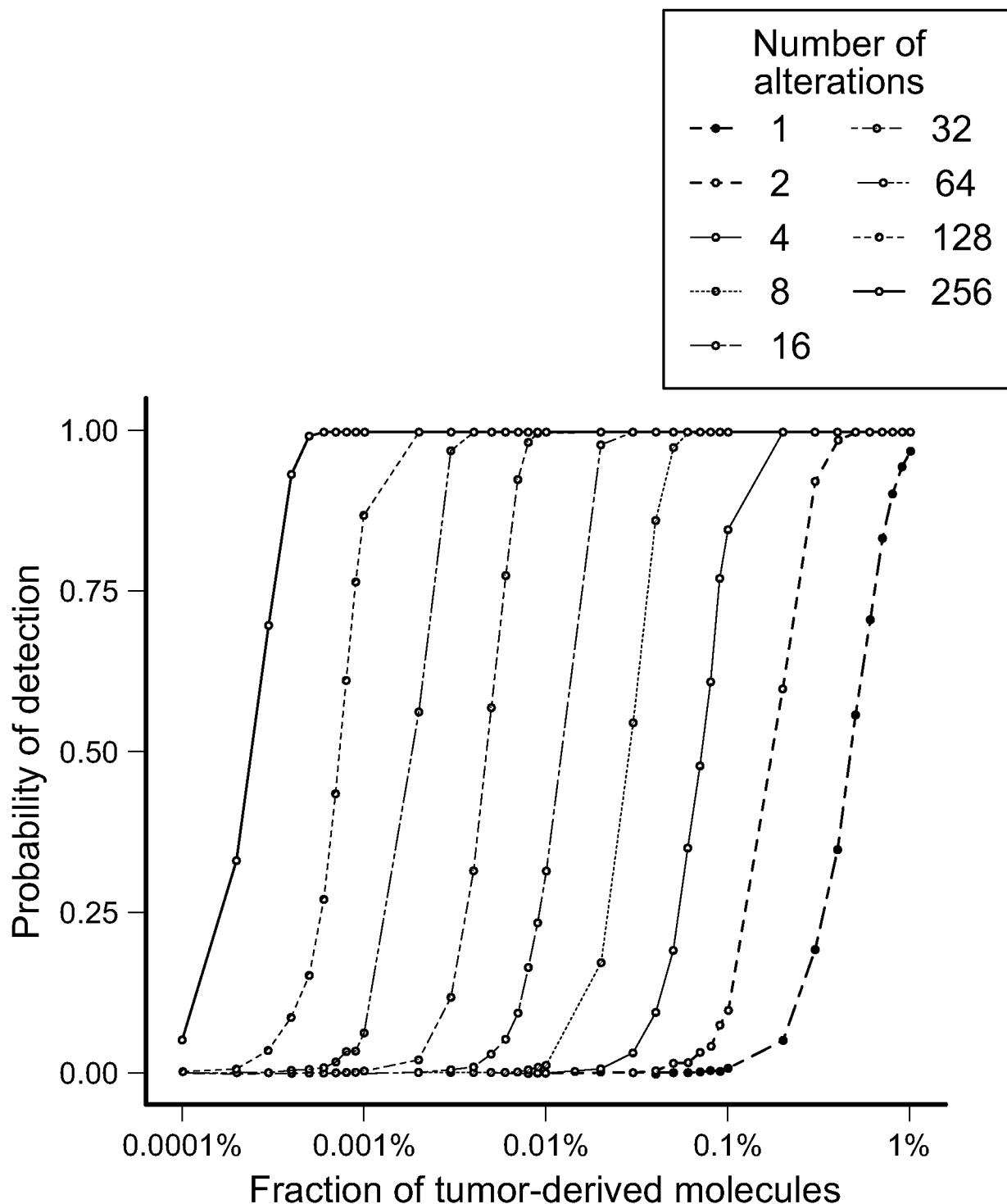


FIG. 2

3/33

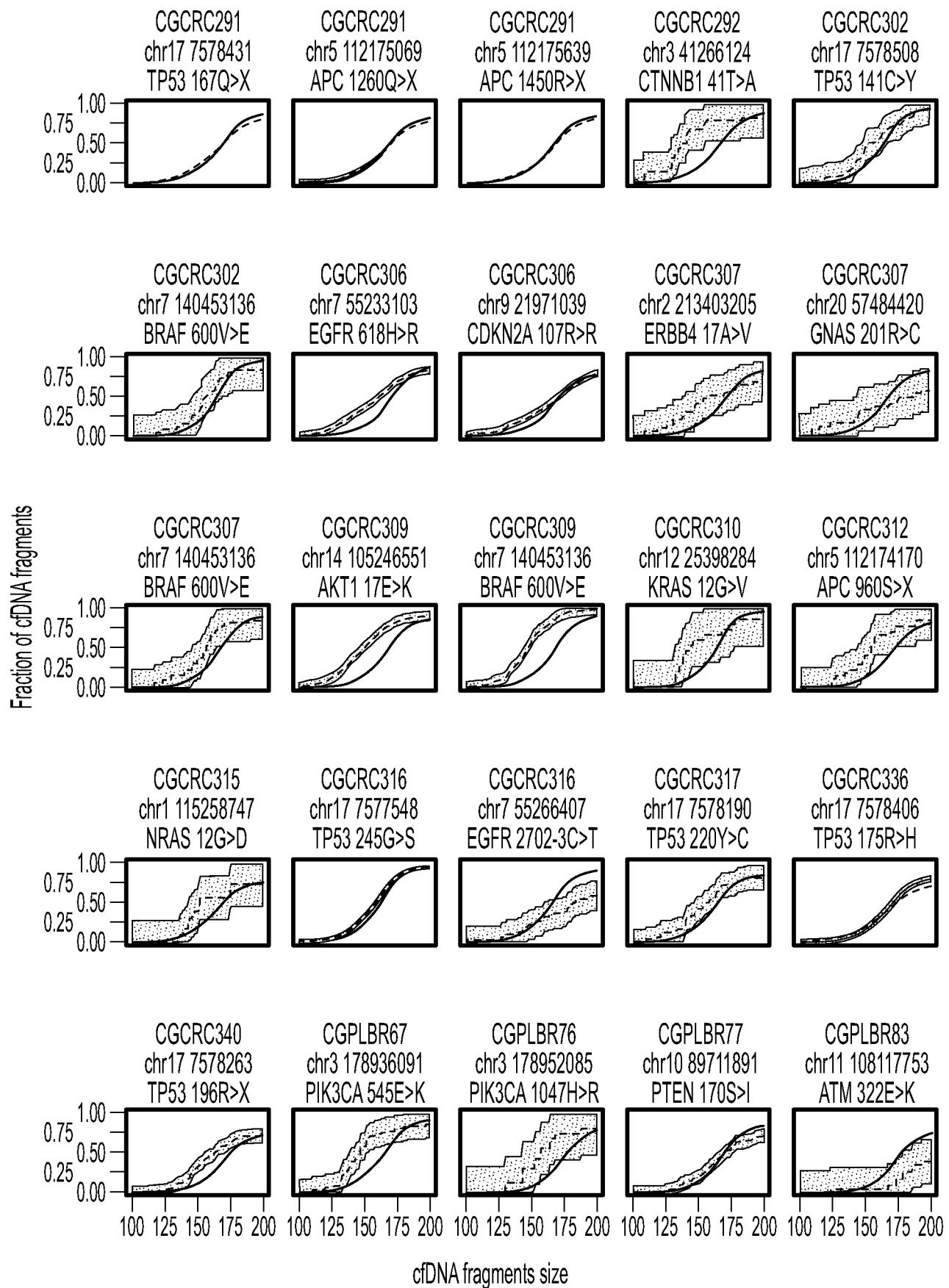


FIG. 3

4/33

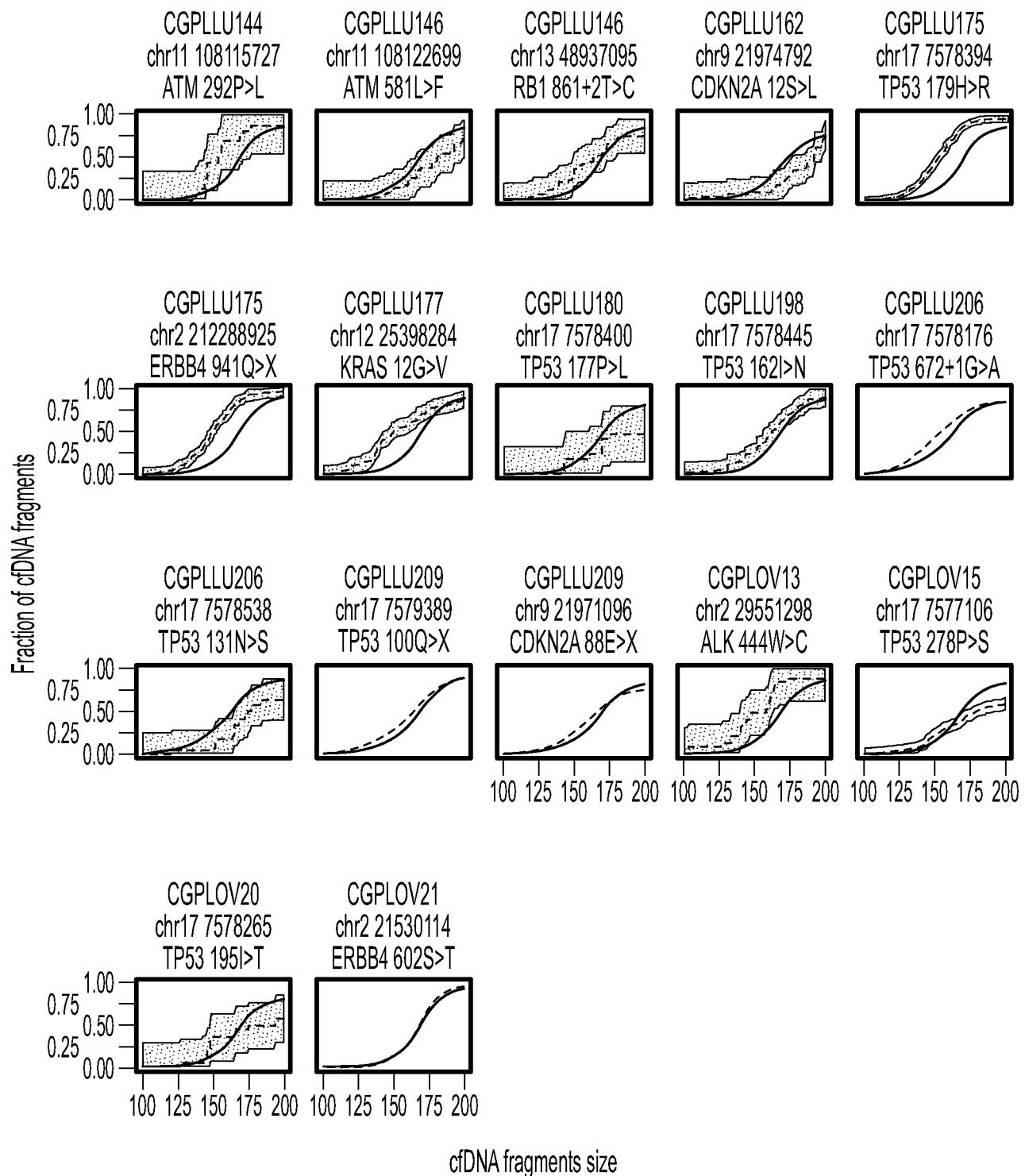


FIG. 3 (Cont.)

5/33

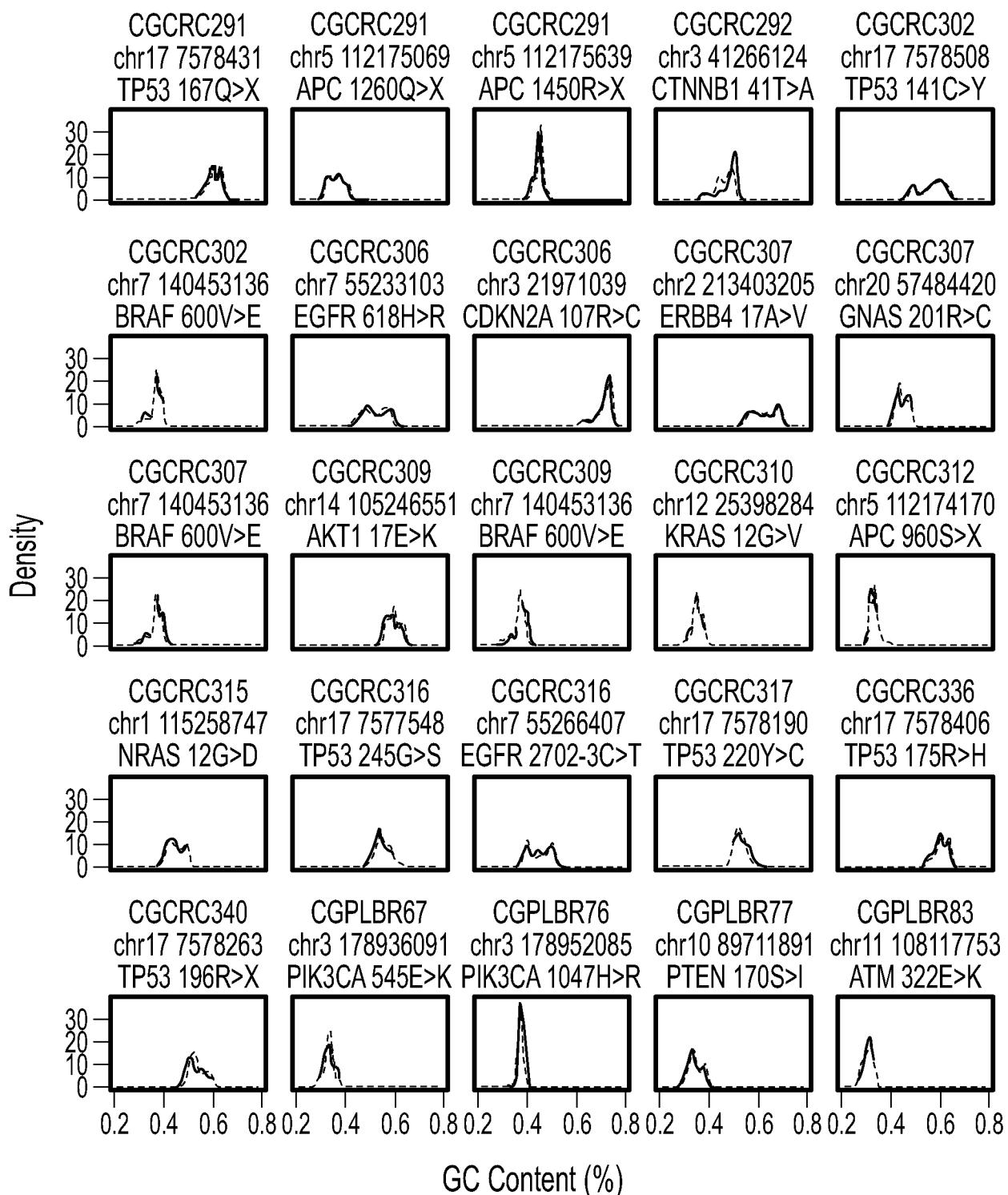


FIG. 4A

6/33

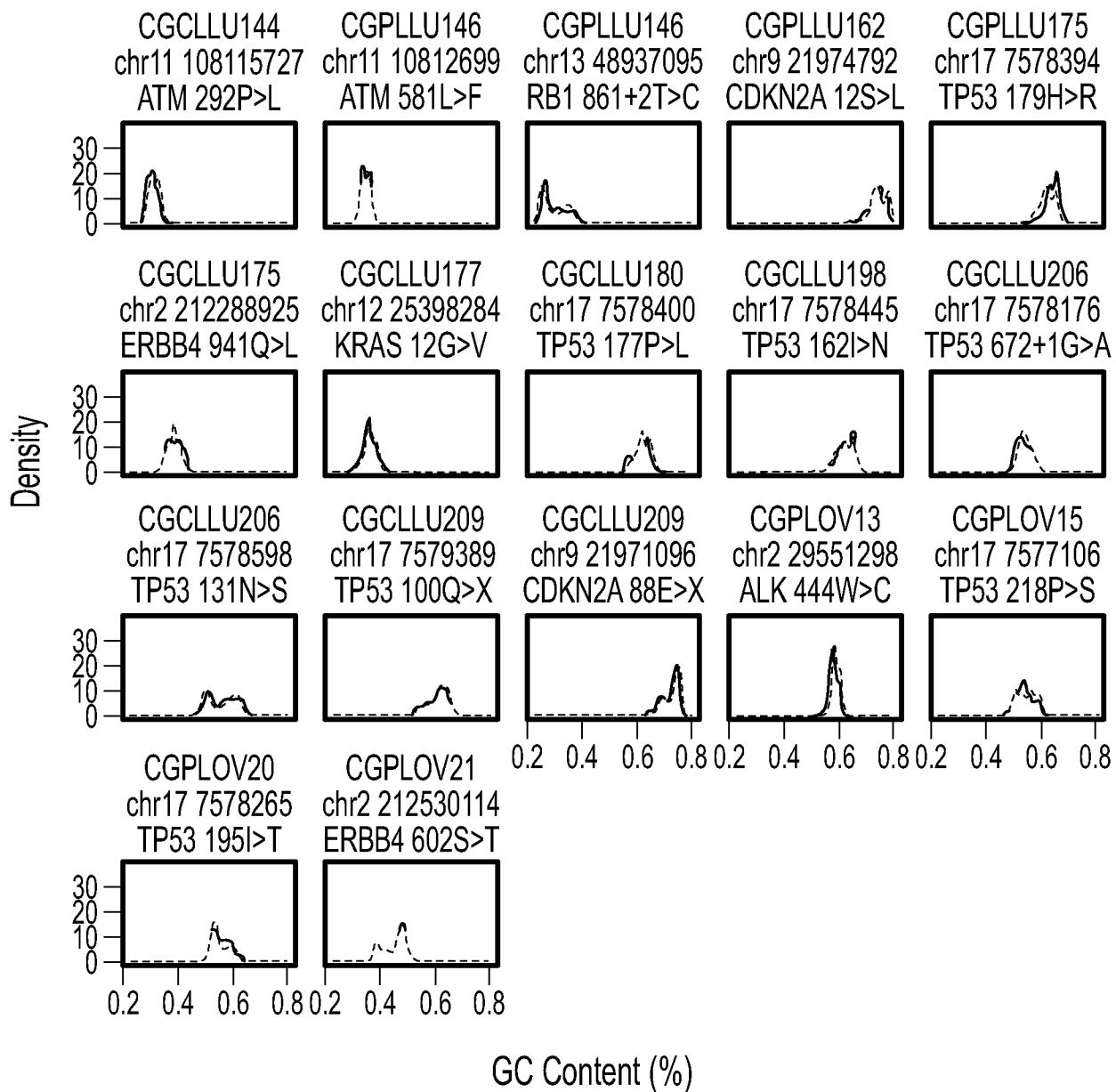


FIG. 4A (Cont.)

7/33

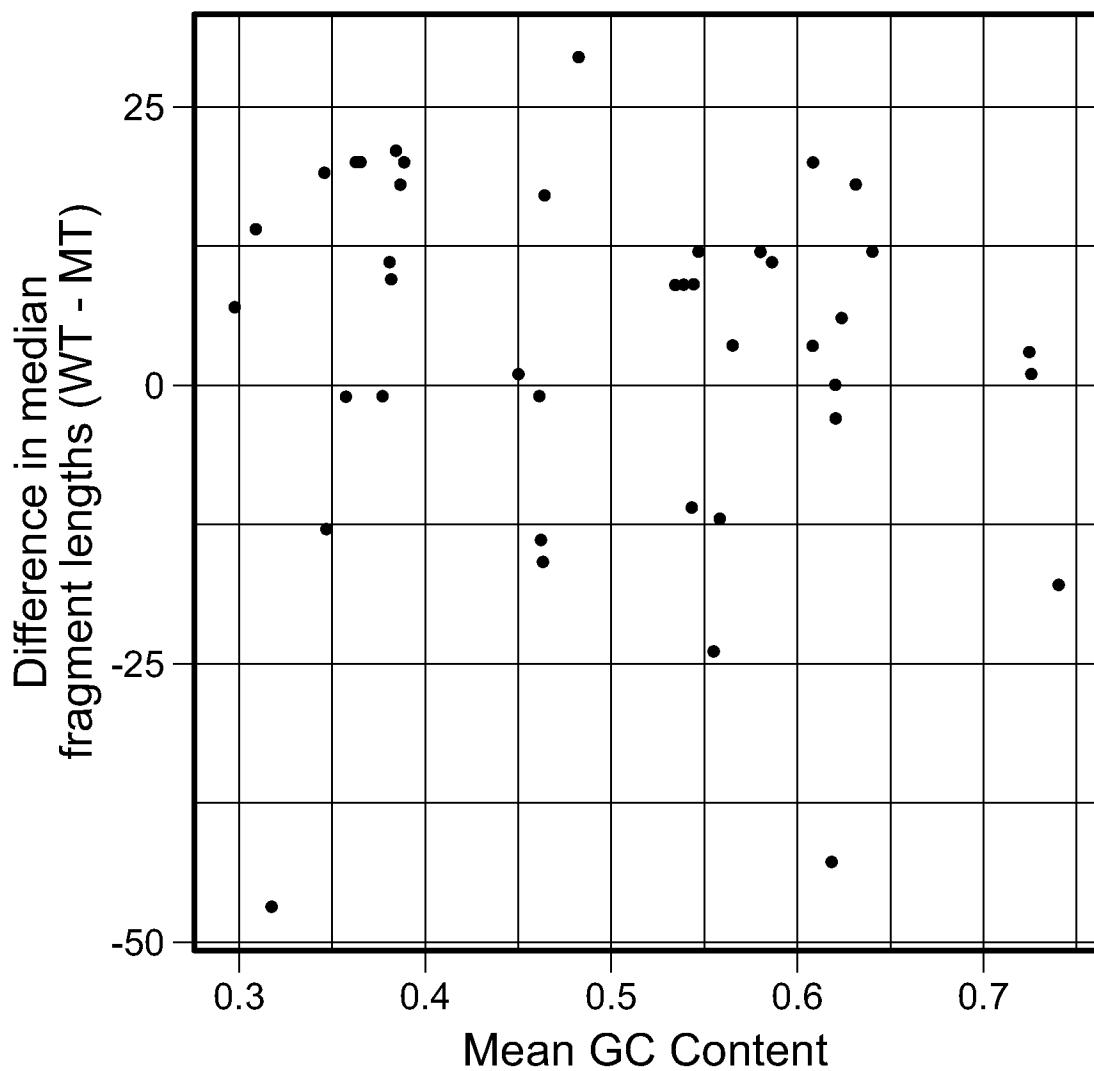


FIG. 4B

8/33

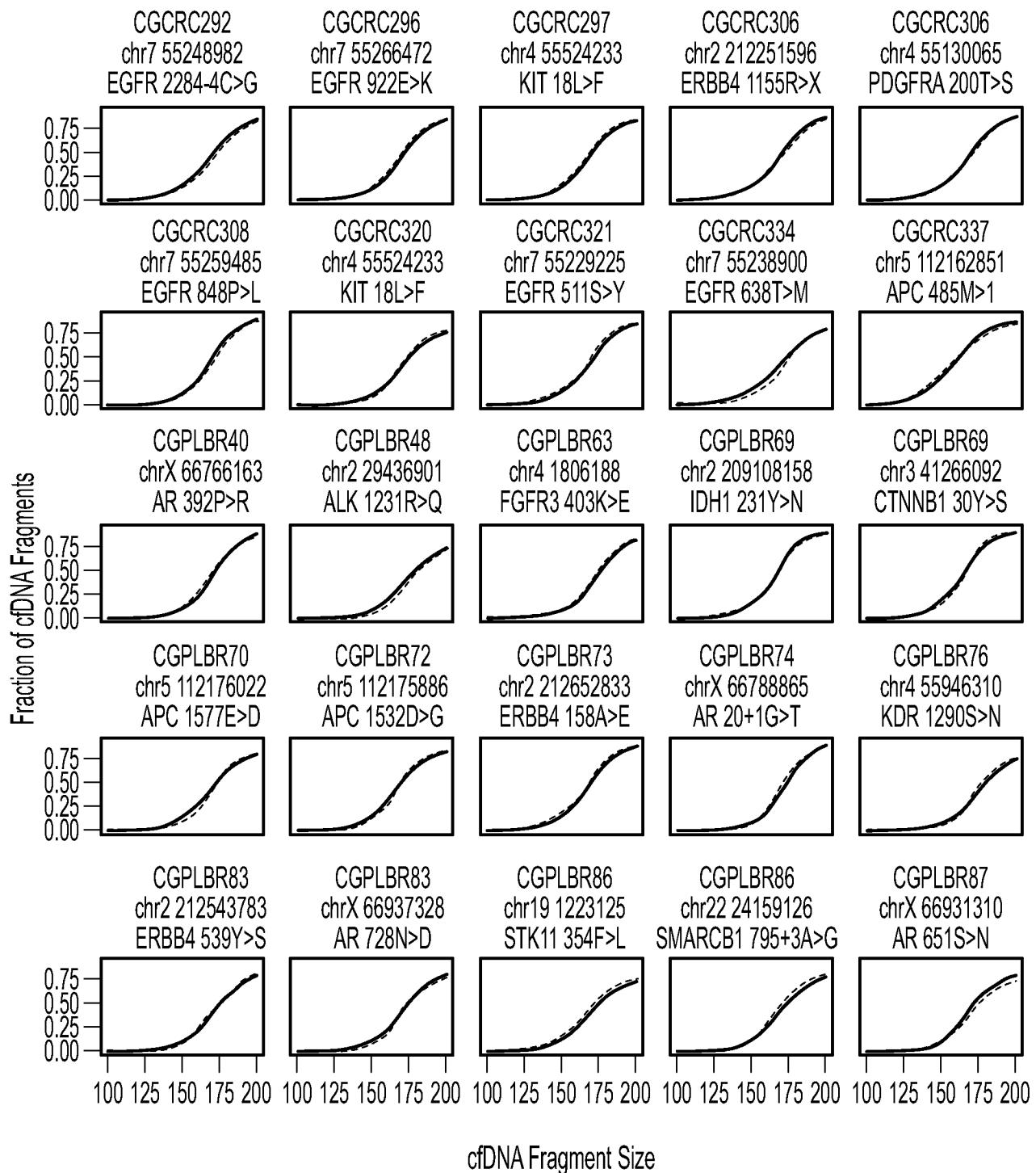


FIG. 5

9/33

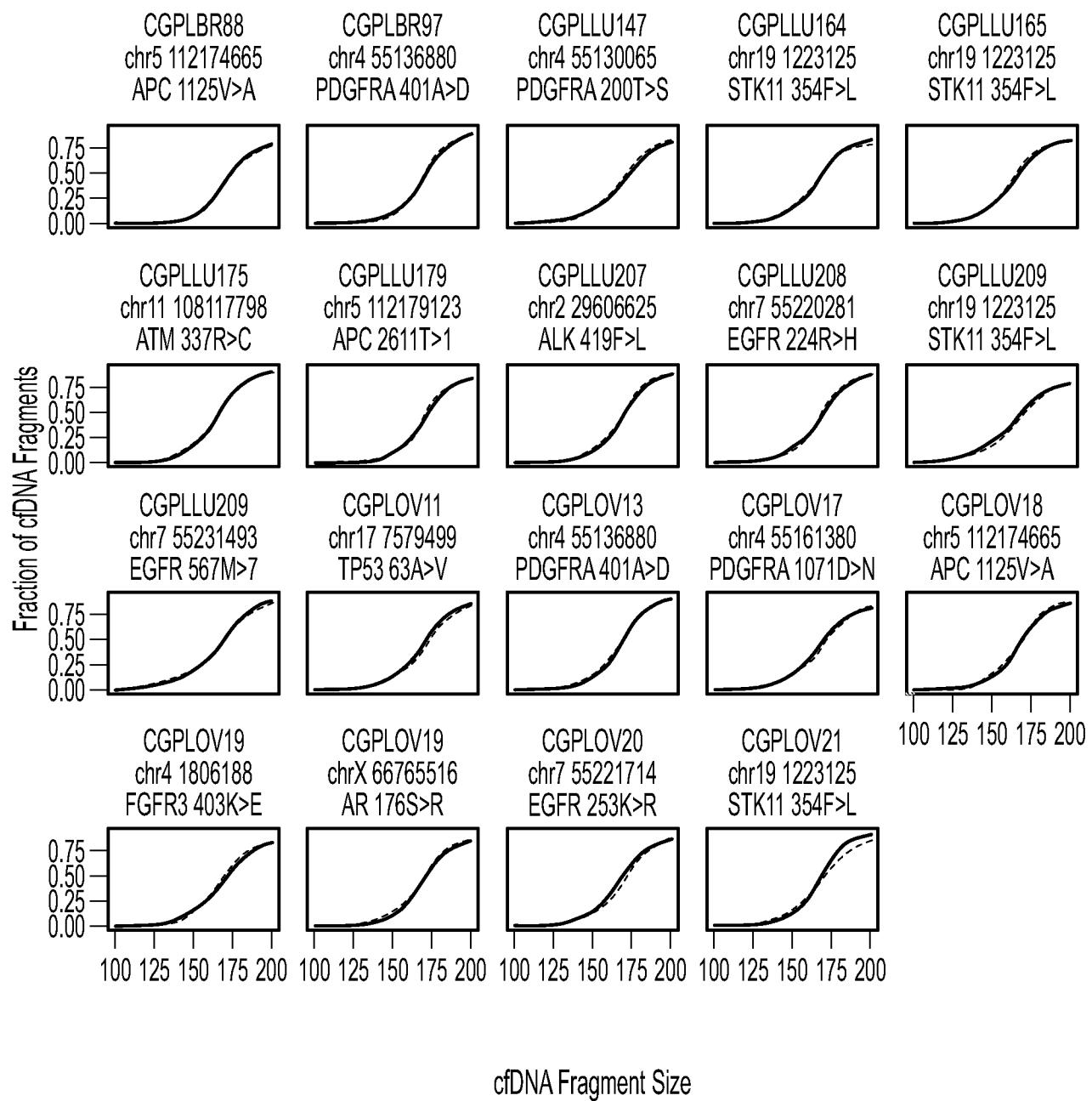
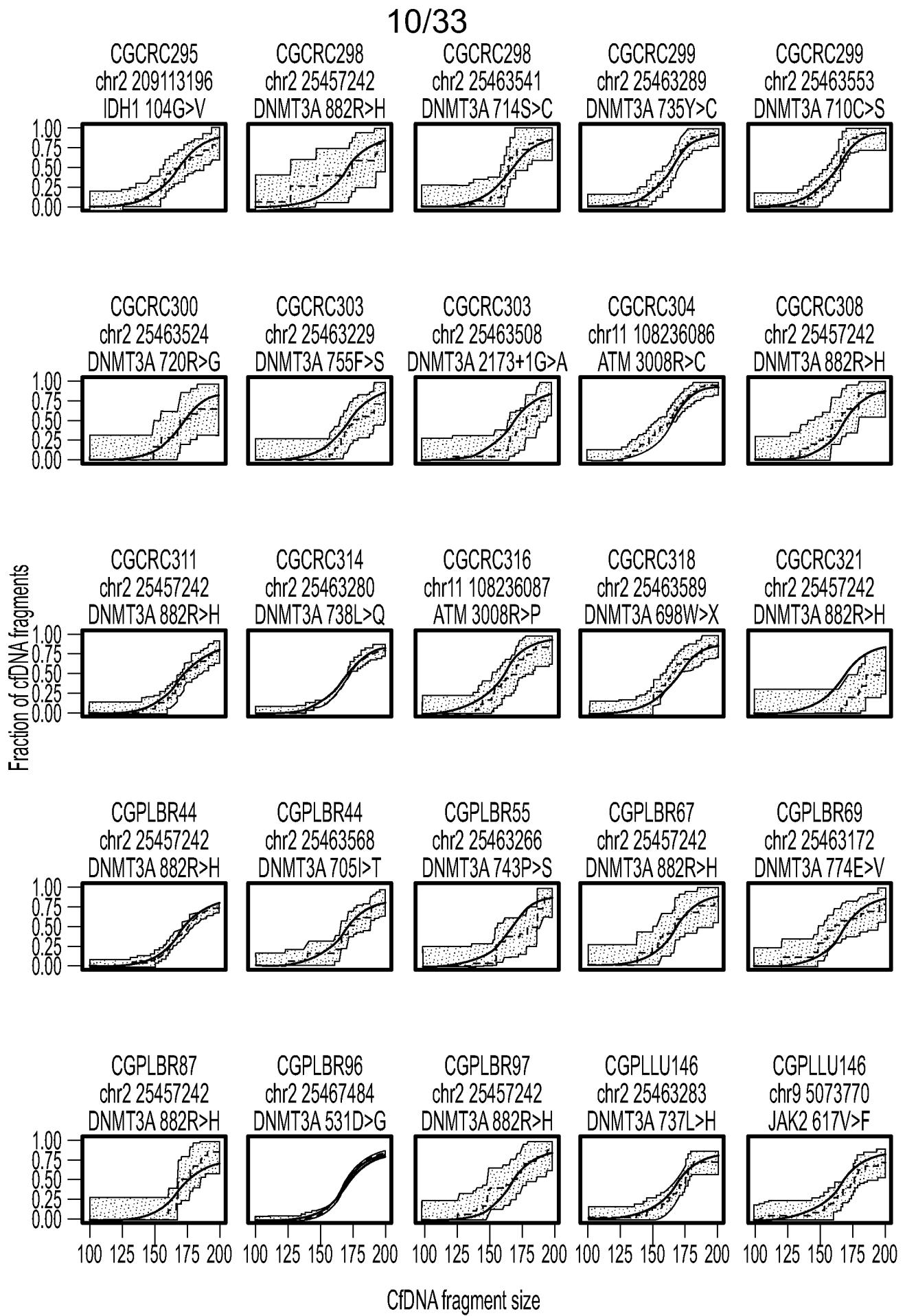


FIG. 5(Cont.)

**FIG. 6**

SUBSTITUTE SHEET (RULE 26)

11/33

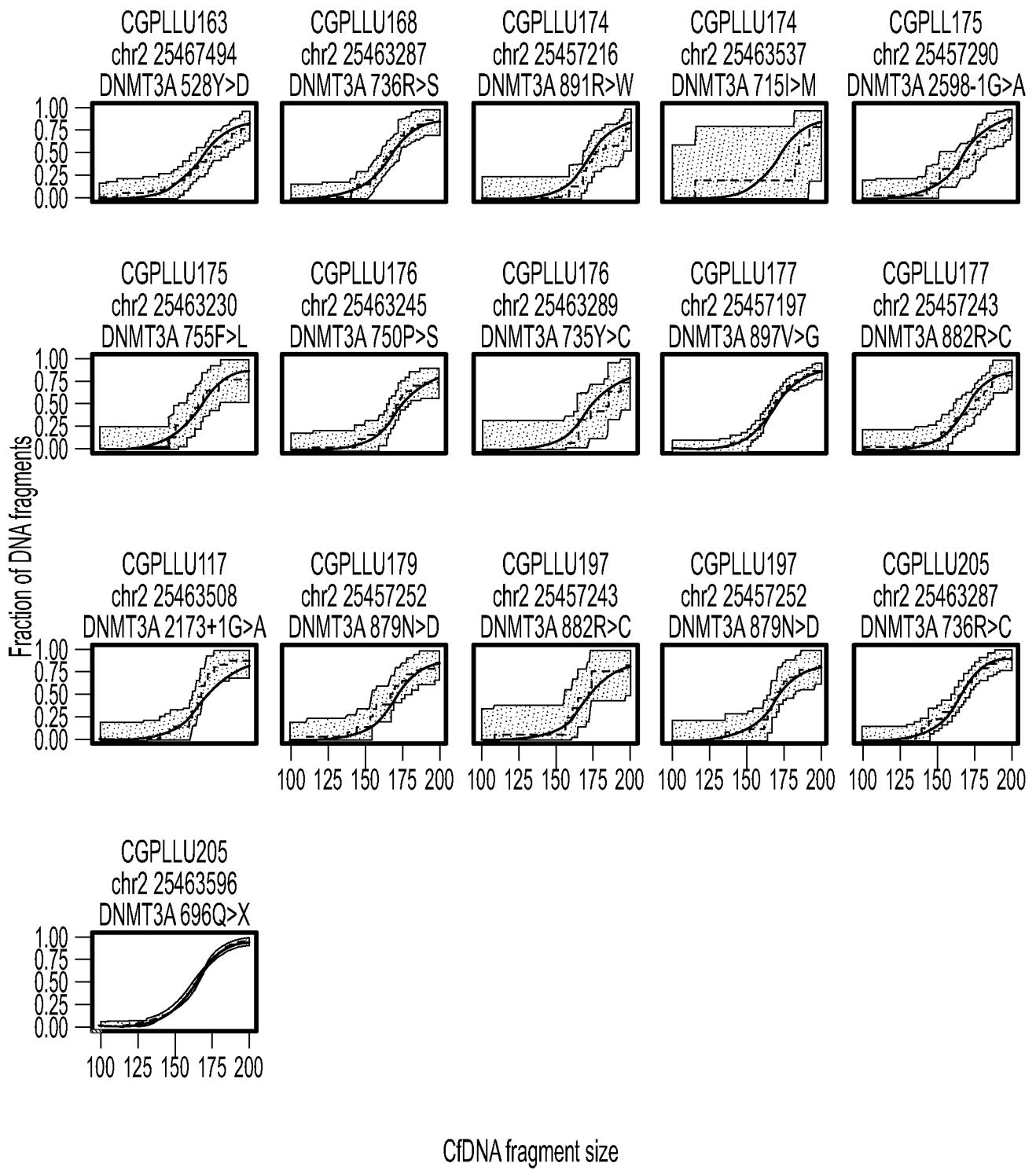
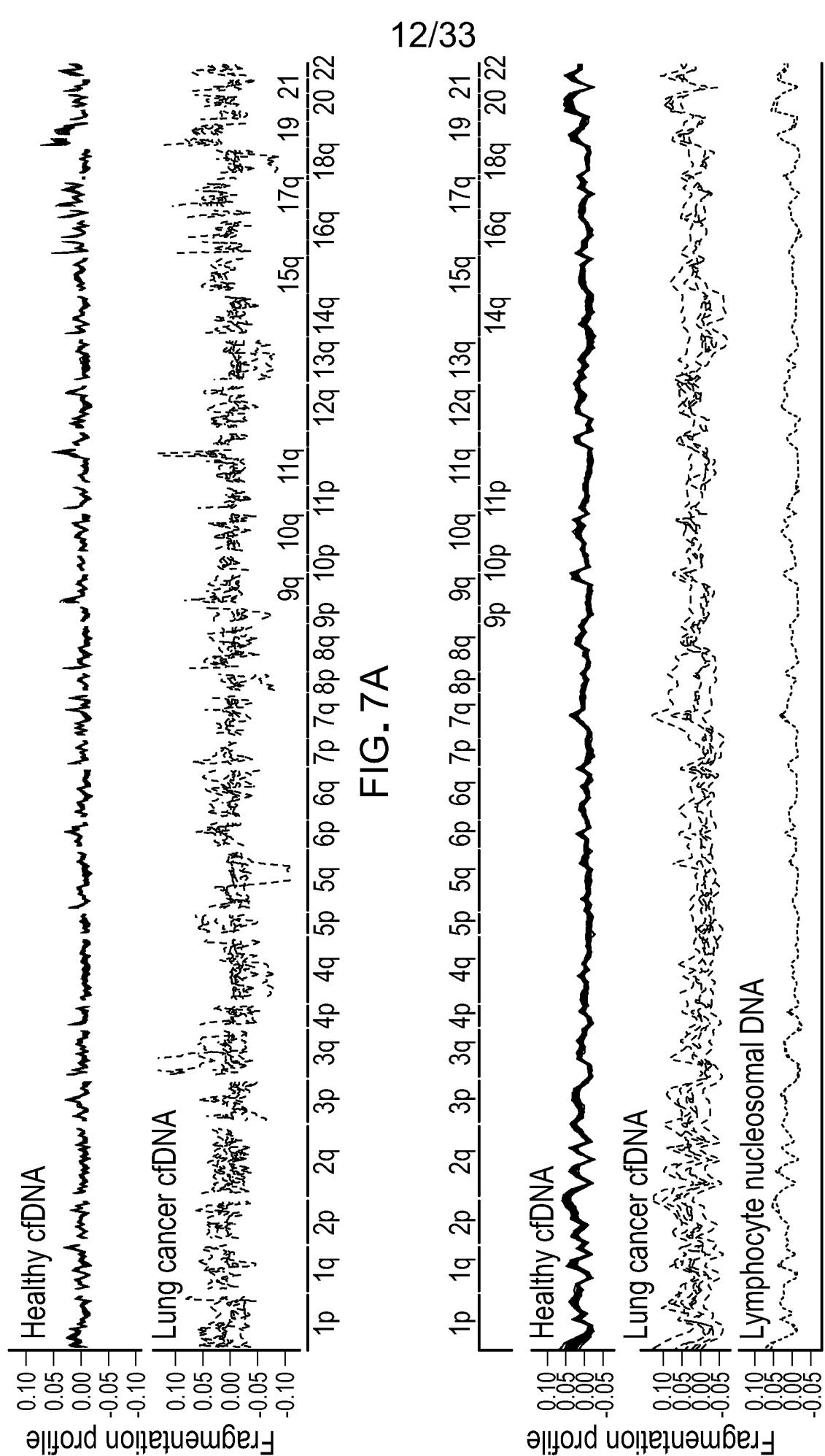
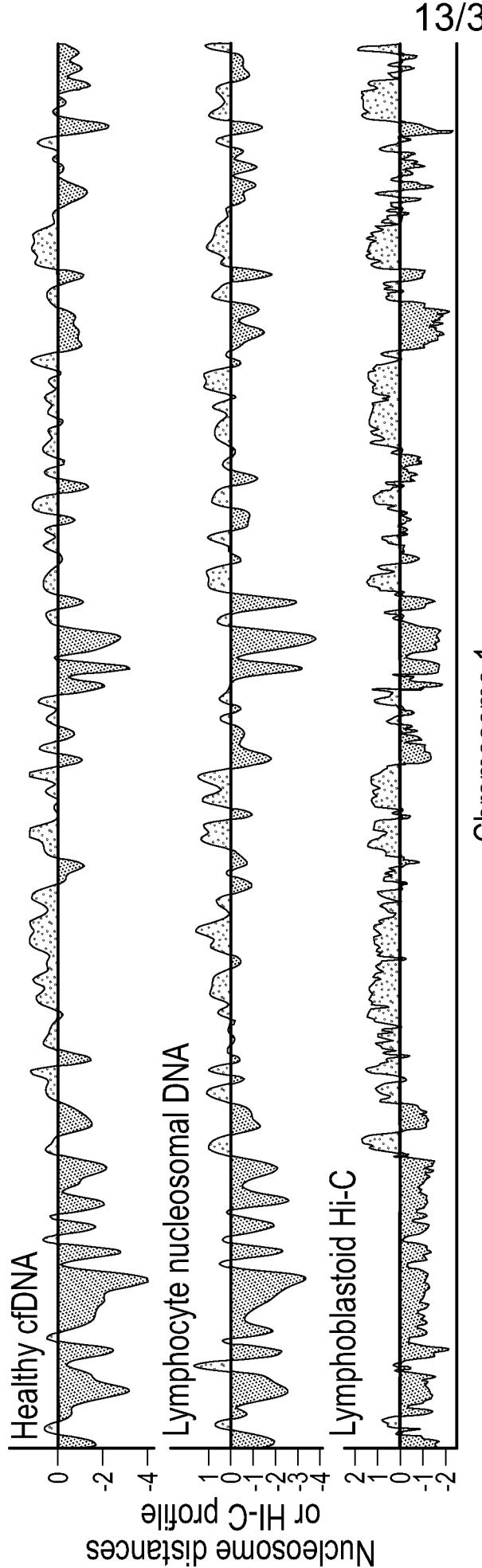


FIG. 6(Cont.)





Chromosome 1
FIG. 7C

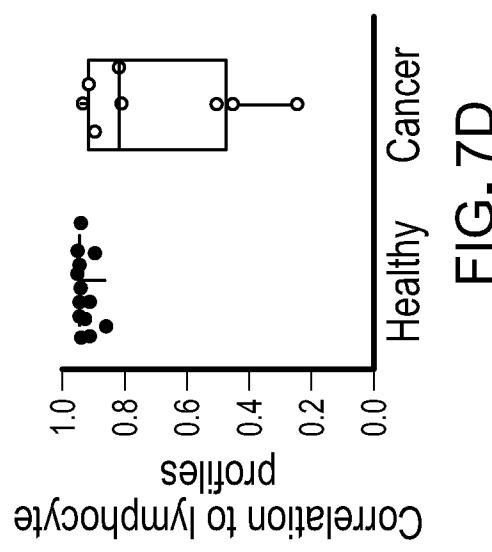
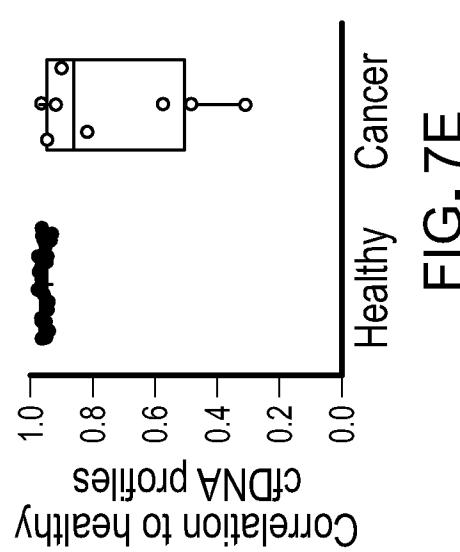
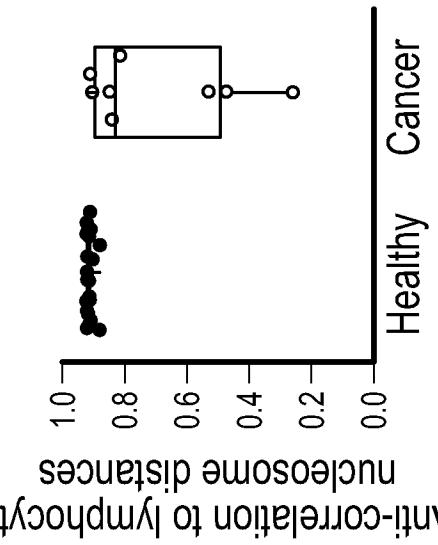


FIG. 7E

FIG. 7D

FIG. 7F

14/33

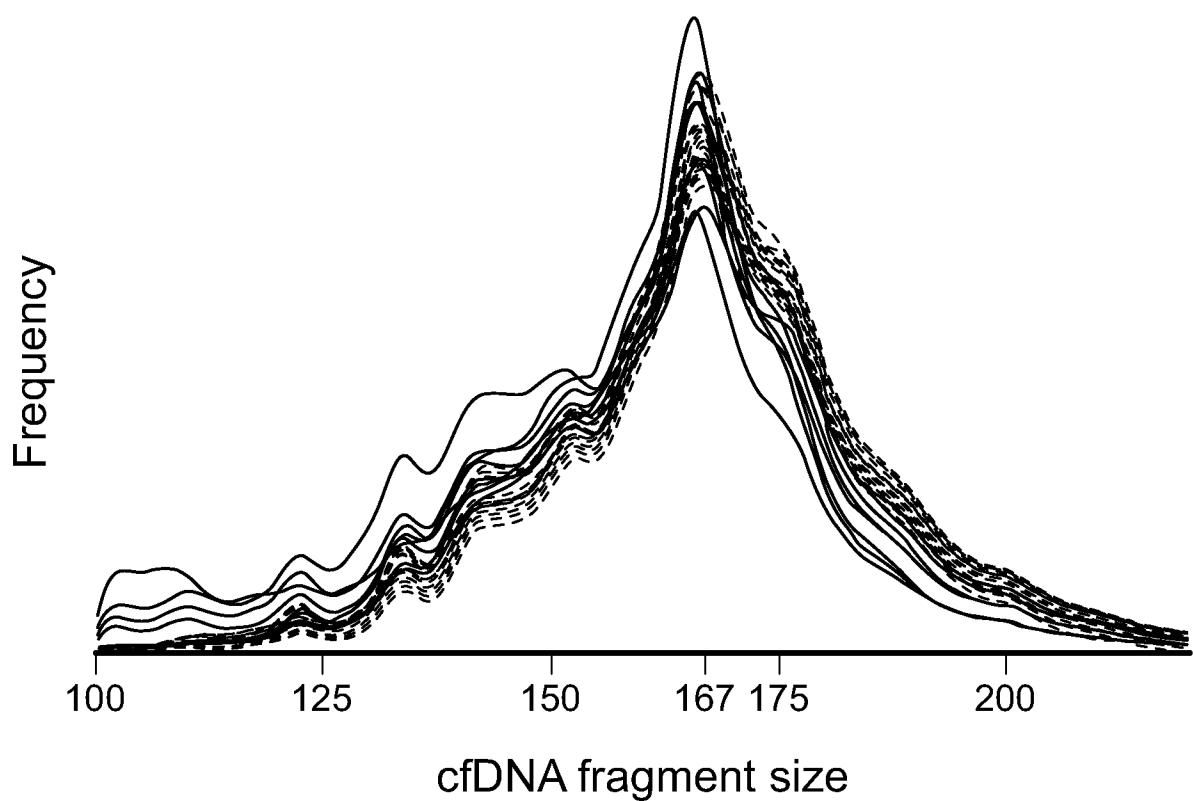
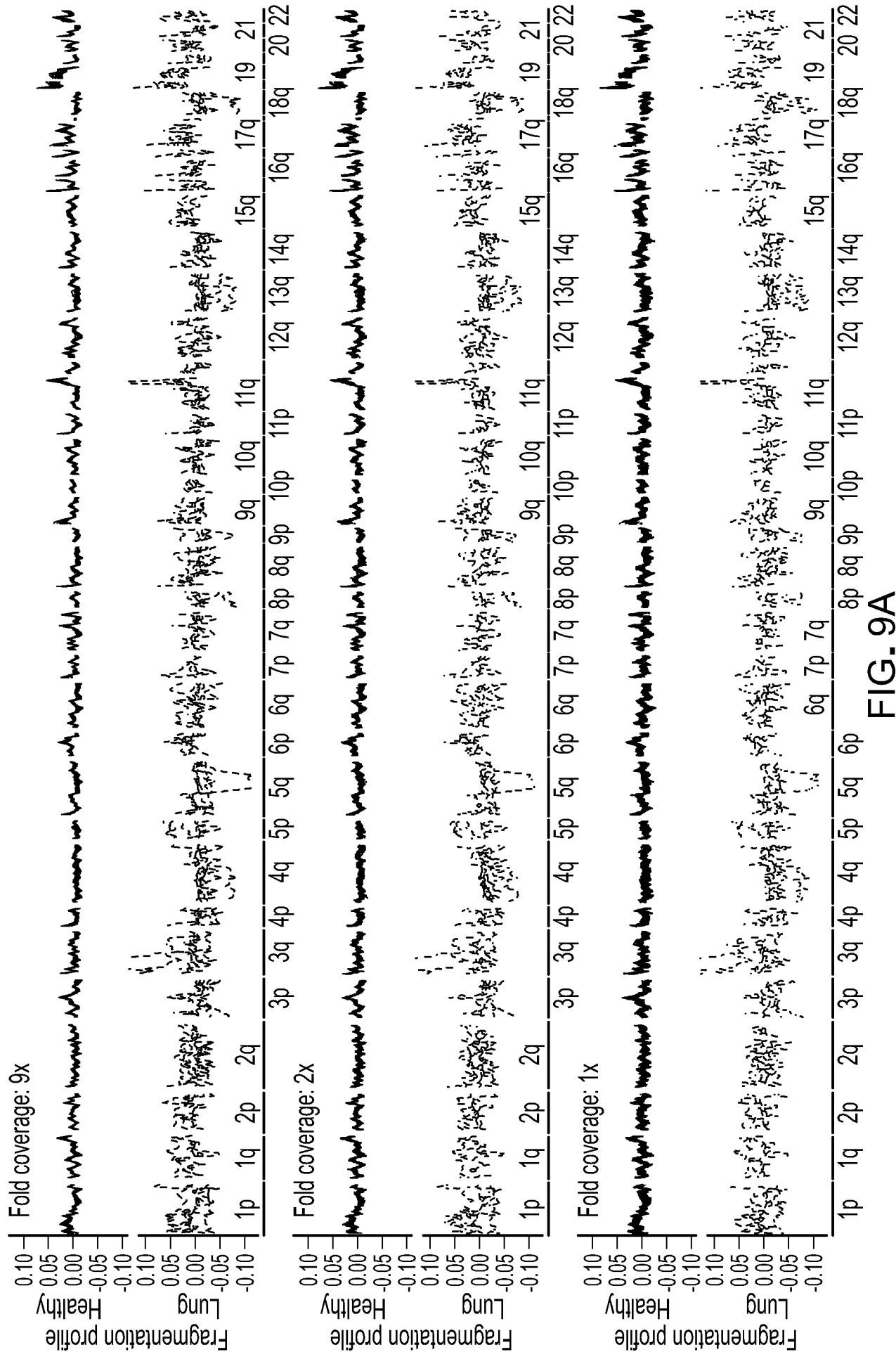
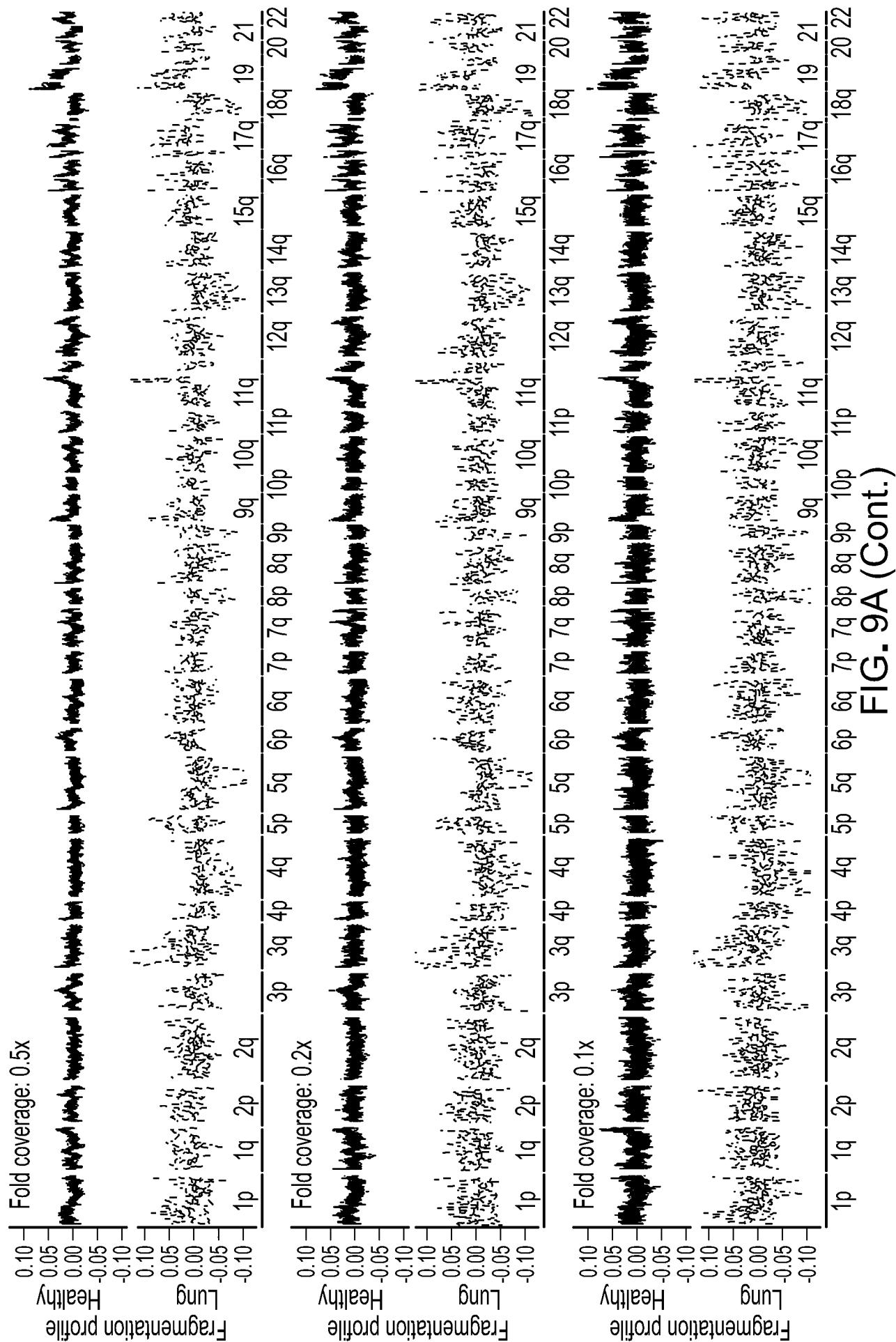


FIG. 8

15/33

**FIG. 9A**

16/33

**FIG. 9A (Cont.)**

17/33

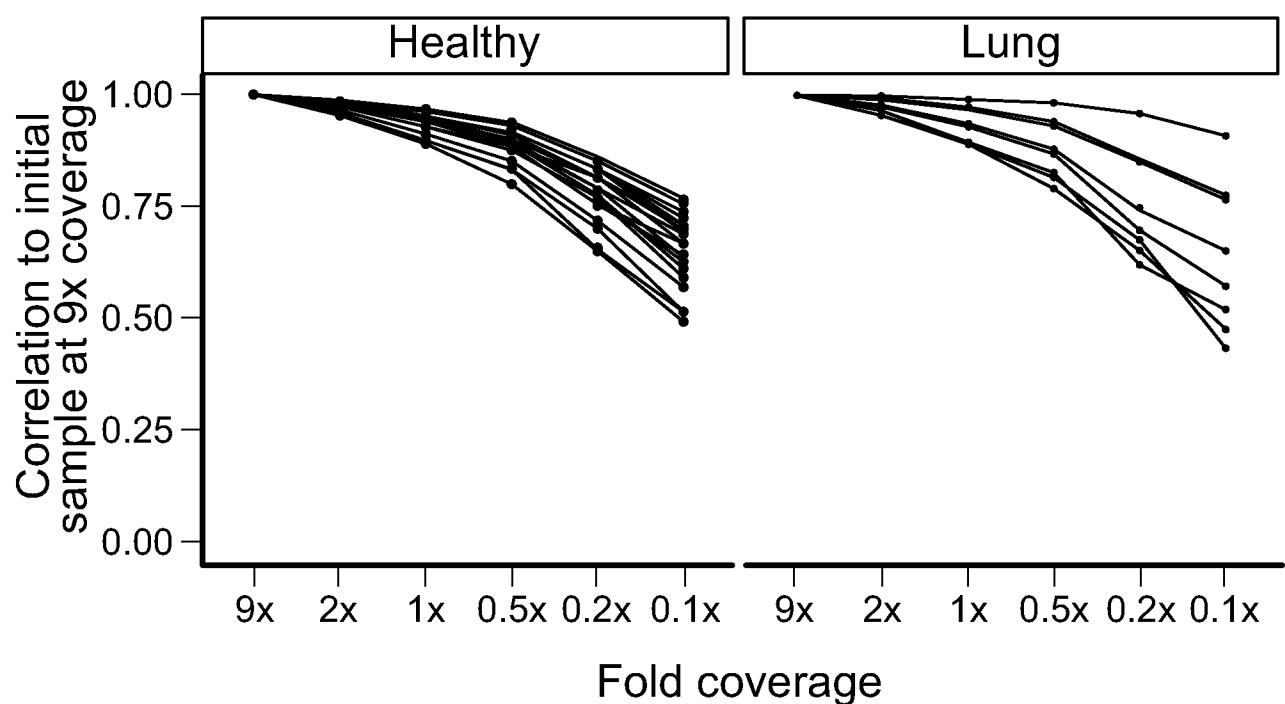


FIG. 9B

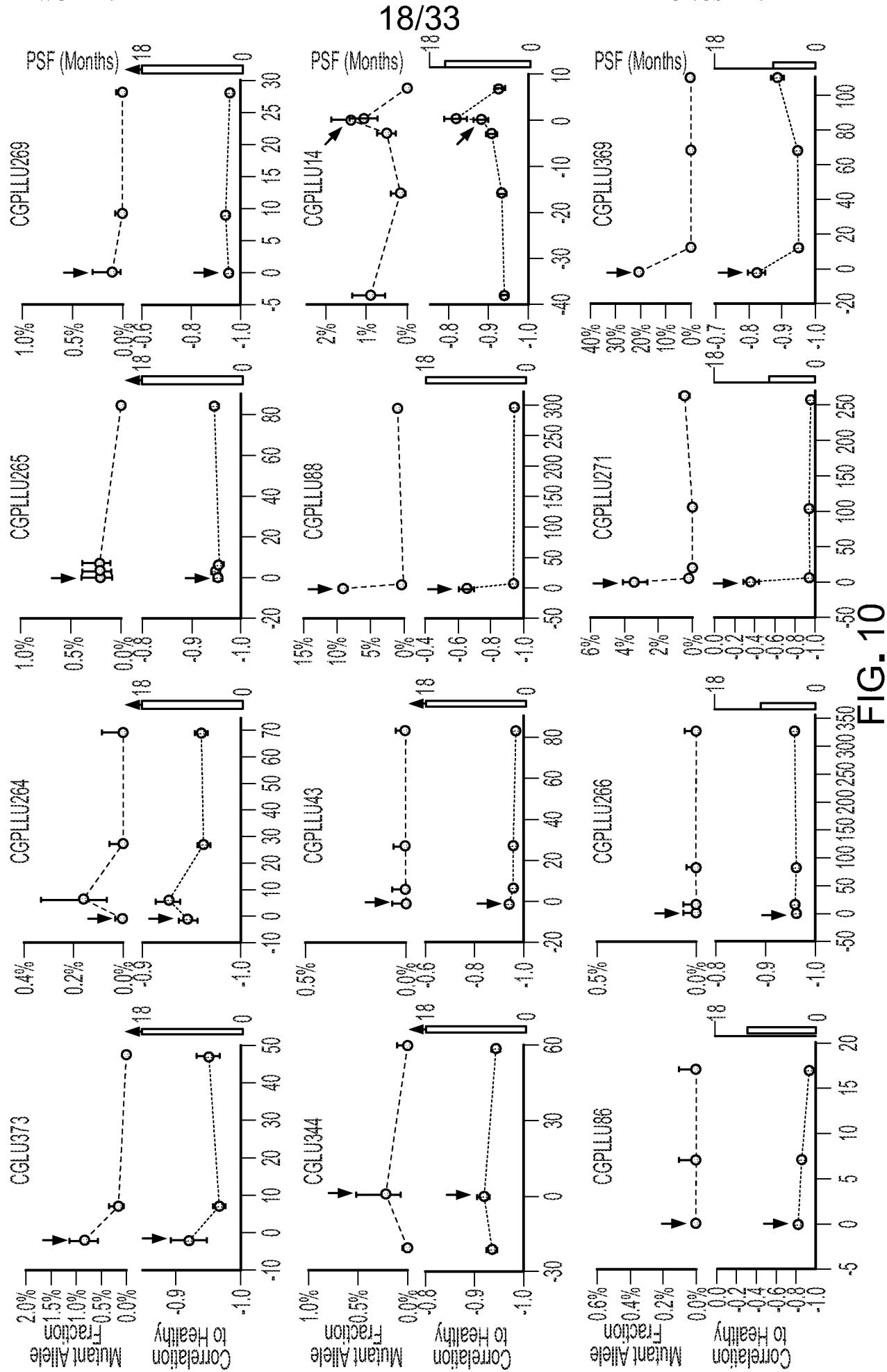
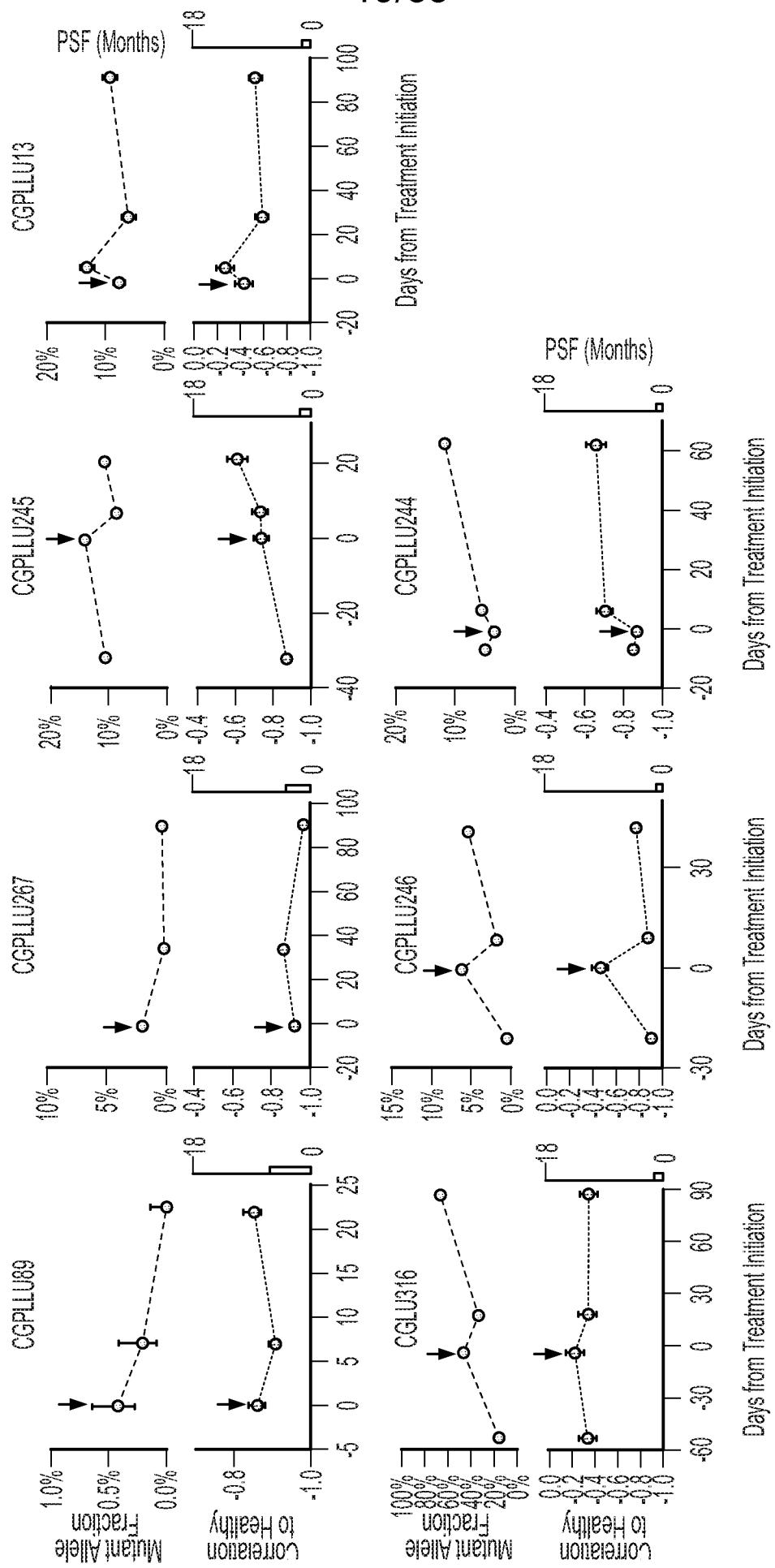


FIG. 10

19/33



SUBSTITUTE SHEET (RULE 26)

FIG. 10(Cont.)

20/33

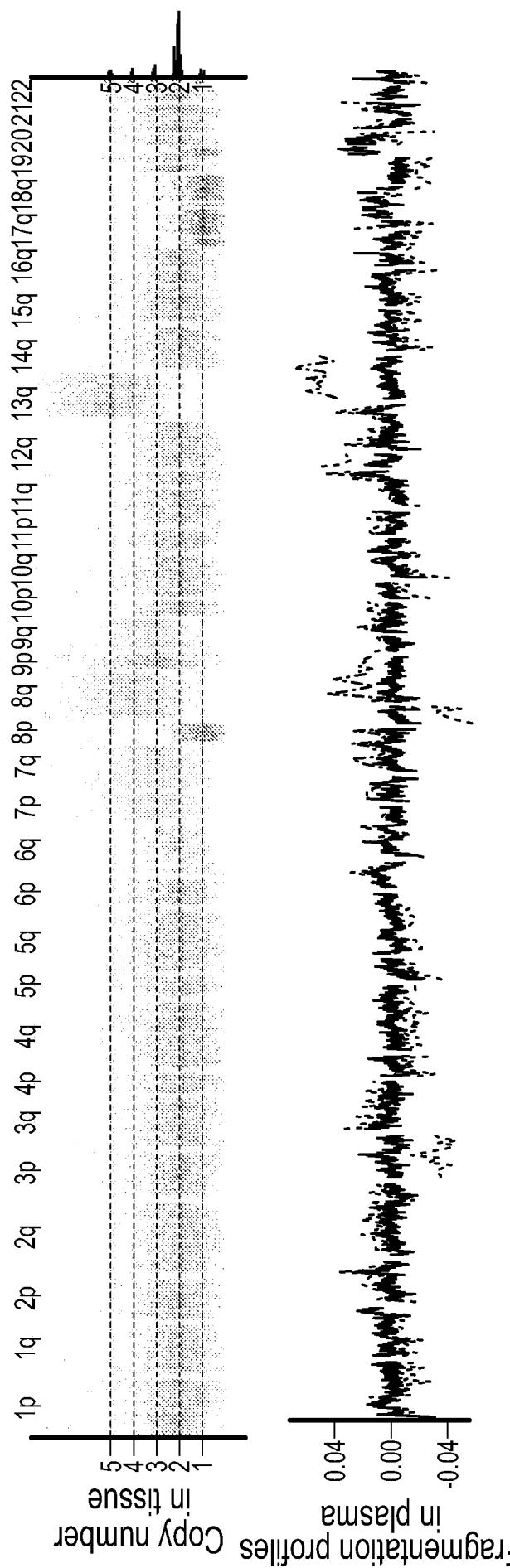


FIG. 11A

21/33

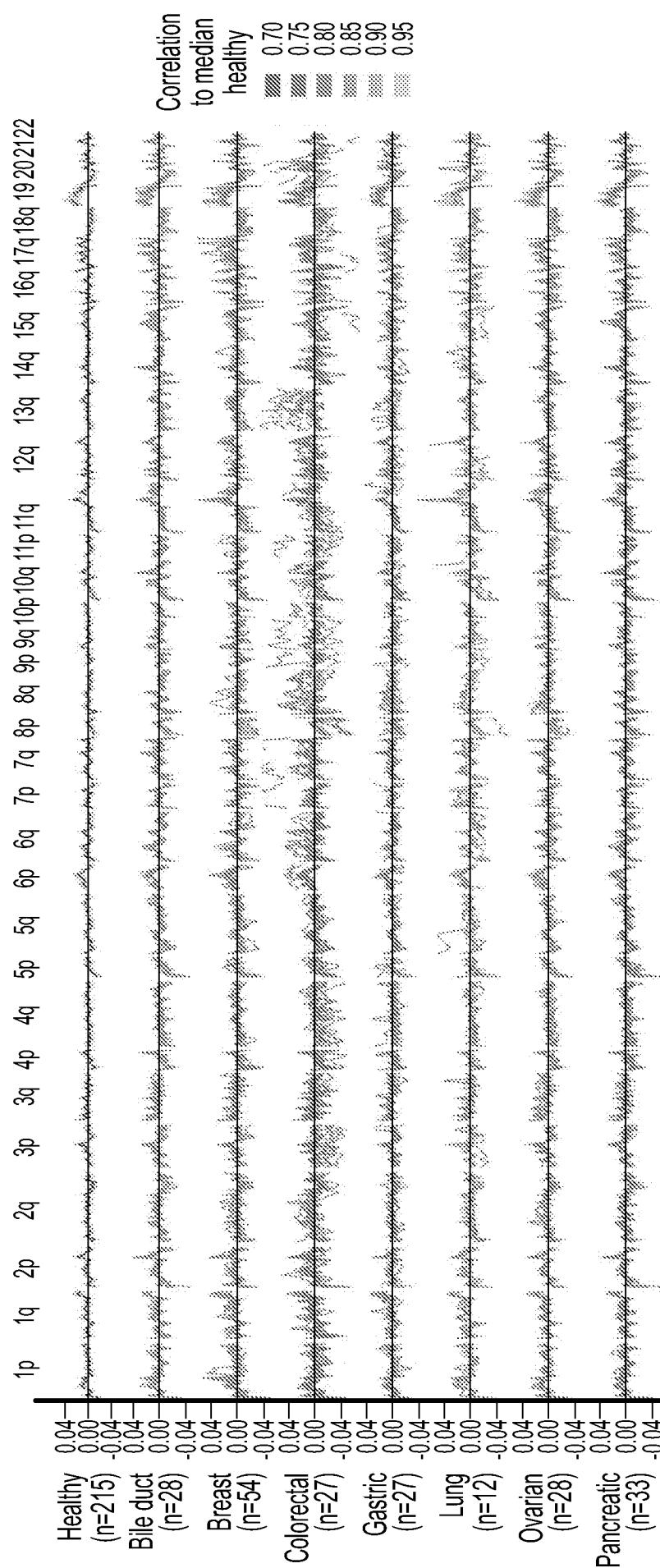


FIG. 11B

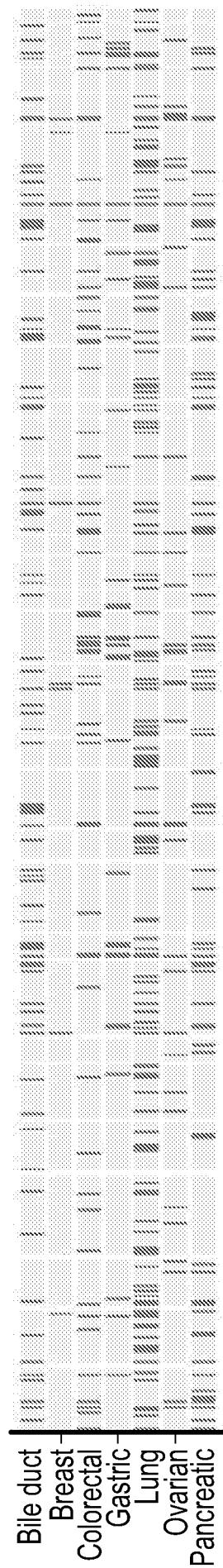


FIG. 11C

22/33

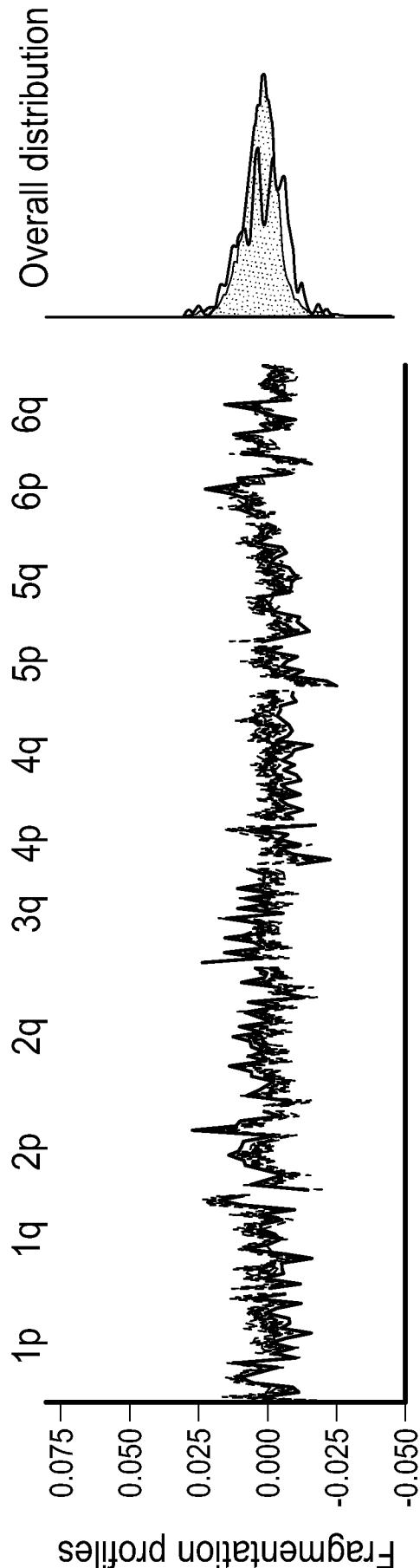


FIG. 12A

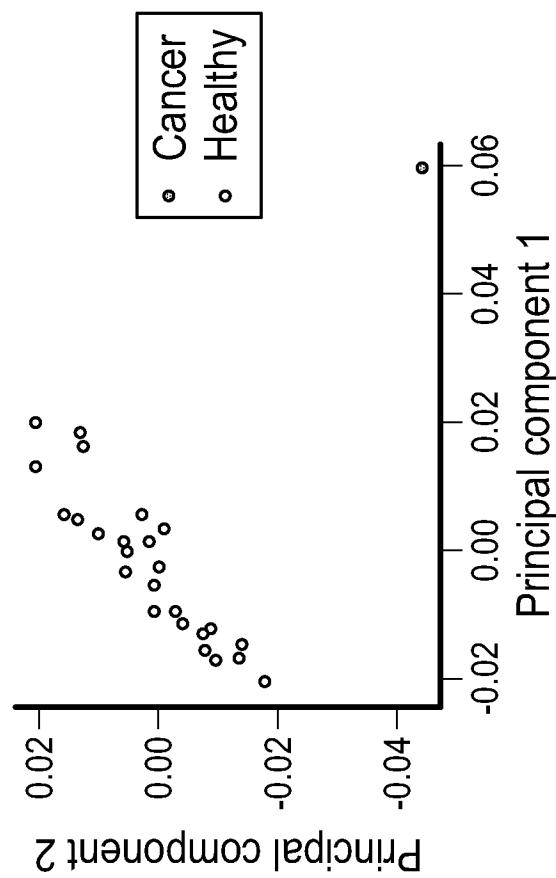


FIG. 12B

23/33

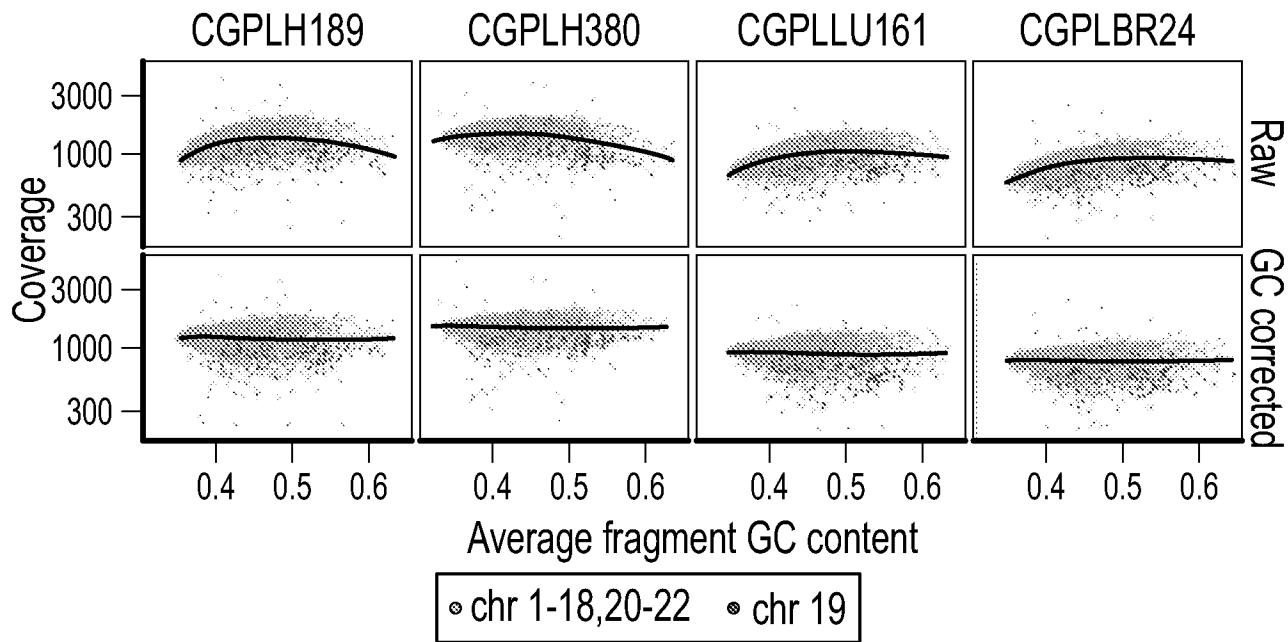


FIG. 13A

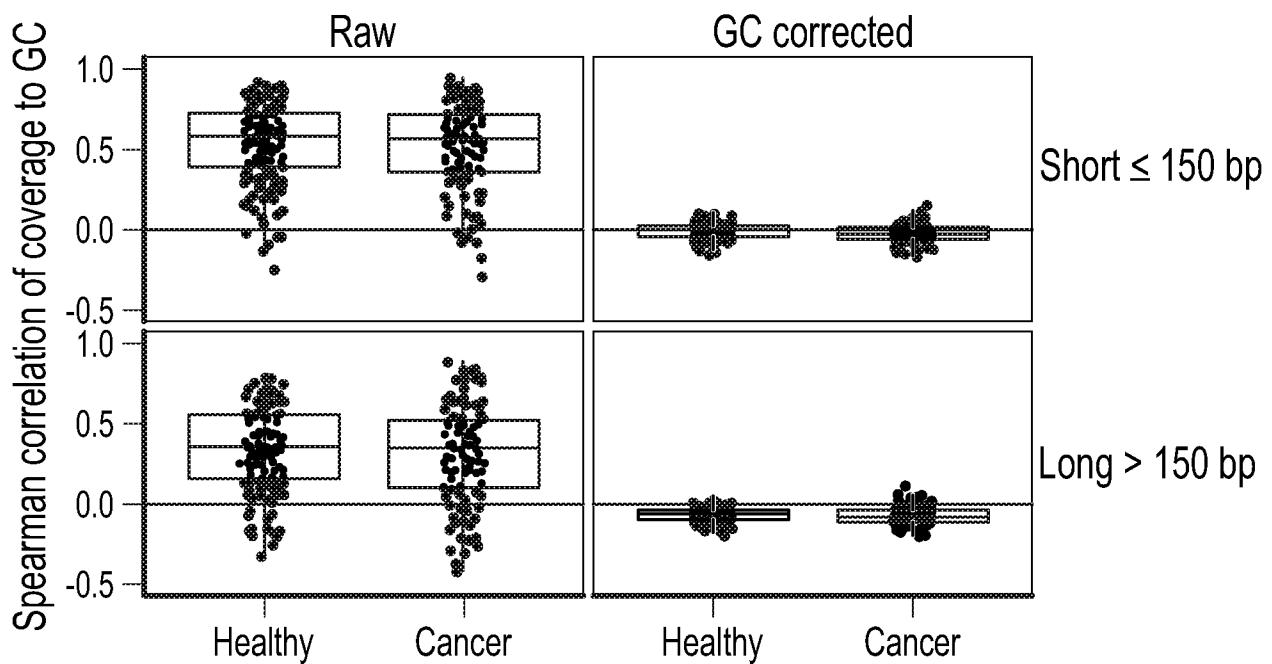


FIG. 13B

24/33

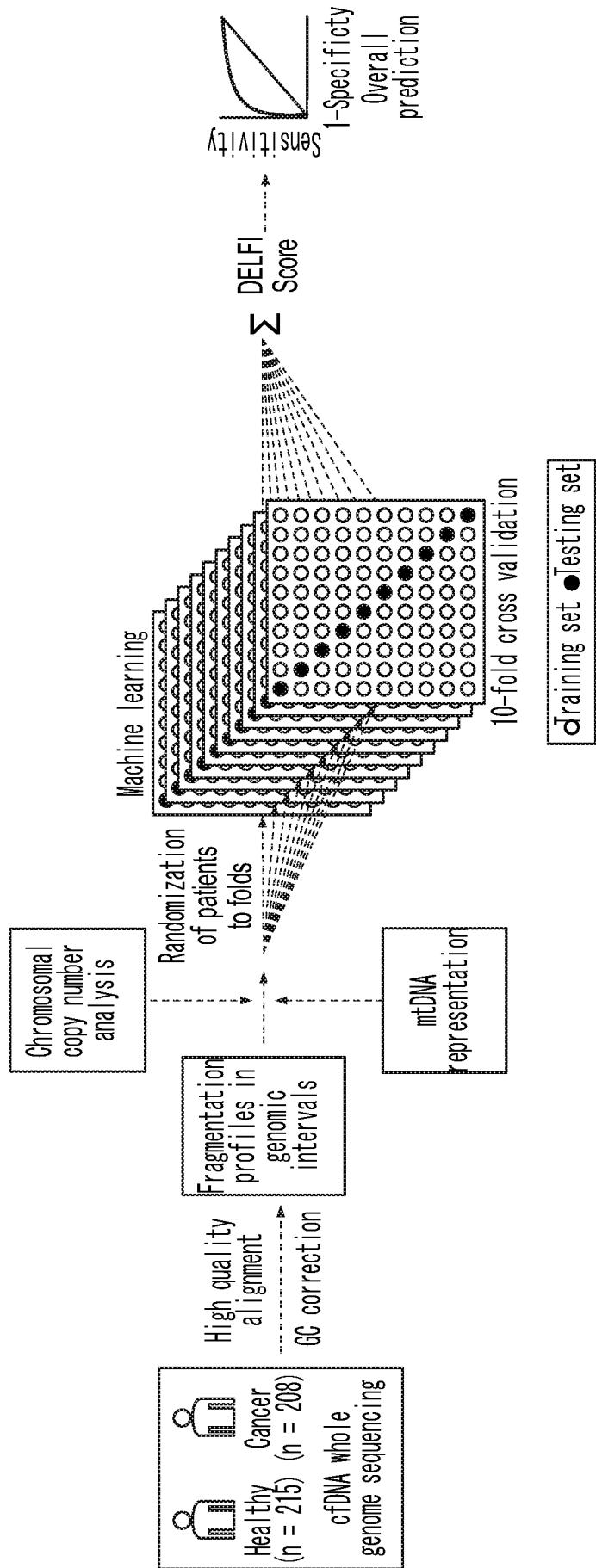


FIG. 14

25/33

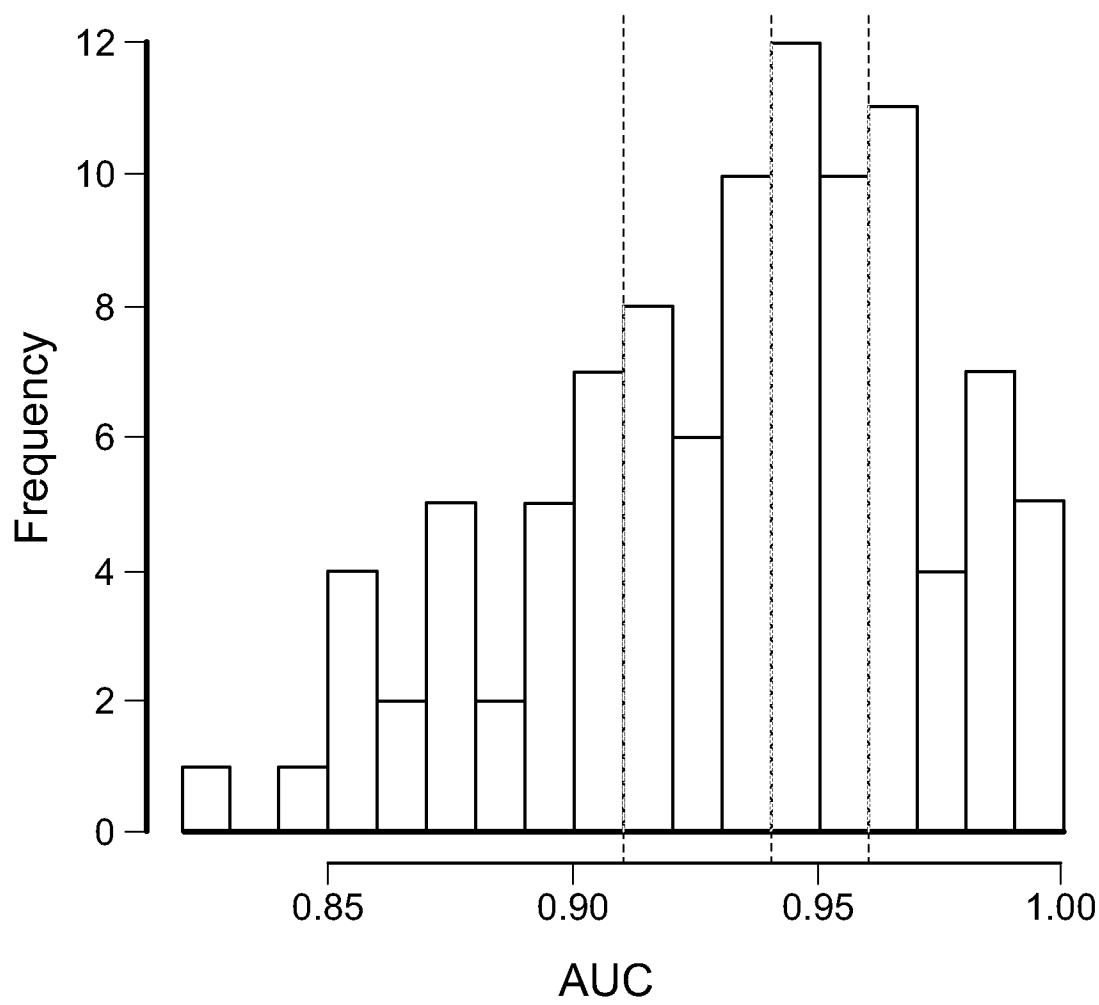


FIG. 15

26/33

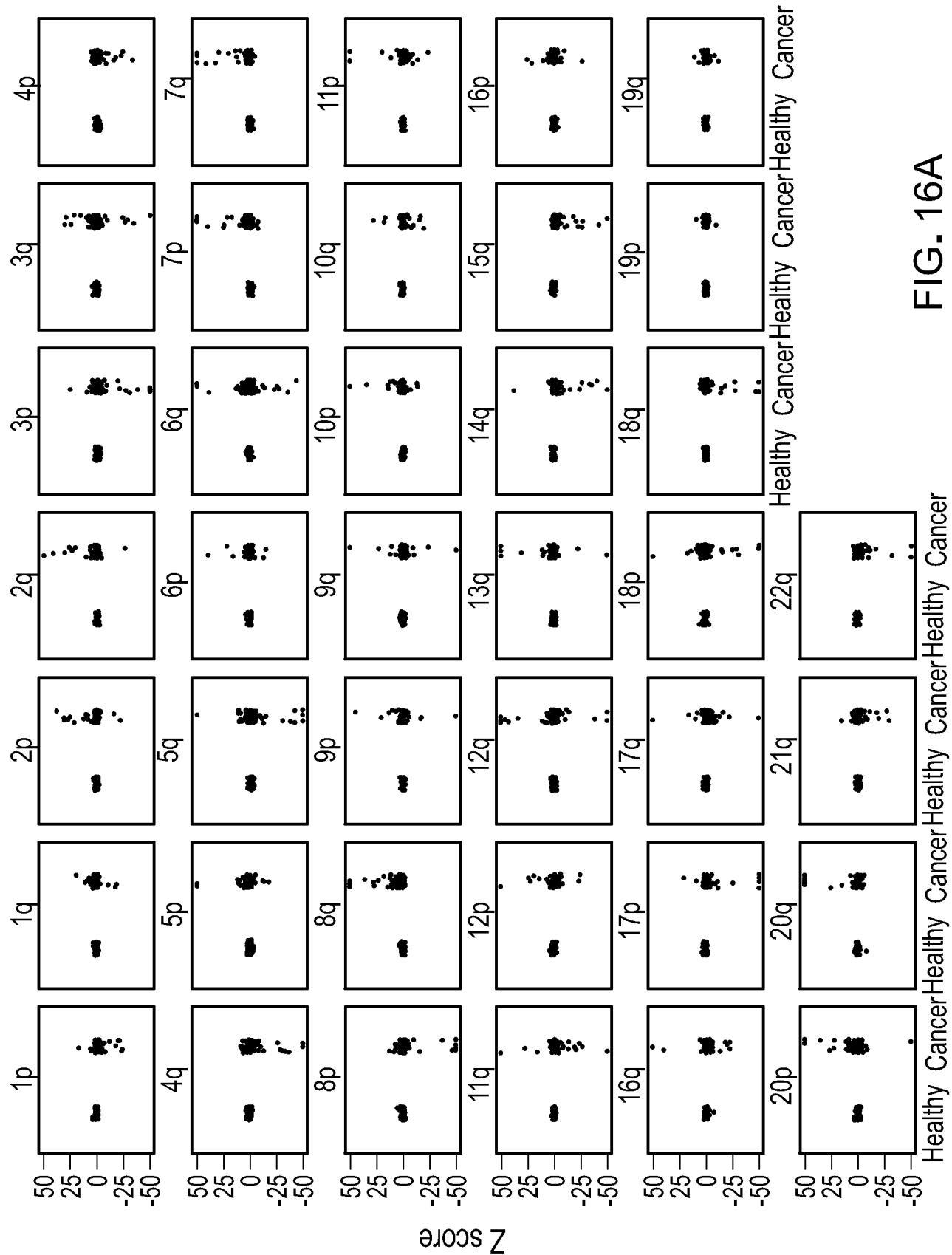


FIG. 16A

Healthy Cancer Healthy Cancer Healthy Cancer Healthy Cancer

27/33

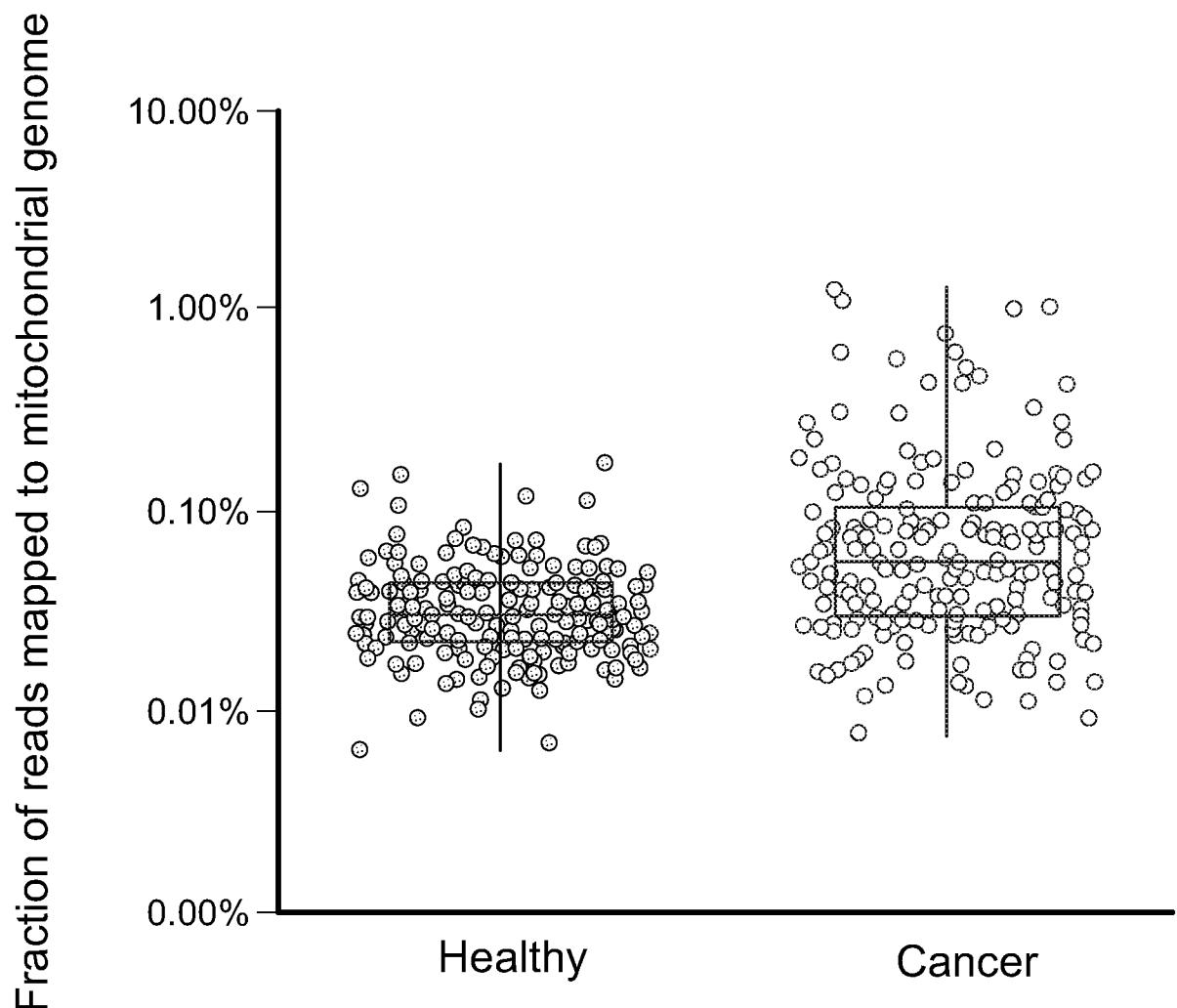


FIG. 16B

28/33

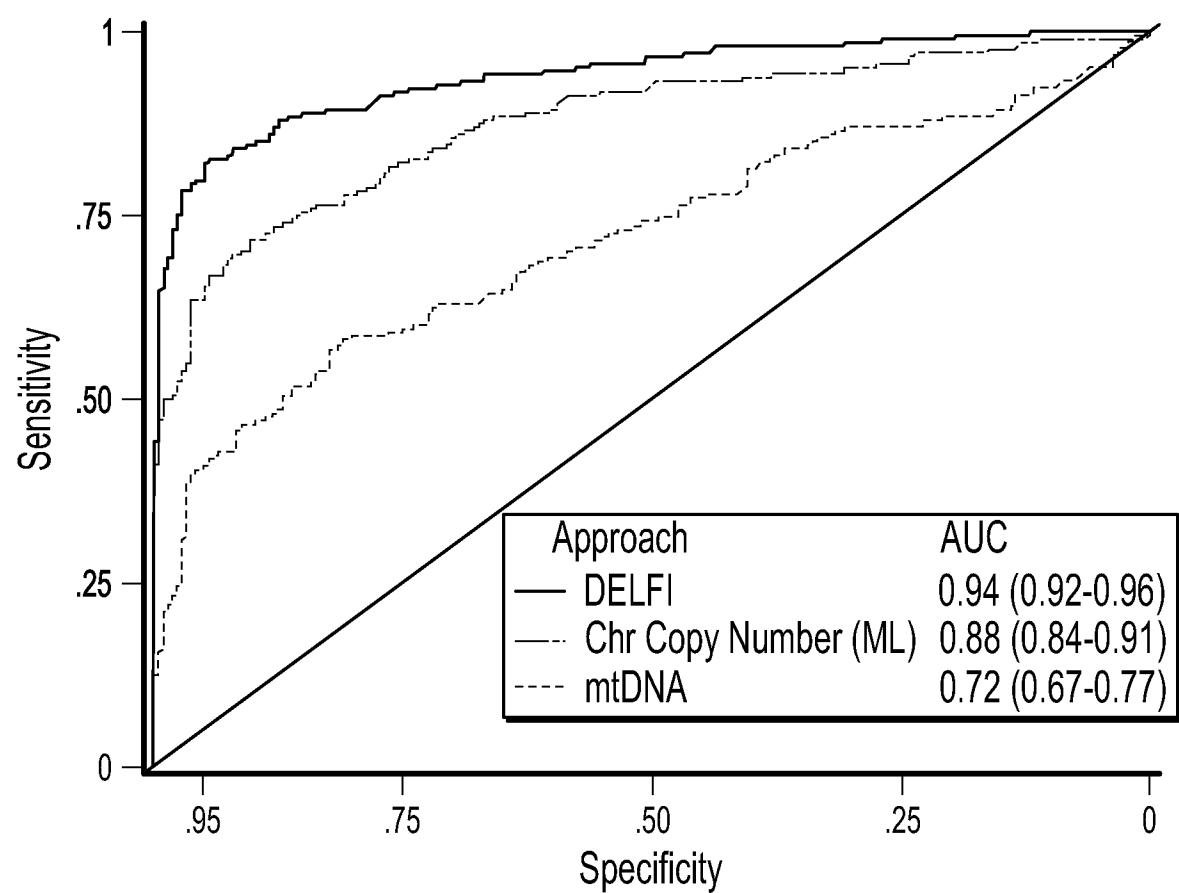
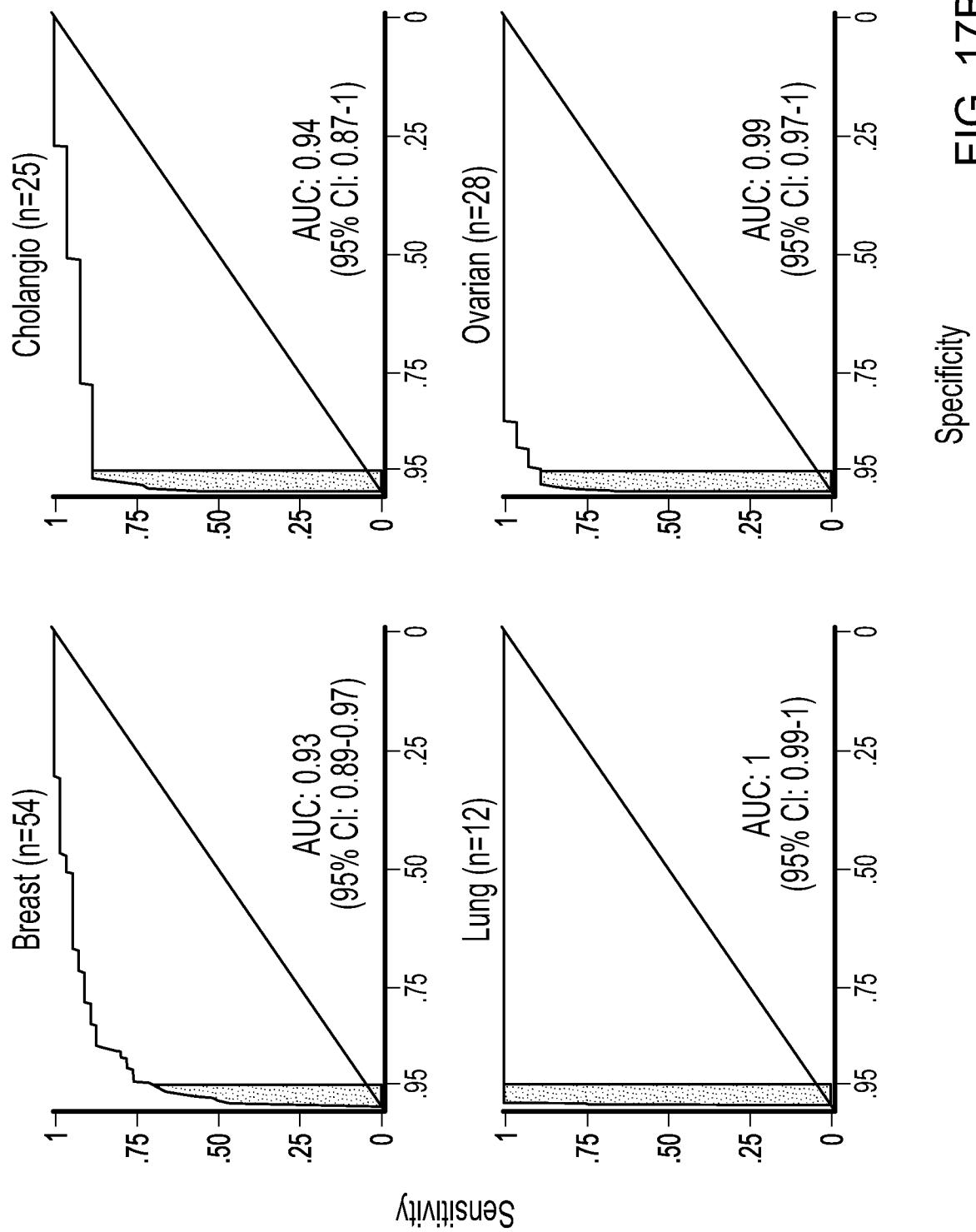


FIG. 17A

29/33



30/33

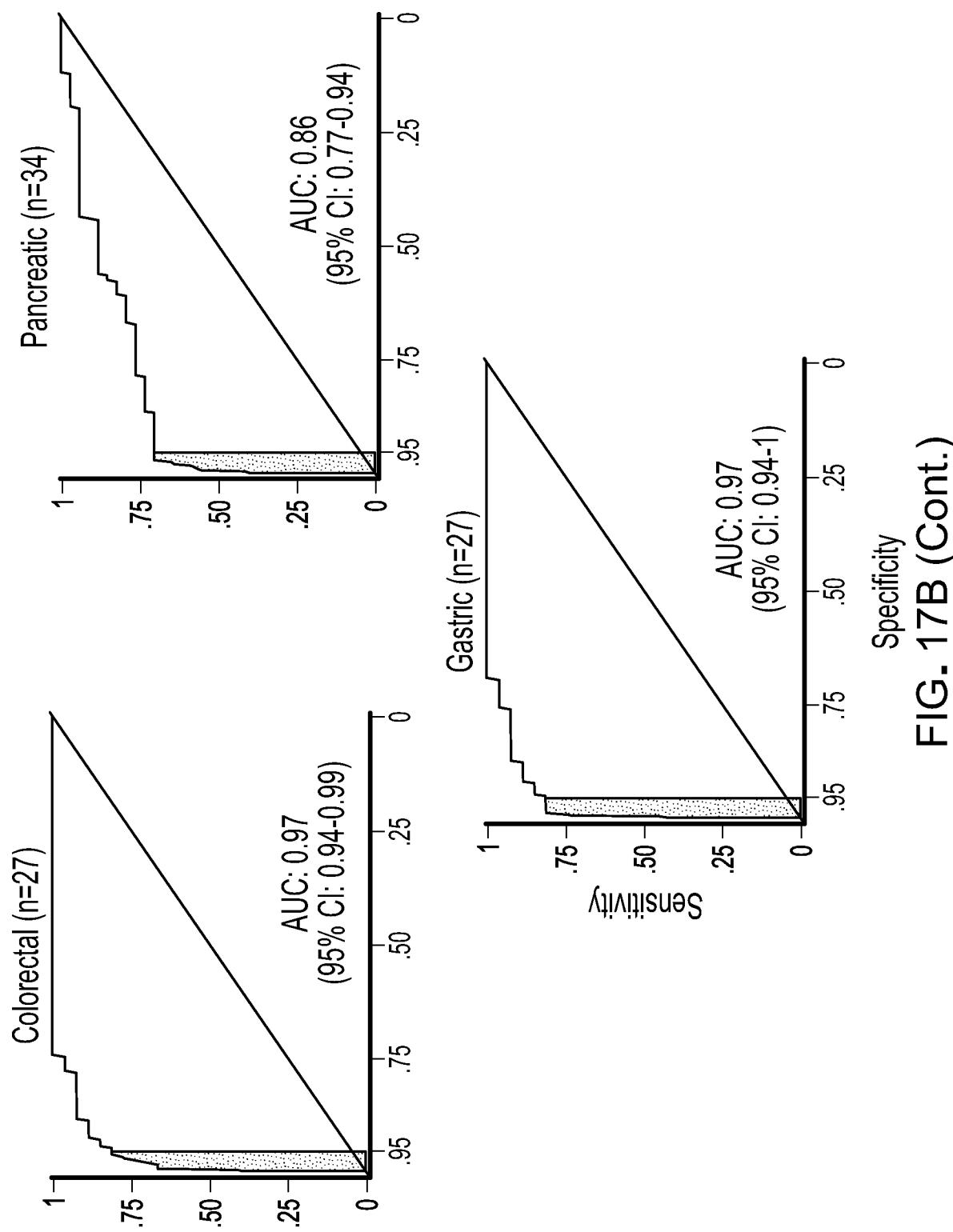


FIG. 17B (Cont.)

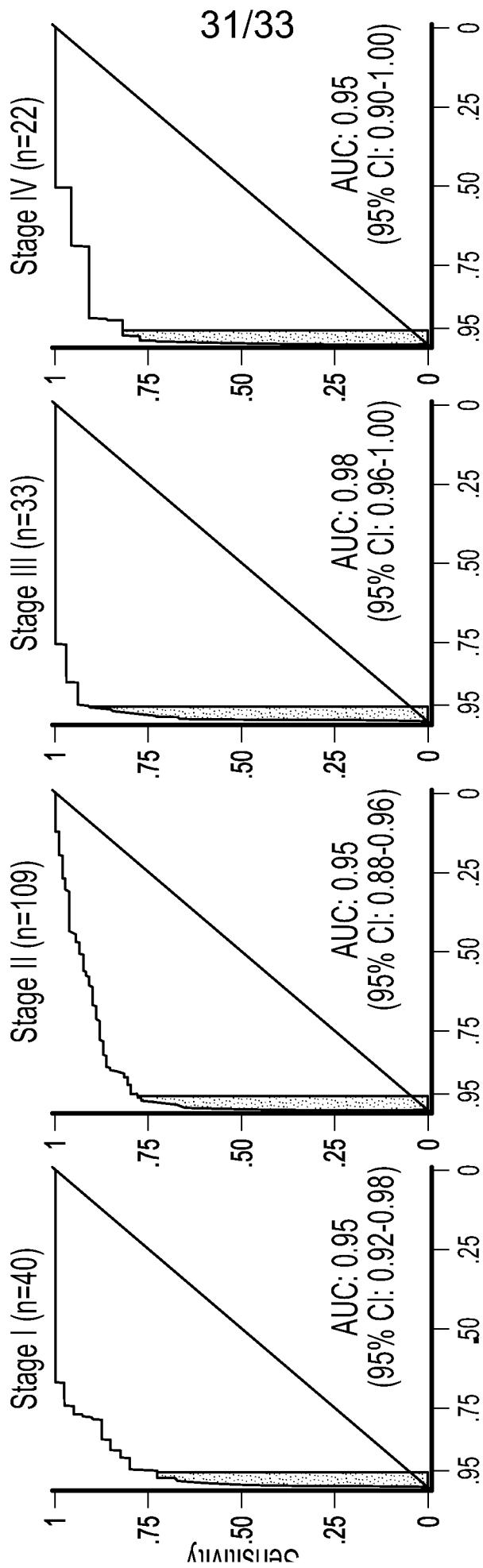


FIG. 18

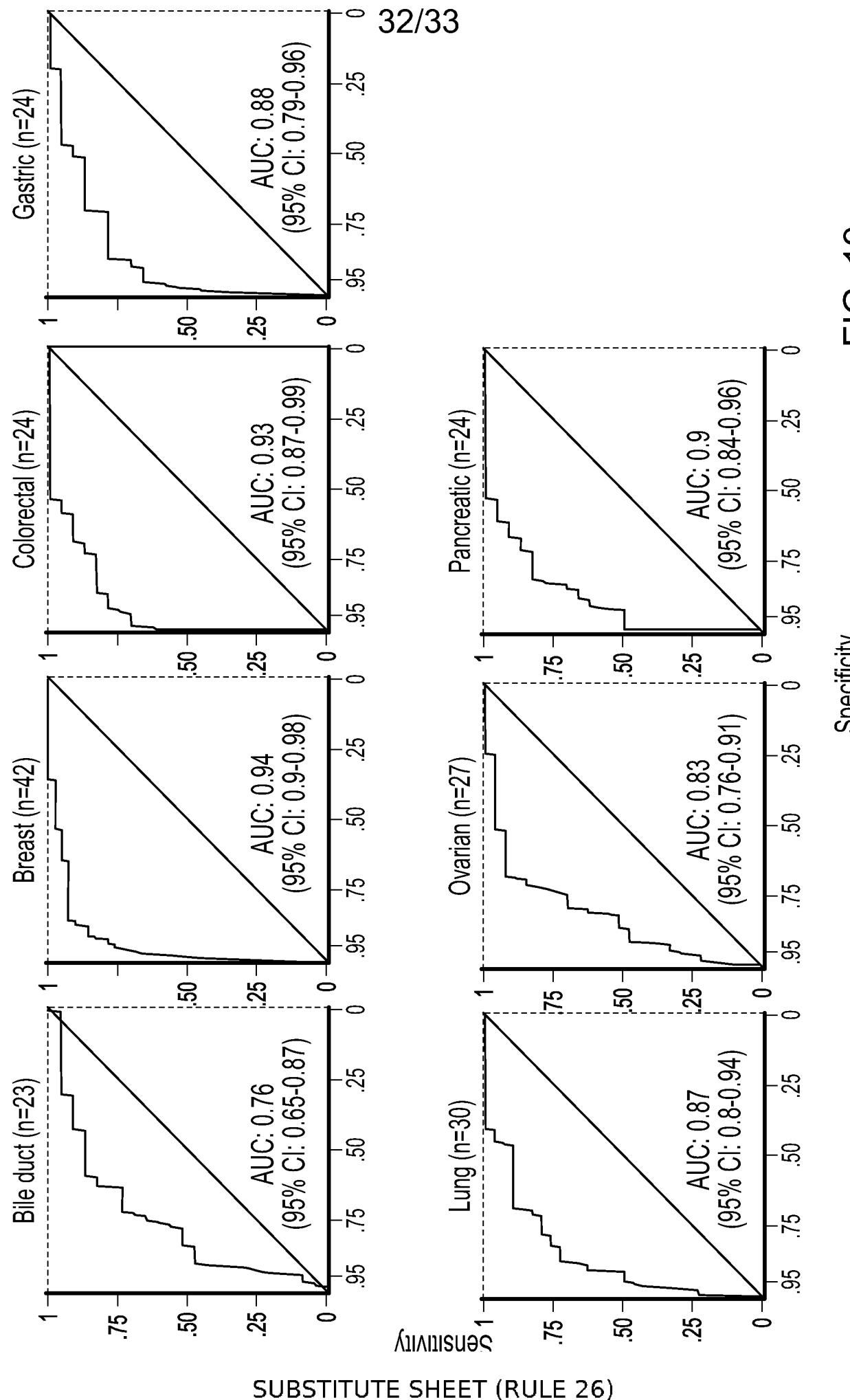
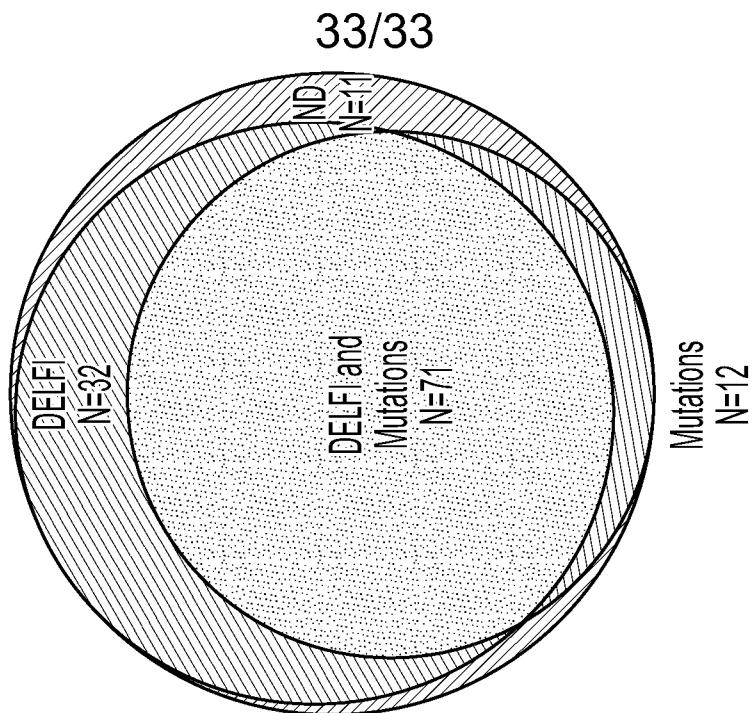


FIG. 19



Detection Approach*	Patients Analyzed	Patients Detected	Fraction of Patients Detected	95% CI
DELF1	126	103	82%	74%-88%
Mutations	126	83	66%	57%-74%
DELF1 and Mutations	126	115	91%	85%-96%
Stage I	32	27	84%	67%-95%
II	52	48	92%	81%-98%
III	25	23	92%	74%-99%
IV	16	16	100%	79%-100%

*Cancer Detection Using DELFI, Sequence Mutations, and the Combination of DELFI and Mutations was performed at Specificities of 98%, >99%, and 98%, respectively. Per Stage Sensitivities are included for all Cases Except for one Patient with Stage X.

FIG. 20

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US 2019/032914

A. CLASSIFICATION OF SUBJECT MATTER

G01N 33/50 (2006.01)
A61P 35/00 (2006.01)
C12Q 1/6869 (2018.01)
C12N 15/07 (2006.01)

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

G01N 33/50, A61P 35/00, C12Q 1/6869, C12N 15/07

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

PatSearch (RUPTO internal), USPTO, PAJ, Esp@cenet, Information Retrieval System of FIPS

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	TAYLOR F. et al. Unbiased Detection of Somatic Copy Number Aberrations in cfDNA of Lung Cancer Cases and High-Risk Controls with Low Coverage Whole Genome Sequencing. <i>Adv Exp Med Biol.</i> 2016;924:29-32., abstract, p. 30	1-3, 14-26, 34-46, 50-52
A	WO 2017/190067 A1 (IMPACT GENOMICS, INC) 02.11.2017, claims 1-33	1-3, 14-26, 34-46, 50-52
A	US 2017/0211143 A1 (UNIVERSITY OF WASHINGTON) 27.07.2017, claims 1, 48-52, 92, 95-96	1-3, 14-26, 34-46, 50-52
A	WO 2018/027176 A1 (THE BROAD INSTITUTE, INC) 08.02.2018, claims 1-52	1-3, 14-26, 34-46, 50-52
A	US 2017/0024513 A1 (THE CHINESE UNIVERSITY OF HONG KONG) 26.01.2017, claims 1-63	1-3, 14-26, 34-46, 50-52

 Further documents are listed in the continuation of Box C.

 See patent family annex.

* Special categories of cited documents:	"T"	later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
"A" document defining the general state of the art which is not considered to be of particular relevance	"X"	document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
"E" earlier document but published on or after the international filing date	"Y"	document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art
"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)	"&"	document member of the same patent family
"O" document referring to an oral disclosure, use, exhibition or other means		
"P" document published prior to the international filing date but later than the priority date claimed		

 Date of the actual completion of the international search
 05 September 2019 (05.09.2019)

 Date of mailing of the international search report
 05 September 2019 (05.09.2019)

 Name and mailing address of the ISA/RU:
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 Authorized officer
 D. Igumnov
 Telephone No. (495) 531-64-81

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US 2019/032914

Box No. II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)

This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. Claims Nos.:
because they relate to subject matter not required to be searched by this Authority, namely:

2. Claims Nos.:
because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:

3. Claims Nos.: 4-13, 27-33, 47-49, 53-67
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box No. III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

1. As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying additional fees, this Authority did not invite payment of additional fees.
3. As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:

4. No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

Remark on Protest

- The additional search fees were accompanied by the applicant's protest and, where applicable, the payment of a protest fee.
- The additional search fees were accompanied by the applicant's protest but the applicable protest fee was not paid within the time limit specified in the invitation.
- No protest accompanied the payment of additional search fees.