Disclaimer

Data obtained through Sentinel are intended to complement other types of evidence such as preclinical studies, clinical trials, postmarket studies, and adverse event reports, all of which are used by FDA to inform regulatory decisions regarding medical product safety. The information contained in this report is provided as part of FDA's commitment to place knowledge acquired from Sentinel in the public domain as soon as possible. Any public health actions taken by FDA regarding products involved in Sentinel queries will continue to be communicated through existing channels.

FDA wants to emphasize that the fact that FDA has initiated a query involving a medical product and is reporting findings related to that query does not mean that FDA is suggesting health care practitioners should change their prescribing practices for the medical product or that patients taking the medical product should stop using it. Patients who have questions about the use of an identified medical product should contact their health care practitioners.

The following report contains a description of the request, request specifications, and results from the modular program run(s).

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Overview for Request cder_mpl1r_wp021_nsdp_v01

Request ID: cder_mpl1r_wp021_nsdp_v01

Query Description: This report contains estimates of drug use among patients with a prior genetic test and/or relevant cancer diagnosis.

Sentinel Modular Program Tool Used: Cohort Identification and Descriptive Analysis (CIDA) tool, version 2.2.1

Data Source: The query was run against the Sentinel Distributed Database (SDD) for the time period of January 1, 2013 to December 31, 2015. The request was distributed to 14 Data Partners on June 17, 2016. See Appendix A for a list of the latest dates of available data for each Data Partner.

Study Design: This request was designed to calculate background rates. The number of qualifying patients with the exposure of interest were calculated overall and stratified by age group, sex, and year.

Exposure of Interest: The exposures of interest were cancer treatments (Cetuximab, Panitumumab, Trametinib, Dabrafenib, Vemurafenib, Cobimetinib, Afatinib, Erlotinib, Tagrisso, Gefitinib, Dasatinib, Imatinib, Bosutinib, Nilotinib, and Ponatinib), which were defined using National Drug Codes (NDCs) and Healthcare Common Procedure Coding System (HCPCS) Level II procedure codes. Please see Appendix B, C, and D for specific codes.

Cohort Eligibility Criteria: Patients were required to be continuously enrolled in plans with both medical and drug coverage for at least 183 days before their testing date, during which gaps in coverage of up to 45 days were allowed. Half of the scenarios restricted inclusion to patients who also had a relevant cancer indication and/or had one of the following genetic tests in the prior 183 days: V-Ki-ras2 Kirsten rat sarcoma viral oncogene (KRAS), v-raf murine sarcoma viral oncogene homolog B1 (BRAF), epidermal growth factor receptor (EGFR), breakpoint cluster region-abelson (BCR-ABL), and breast cancer susceptibility gene (BRCA). Cancer indications were defined using International Classification of Diseases, Ninth Revision, Clinical Modification (ICD-9-CM) diagnosis codes. Please refer to Appendix D for specific codes. Genetic tests were defined using HCPCS and Current Procedural Terminology (CPT-4) procedure codes. Please refer to Appendix E for specific codes. The following age groups were included in the cohort: 0-21, 22-44, 45-64, and 65+ years.

Limitations: Algorithms to define exposures and events are imperfect and, therefore, may be misclassified.

Please see the *Appendix F* for the specifications of parameters used in the analyses for this request.

Notes: Please contact the Sentinel Operations Center Query Fulfillment Team (production@mini-sentinel.org) for questions and to provide comments/suggestions for future enhancements to this document.

Sentinel

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Glossary of Terms for Analyses Using Cohort Identification and Descriptive Analysis (CIDA) Tool*

Amount Supplied - number of units (pills, tablets, vials) dispensed. Net amount per NDC per dispensing. This is equivalent to the "RxAmt" value in the MSCDM.

Blackout Period - number of days at the beginning of a treatment episode that events are to be ignored. If an event occurs during the blackout period, the episode is excluded.

Care Setting - type of medical encounter or facility where the exposure, event, or condition code was recorded. Possible care settings include: Inpatient Hospital Stay (IP), Non-Acute Institutional Stay (IS), Emergency Department (ED), Ambulatory Visit (AV), and Other Ambulatory Visit (OA). For laboratory results, possible care settings include: Emergency department (E), Home (H), Inpatient (I), Outpatient (O), or Unknown or Missing (U). Along with the Principal Diagnosis Indicator, forms the Care Setting/PDX parameter.

Cohort Definition (drug/exposure) - indicates how the cohort will be defined: 1: Cohort includes only the first valid incident treatment episode during the query period; 2: Cohort includes all valid incident treatment episodes during the query period; 3: Cohort includes all valid incident treatment episodes during the query period until an event occurs.

Days Supplied - number of days supplied for all dispensings in qualifying treatment episodes.

Episodes - treatment episodes; length of episode is determined by days supplied in one dispensing or consecutive dispensings bridged by

Eligible Members - number of members eligible for an incident treatment episode (defined by the drug/exposure and event washout periods) with drug and medical coverage during the query period.

Years at Risk - number of days supplied plus any episode gaps and exposure extension periods all divided by 365.25.

Enrollment Gap - number of days allowed between two consecutive enrollment periods without breaking a “continuously enrolled”

Episode Gap - number of days allowed between two (or more) consecutive exposures (dispensings/procedures) to be considered the same treatment episode.

Event Deduplication - specifies how events are counted by the MP algorithm: 0: Counts all occurrences of an HOI during an exposure episode; 1: de-duplicates occurrences of the same HOI code and code type on the same day; 2: de-duplicates occurrences of the same HOI group on the same day (e.g., de-duplicates at the group level).

Exposure Extension Period - number of days post treatment period in which the outcomes/events are counted for a treatment episode.

Exposure Episode Length - number of days after exposure initiation that is considered "exposed time."

Lookback Period (pre-existing condition) - number of days wherein a member is required to have evidence of pre-existing condition (diagnosis/procedure/drug dispensing).

Member-Years - sum of all days of enrollment with medical and drug coverage** in the query period preceded by an exposure washout

Minimum Days Supplied - specifies a minimum number of days in length of the days supplied for the episode to be considered.

Minimum Episode Duration - specifies a minimum number of days in length of the episode for it to be considered.

Principal Diagnosis (PDX) - diagnosis or condition established to be chiefly responsible for admission of the patient to the hospital. 'P' = principal diagnosis, 'S' = secondary diagnosis, 'X' = unspecified diagnosis, '.' = blank. Along with the Care Setting values, forms the

Treatment Episode Truncation Indicator - indicates whether observation of the incident query code during follow-up requires truncation of valid treatment episodes. A value of Y indicates that the treatment episodes should be truncated at the first occurrence of an incident query code. A value of N indicates that the treatment episodes should not be truncated at the occurrence of the incident query code.

Users - number of members with exposure during the query period. Member must have no evidence of exposure(s) of interest (defined by incidence criteria) in the prior washout period. A user may only be counted once in a query period.

Washout Period (drug/exposure) -** number of days a user is required to have no evidence of prior exposure (drug dispensing/procedure) and continuous drug and medical coverage prior to an incident treatment episode.

Washout Period (event/outcome) -** number of days a user is required to have no evidence of a prior event (procedure/diagnosis) and continuous drug and medical coverage prior to an incident treatment episode.

*all terms may not be used in this report

**incident treatment episodes must be incident to both the exposure and the event

Table 1: Summary of Incident Cancer Treatment in the SDD between January 1, 2013 and December 31, 2015, by Drug Group, Cancer Diagnosis, and Genetic Test

	New Users	Percentage of All New Users	Eligible Members	Member- Years	New Users / 1K Eligible Members
Cetuximab and Panitumumab					
All New Users					
	5,220	100.00%	63,192,462	94,429,489.8	0.08
New Users with Colorectal Cancer					
	2,655	50.86%	175,920	143,775.3	15.09
New Users with a prior KRAS test					
	710	13.60%	17,020	6,343.3	41.72
New Users with Colorectal Cancer and a prior KRAS test					
	647	12.39%	Not Available	Not Available	Not Available
Dasatinib, Imatinib, Bosutinib, Nilotinib, Ponatinib					
All New Users					
	3,545	100.00%	63,187,120	94,419,678.6	0.06
New Users with Leukemia					
	2,139	60.34%	98,166	93,499.4	21.79
New Users with a Previous BCR-ABL Test					
	1,191	33.60%	19,763	8,103.4	60.26
New Users with Leukemia and a prior BCR-ABL test					
	1,115	31.45%	Not Available	Not Available	Not Available
Olaparib					
All New Users					
	141	100.00%	63,193,377	94,434,448.9	0.00
New Users with Ovarian Cancer					
	125	88.65%	49,746	40,840.9	2.51
New Users with a Prior BRCA test					
	30	21.28%	111,916	43,524.0	0.27
New Users with a Prior BRCA test and Ovarian Cancer					
	23	16.31%	Not Available	Not Available	Not Available

Table 1: Summary of Incident Cancer Treatment in the SDD between January 1, 2013 and December 31, 2015, by Drug Group, Cancer Diagnosis, and Genetic Test

	New Users	Percentage of All New Users	Eligible Members	Member- Years	New Users / 1K Eligible Members
Cetuximab, Panitumumab, Afatinib, Erlotinib, Tagrisso, Gefitinib					
All New Users					
	9,179	100.00%	63,190,877	94,424,568.7	0.15
New Users with Lung or Colorectal Cancer					
	4,189	45.64%	163,059	117,240.4	25.69
New Users with a Prior EGFR test					
	1,022	11.13%	18,167	6,472.8	56.26
New Users with Lung or Colorectal Cancer and a Prior EGFR test					
	879	9.58%	Not Available	Not Available	Not Available
Trametinib, Dabrafenib, Vemurafenib, Cobimetinib					
All New Users					
	1,078	100.00%	63,193,109	94,433,523.9	0.02
New Users with Melanoma					
	913	84.69%	160,176	98,127.2	5.70
New Users with a Prior BRAF test					
	338	31.35%	16,048	5,975.0	21.06
New Users with Melanoma and a Prior BRAF test					
	287	26.62%	Not available	Not available	Not available

Table 2: Summary of Incident Cancer Treatment in the SDD between January 1, 2013 and December 31, 2015, by Drug Group, Cancer Diagnosis, Genetic Test, and Age Group

	New Users	Percentage of All New Users	Eligible Members	Member- Years	New Users / 1K Eligible Members
Cetuximab and Panitumumab					
All New Users					
0-21 Years	10	100.00%	16,837,978	23,987,670.6	0.00
22-44 Years	315	100.00%	23,352,856	29,366,624.6	0.01
45-64 Years	2,443	100.00%	19,195,622	28,412,051.4	0.13
65+ Years	2,452	100.00%	7,364,240	12,663,143.2	0.33
New Users with Colorectal Cancer					
0-21 Years	2	20.00%	390	183.0	5.13
22-44 Years	226	71.75%	9,955	7,130.1	22.70
45-64 Years	1,316	53.87%	72,565	56,307.1	18.14
65+ Years	1,111	45.31%	97,456	80,155.1	11.40
New Users with a prior KRAS test					
0-21 Years	3	30.00%	258	99.9	11.63
22-44 Years	65	20.63%	1,986	733.1	32.73
45-64 Years	369	15.10%	9,031	3,344.8	40.86
65+ Years	273	11.13%	5,977	2,165.4	45.68
New Users with Colorectal Cancer and a prior KRAS test					
0-21 Years	2	20.00%	Not Available	Not Available	Not Available
22-44 Years	56	17.78%	Not Available	Not Available	Not Available
45-64 Years	335	13.71%	Not Available	Not Available	Not Available
65+ Years	254	10.36%	Not Available	Not Available	Not Available
Dasatinib, Imatinib, Bosutinib, Nilotinib, Ponatinib					
All New Users					
0-21 Years	98	100.00%	16,837,861	23,987,420.4	0.01
22-44 Years	744	100.00%	23,351,739	29,364,477.4	0.03
45-64 Years	1,599	100.00%	19,192,730	28,407,369.5	0.08
65+ Years	1,104	100.00%	7,362,590	12,660,411.4	0.15

Table 2: Summary of Incident Cancer Treatment in the SDD between January 1, 2013 and December 31, 2015, by Drug Group, Cancer Diagnosis, Genetic Test, and Age Group

	New Users	Percentage of All New Users	Eligible Members	Member- Years	New Users / 1K Eligible Members
New Users with Leukemia					
0-21 Years	57	58.16%	8,956	8,538.4	6.36
22-44 Years	525	70.56%	10,842	6,608.9	48.42
45-64 Years	958	59.91%	32,835	27,664.7	29.18
65+ Years	599	54.26%	48,354	50,687.4	12.39
New Users with a Previous BCR-ABL Test					
0-21 Years	15	15.31%	360	155.0	41.67
22-44 Years	293	39.38%	4,615	1,793.4	63.49
45-64 Years	582	36.40%	9,721	3,890.8	59.87
65+ Years	301	27.26%	5,371	2,264.2	56.04
New Users with Leukemia and a prior BCR-ABL test					
0-21 Years	13	13.27%	Not Available	Not Available	Not Available
22-44 Years	281	37.77%	Not Available	Not Available	Not Available
45-64 Years	542	33.90%	Not Available	Not Available	Not Available
65+ Years	279	25.27%	Not Available	Not Available	Not Available

Olaparib

All New Users					
0-21 Years	0	100.00%	16,837,979	23,987,679.4	0.00
22-44 Years	12	100.00%	23,352,936	29,366,917.2	0.00
45-64 Years	100	100.00%	19,196,142	28,414,362.2	0.01
65+ Years	29	100.00%	7,364,710	12,665,490.1	0.00
New Users with Ovarian Cancer					
0-21 Years	0	N/A	593	403.3	0.00
22-44 Years	8	66.67%	7,595	4,789.8	1.05
45-64 Years	91	91.00%	25,302	20,623.0	3.60
65+ Years	26	89.66%	17,809	15,024.8	1.46

Table 2: Summary of Incident Cancer Treatment in the SDD between January 1, 2013 and December 31, 2015, by Drug Group, Cancer Diagnosis, Genetic Test, and Age Group

	New Users	Percentage of All New Users	Eligible Members	Member- Years	New Users / 1K Eligible Members
New Users with a Prior BRCA test					
0-21 Years	0	N/A	1,219	438.4	0.00
22-44 Years	3	25.00%	43,929	16,451.2	0.07
45-64 Years	21	21.00%	60,595	23,584.2	0.35
65+ Years	6	20.69%	8,137	3,050.2	0.74
New Users with a Prior BRCA test and Ovarian Cancer					
0-21 Years	0	N/A	Not Available	Not Available	Not Available
22-44 Years	1	8.33%	Not Available	Not Available	Not Available
45-64 Years	17	17.00%	Not Available	Not Available	Not Available
65+ Years	5	17.24%	Not Available	Not Available	Not Available

Cetuximab, Panitumumab, Afatinib, Erlotinib, Tagrisso, Gefitinib

All New Users					
0-21 Years	19	100.00%	16,837,973	23,987,659.6	0.00
22-44 Years	446	100.00%	23,352,804	29,366,474.5	0.02
45-64 Years	4,009	100.00%	19,194,850	28,410,106.4	0.21
65+ Years	4,705	100.00%	7,363,332	12,660,328.2	0.64
New Users with Lung or Colorectal Cancer					
0-21 Years	1	5.26%	451	222.3	2.22
22-44 Years	122	27.35%	4,646	2,314.8	26.26
45-64 Years	1,652	41.21%	54,144	35,424.9	30.51
65+ Years	2,414	51.31%	106,950	79,278.4	22.57
New Users with a Prior EGFR test					
0-21 Years	2	10.53%	129	50.7	15.50
22-44 Years	56	12.56%	1,239	444.1	45.20
45-64 Years	430	10.73%	8,382	2,982.6	51.30
65+ Years	534	11.35%	8,613	2,995.5	62.00

Table 2: Summary of Incident Cancer Treatment in the SDD between January 1, 2013 and December 31, 2015, by Drug Group, Cancer Diagnosis, Genetic Test, and Age Group

	New Users	Percentage of All New Users	Eligible Members	Member- Years	New Users / 1K Eligible Members
New Users with Lung or Colorectal Cancer and a Prior EGFR test					
0-21 Years	0	0.00%	Not Available	Not Available	Not Available
22-44 Years	38	8.52%	Not Available	Not Available	Not Available
45-64 Years	353	8.81%	Not Available	Not Available	Not Available
65+ Years	488	10.37%	Not Available	Not Available	Not Available

Trametinib, Dabrafenib, Vemurafenib, Cobimetinib

All New Users					
0-21 Years	20	100.00%	16,837,975	23,987,660.6	0.00
22-44 Years	176	100.00%	23,352,873	29,366,744.6	0.01
45-64 Years	556	100.00%	19,195,967	28,413,911.3	0.03
65+ Years	326	100.00%	7,364,653	12,665,207.4	0.04
New Users with Melanoma					
0-21 Years	7	35.00%	1,369	678.8	5.11
22-44 Years	156	88.64%	22,629	12,251.0	6.89
45-64 Years	467	83.99%	72,951	43,264.5	6.40
65+ Years	283	86.81%	66,739	41,932.9	4.24
New Users with a Prior BRAF test					
0-21 Years	5	25.00%	342	134.6	14.62
22-44 Years	43	24.43%	2,381	880.4	18.06
45-64 Years	176	31.65%	8,284	3,065.8	21.25
65+ Years	114	34.97%	5,261	1,894.1	21.67
New Users with Melanoma and a Prior BRAF test					
0-21 Years	2	10.00%	Not Available	Not Available	Not Available
22-44 Years	38	21.59%	Not Available	Not Available	Not Available
45-64 Years	148	26.62%	Not Available	Not Available	Not Available
65+ Years	99	30.37%	Not Available	Not Available	Not Available

Table 3: Summary of Incident Cancer Treatment in the SDD between January 1, 2013 and December 31, 2015, by Drug Group, Cancer Diagnosis, Genetic Test, and Sex

	New Users	Percentage of All New Users	Eligible Members	Member- Years	New Users / 1K Eligible Members
Cetuximab and Panitumumab					
All New Users					
Female	1,759	100.00%	32,126,116	48,246,347.6	0.05
Male	3,459	100.00%	31,063,571	46,179,370.5	0.11
Unknown	2	100.00%	2,775	3,771.7	0.72
New Users with Colorectal Cancer					
Female	1,123	63.84%	85,907	69,781.5	13.07
Male	1,532	44.29%	90,000	73,982.6	17.02
Unknown	0	0.00%	13	11.1	0.00
New Users with a prior KRAS test					
Female	307	17.45%	9,292	3,508.2	33.04
Male	403	11.65%	7,724	2,833.0	52.18
Unknown	0	0.00%	4	2.0	0.00
New Users with Colorectal Cancer and a prior KRAS test					
Female	287	16.32%	Not Available	Not Available	Not Available
Male	360	10.41%	Not Available	Not Available	Not Available
Unknown	0	0.00%	Not Available	Not Available	Not Available
Dasatinib, Imatinib, Bosutinib, Nilotinib, Ponatinib					
All New Users					
Female	1,613	100.00%	32,123,691	48,241,414.1	0.05
Male	1,932	100.00%	31,060,654	46,174,491.7	0.06
Unknown	0	100.00%	2,775	3,772.8	0.00
New Users with Leukemia					
Female	927	57.47%	44,464	41,464.8	20.85
Male	1,212	62.73%	53,697	52,030.5	22.57
Unknown	0	N/A	5	4.1	0.00
New Users with a Previous BCR-ABL Test					
Female	521	32.30%	11,161	4,556.1	46.68
Male	670	34.68%	8,600	3,546.7	77.91
Unknown	0	N/A	2	0.7	0.00

Table 3: Summary of Incident Cancer Treatment in the SDD between January 1, 2013 and December 31, 2015, by Drug Group, Cancer Diagnosis, Genetic Test, and Sex

	New Users	Percentage of All New Users	Eligible Members	Member- Years	New Users / 1K Eligible Members
New Users with Leukemia and a prior BCR-ABL test					
Female	486	30.13%	Not Available	Not Available	Not Available
Male	629	32.56%	Not Available	Not Available	Not Available
Unknown	0	N/A	Not Available	Not Available	Not Available

Olaparib

All New Users					
Female	135	100.00%	32,126,399	48,247,974.7	0.00
Male	6	100.00%	31,064,203	46,182,701.0	0.00
Unknown	0	100.00%	2,775	3,773.3	0.00
New Users with Ovarian Cancer					
Female	125	92.59%	49,328	40,653.8	2.53
Male	0	0.00%	412	180.9	0.00
Unknown	0	N/A	6	6.3	0.00
New Users with a Prior BRCA test					
Female	26	19.26%	106,777	41,610.3	0.24
Male	4	66.67%	5,134	1,911.6	0.78
Unknown	0	N/A	5	2.1	0.00
New Users with a Prior BRCA test and Ovarian Cancer					
Female	23	17.04%	Not Available	Not Available	Not Available
Male	0	0.00%	Not Available	Not Available	Not Available
Unknown	0	N/A	Not Available	Not Available	Not Available

Cetuximab, Panitumumab, Afatinib, Erlotinib, Tagrisso, Gefitinib

All New Users					
Female	3,998	100.00%	32,125,144	48,243,291.4	0.12
Male	5,178	100.00%	31,062,958	46,177,505.9	0.17
Unknown	3	100.00%	2,775	3,771.4	1.08

Table 3: Summary of Incident Cancer Treatment in the SDD between January 1, 2013 and December 31, 2015, by Drug Group, Cancer Diagnosis, Genetic Test, and Sex

	New Users	Percentage of All New Users	Eligible Members	Member- Years	New Users / 1K Eligible Members
New Users with Lung or Colorectal Cancer					
Female	2,275	56.90%	Not Available	Not Available	Not Available
Male	1,913	36.94%	Not Available	Not Available	Not Available
Unknown	1	33.33%	Not Available	Not Available	Not Available
New Users with a Prior EGFR test					
Female	587	14.68%	9,981	3,616.1	58.81
Male	435	8.40%	8,183	2,855.2	53.16
Unknown	0	0.00%	3	1.5	0.00
New Users with Lung or Colorectal Cancer and a Prior EGFR test					
Female	528	13.21%	32,125,144	48,243,822.3	0.02
Male	351	6.78%	31,062,958	46,178,395.9	0.01
Unknown	0	0.00%	2,775	3,772.3	0.00

Trametinib, Dabrafenib, Vemurafenib, Cobimetinib

All New Users					
Female	408	100.00%	32,126,289	48,247,637.5	0.01
Male	670	100.00%	31,064,045	46,182,113.1	0.02
Unknown	0	100.00%	2,775	3,773.3	0.00
New Users with Melanoma					
Female	334	81.86%	75,126	45,001.5	4.45
Male	579	86.42%	85,045	53,122.5	6.81
Unknown	0	N/A	5	3.2	0.00
New Users with a Prior BRAF test					
Female	124	30.39%	8,931	3,353.3	13.88
Male	214	31.94%	7,115	2,621.2	30.08
Unknown	0	N/A	2	0.5	0.00
New Users with Melanoma and a Prior BRAF test					
Female	103	25.25%	Not Available	Not Available	Not Available
Male	184	27.46%	Not Available	Not Available	Not Available
Unknown	0	N/A	Not Available	Not Available	Not Available

Table 4: Summary of Incident Cancer Treatment in the SDD between January 1, 2013 and December 31, 2015, by Drug Group, Cancer Diagnosis, Genetic Test, and Year

	New Users	Percentage of All New Users	Eligible Members	Member- Years	New Users / 1K Eligible Members
Cetuximab and Panitumumab					
All New Users					
2013	1,974	100.00%	44,428,764	35,277,704.1	0.04
2014	1,979	100.00%	45,834,292	35,215,004.2	0.04
2015	1,267	100.00%	40,569,004	23,936,781.5	0.03
New Users with Colorectal Cancer					
2013	1,003	50.81%	101,791	53,775.6	9.85
2014	996	50.33%	104,587	53,931.5	9.52
2015	656	51.78%	87,083	36,068.2	7.53
New Users with a prior KRAS test					
2013	231	11.70%	5,202	1,594.7	44.41
2014	306	15.46%	9,463	2,743.8	32.34
2015	173	13.65%	7,561	2,004.8	22.88
New Users with Colorectal Cancer and a prior KRAS test					
2013	217	10.99%	Not Available	Not Available	Not Available
2014	279	14.10%	Not Available	Not Available	Not Available
2015	151	11.92%	Not Available	Not Available	Not Available
Dasatinib, Imatinib, Bosutinib, Nilotinib, Ponatinib					
All New Users					
2013	1,368	100.00%	44,424,405	35,274,061.0	0.03
2014	1,326	100.00%	45,830,060	35,211,365.8	0.03
2015	851	100.00%	40,565,080	23,934,251.9	0.02
New Users with Leukemia					
2013	845	61.77%	58,935	34,182.2	14.34
2014	818	61.69%	61,126	35,081.6	13.38
2015	476	55.93%	52,116	24,235.5	9.13
New Users with a Previous BCR-ABL Test					
2013	417	30.48%	6,148	2,007.1	67.83
2014	473	35.67%	11,212	3,512.3	42.19
2015	301	35.37%	9,039	2,584.0	33.30

Table 4: Summary of Incident Cancer Treatment in the SDD between January 1, 2013 and December 31, 2015, by Drug Group, Cancer Diagnosis, Genetic Test, and Year

	New Users	Percentage of All New Users	Eligible Members	Member- Years	New Users / 1K Eligible Members
New Users with Leukemia and a prior BCR-ABL test					
2013	386	28.22%	Not Available	Not Available	Not Available
2014	456	34.39%	Not Available	Not Available	Not Available
2015	273	32.08%	Not Available	Not Available	Not Available

Olaparib

All New Users					
2013	0	100.00%	44,429,494	35,279,176.2	0.00
2014	0	100.00%	45,835,954	35,216,961.5	0.00
2015	141	100.00%	40,571,096	23,938,311.2	0.00
New Users with Ovarian Cancer					
2013	0	N/A	29,349	15,690.8	0.00
2014	0	N/A	29,232	15,185.2	0.00
2015	125	88.65%	23,649	9,964.9	5.29
New Users with a Prior BRCA test					
2013	0	N/A	44,727	14,289.1	0.00
2014	0	N/A	57,315	17,393.0	0.00
2015	30	21.28%	43,858	11,841.9	0.68
New Users with a Prior BRCA test and Ovarian Cancer					
2013	0	N/A	Not Available	Not Available	Not Available
2014	0	N/A	Not Available	Not Available	Not Available
2015	23	16.31%	Not Available	Not Available	Not Available

Cetuximab, Panitumumab, Afatinib, Erlotinib, Tagrisso, Gefitinib

All New Users					
2013	3,631	100.00%	44,427,377	35,275,972.5	0.08
2014	3,449	100.00%	45,832,487	35,213,161.4	0.08
2015	2,099	100.00%	40,567,154	23,935,434.8	0.05
New Users with Lung or Colorectal Cancer					
2013	1,697	46.74%	88,031	43,206.7	19.28
2014	1,590	46.10%	91,124	44,218.9	17.45
2015	902	42.97%	74,970	29,814.8	12.03

Table 4: Summary of Incident Cancer Treatment in the SDD between January 1, 2013 and December 31, 2015, by Drug Group, Cancer Diagnosis, Genetic Test, and Year

	New Users	Percentage of All New Users	Eligible Members	Member- Years	New Users / 1K Eligible Members
Patients with a Prior EGFR test					
2013	293	8.07%	5,372	1,551.2	54.54
2014	434	12.58%	10,035	2,875.9	43.25
2015	295	14.05%	7,961	2,045.7	37.06
New Users with Lung or Colorectal Cancer and a Prior EGFR test					
2013	267	7.35%	Not Available	Not Available	Not Available
2014	371	10.76%	Not Available	Not Available	Not Available
2015	241	11.48%	Not Available	Not Available	Not Available

Trametinib, Dabrafenib, Vemurafenib, Cobimetinib

All New Users					
2013	373	100.00%	44,429,292	35,278,898.1	0.01
2014	451	100.00%	45,835,609	35,216,579.5	0.01
2015	254	100.00%	40,570,673	23,938,046.4	0.01
New Users with Melanoma					
2013	329	88.20%	85,071	36,660.6	3.87
2014	382	84.70%	86,599	36,600.5	4.41
2015	202	79.53%	70,316	24,866.0	2.87
New Users with a Prior BRAF test					
2013	100	26.81%	4,171	1,291.0	23.98
2014	146	32.37%	8,666	2,559.7	16.85
2015	92	36.22%	7,968	2,124.3	11.55
New Users with Melanoma and a Prior BRAF test					
2013	87	23.32%	Not Available	Not Available	Not Available
2014	122	27.05%	Not Available	Not Available	Not Available
2015	78	30.71%	Not Available	Not Available	Not Available

Appendix A: Latest Date of Available Data for Each Data Partner up to Request End Date (5/15/2016)

DP ID	End Date
DP0001	6/30/2015
DP0002	4/30/2015
DP0003	12/31/2014
DP0004	10/31/2014
DP0005	11/30/2015
DP0006	2/28/2015
DP0007	12/31/2015
DP0008	9/30/2015
DP0009	11/30/2015
DP0010	7/31/2015
DP0011	7/31/2014
DP0012	9/30/2015
DP0013	6/30/2015
DP0014	10/31/2015

Appendix B: Generic and Brand Names used to Define Exposures in this Request

Generic Name	Brand Name
CETUXIMAB	Erbitux
PANITUMUMAB	Vectibix
COBIMETINIB FUMARATE	Cotellic
TRAMETINIB DIMETHYL SULFOXIDE	Mekinist
DABRAFENIB MESYLATE	Tafinlar
VEMURAFENIB	Zelboraf
AFATINIB DIMALEATE	Gilotrif
GEFITINIB	Iressa
OSIMERTINIB MESYLATE	Tagrisso
ERLOTINIB HCL	Tarceva
BOSUTINIB	Bosulif
IMATINIB MESYLATE	Gleevec
PONATINIB HCL	Iclusig
DASATINIB	Sprycel
NILOTINIB HCL	Tasigna
OLAPARIB	Lynparza

Appendix C: List of Procedure Codes Used to Define Exposures in this Request

Code	Description	Code Type
KRAS Drug Pairs		
J9055	Injection, cetuximab, 10 mg	HCPCS Procedure
C9235	Injection, panitumumab, 10 mg	HCPCS Procedure
C9215	Injection, cetuximab, per 10 mg	HCPCS Procedure
J9303	Injection, panitumumab, 10 mg	HCPCS Procedure
EGFR Drug Pairs		
J9055	Injection, cetuximab, 10 mg	HCPCS Procedure
C9235	Injection, panitumumab, 10 mg	HCPCS Procedure
C9215	Injection, cetuximab, per 10 mg	HCPCS Procedure
J9303	Injection, panitumumab, 10 mg	HCPCS Procedure
J8565	Gefitinib, oral, 250 mg	HCPCS Procedure
BCR-ABL Drug Pairs		
S0088	Imatinib, 100 mg	HCPCS Procedure

Appendix D: List of Diagnosis Codes used to Define Cancer Inclusion Criteria in this Request

Code	Description
Colorectal Cancer	
153	Malignant neoplasm of colon
153.1	Malignant neoplasm of transverse colon
153.2	Malignant neoplasm of descending colon
153.3	Malignant neoplasm of sigmoid colon
153.6	Malignant neoplasm of ascending colon
153.9	Malignant neoplasm of colon, unspecified site
154	Malignant neoplasm of rectum, rectosigmoid junction, and anus
154.1	Malignant neoplasm of rectum
154.8	Malignant neoplasm of other sites of rectum, rectosigmoid junction, and anus
230.3	Carcinoma in situ of colon
230.4	Carcinoma in situ of rectum
Melanoma	
172	Malignant melanoma of skin
172.5	Malignant melanoma of skin of trunk, except scrotum
172.3	Malignant melanoma of skin of other and unspecified parts of face
172.8	Malignant melanoma of other specified sites of skin
172.2	Malignant melanoma of skin of ear and external auditory canal
172.6	Malignant melanoma of skin of upper limb, including shoulder
172.4	Malignant melanoma of skin of scalp and neck
172.1	Malignant melanoma of skin of eyelid, including canthus
172.9	Melanoma of skin, site unspecified
172.0	Malignant melanoma of skin of lip
172.7	Malignant melanoma of skin of lower limb, including hip
Lung Cancer	
162	Malignant neoplasm of trachea, bronchus, and lung
162.3	Malignant neoplasm of upper lobe, bronchus, or lung
162.4	Malignant neoplasm of middle lobe, bronchus, or lung
162.5	Malignant neoplasm of lower lobe, bronchus, or lung
162.8	Malignant neoplasm of other parts of bronchus or lung
162.9	Malignant neoplasm of bronchus and lung, unspecified site
231.2	Carcinoma in situ of bronchus and lung
Leukemia	
204	Lymphoid leukemia
204.0	Acute lymphoid leukemia
204.00	Acute lymphoid leukemia, without mention of having achieved remission
204.01	Acute lymphoid leukemia in remission
204.02	Acute lymphoid leukemia, in relapse
204.1	Chronic lymphoid leukemia
204.10	Chronic lymphoid leukemia, without mention of having achieved remission
204.11	Chronic lymphoid leukemia in remission
204.12	Chronic lymphoid leukemia, in relapse
204.2	Subacute lymphoid leukemia
204.20	Subacute lymphoid leukemia, without mention of having achieved remission
204.21	Subacute lymphoid leukemia in remission
204.22	Subacute lymphoid leukemia, in relapse
204.8	Other lymphoid leukemia
204.80	Other lymphoid leukemia, without mention of having achieved remission
204.81	Other lymphoid leukemia in remission
204.82	Other lymphoid leukemia, in relapse

204.9	Unspecified lymphoid leukemia
204.90	Unspecified lymphoid leukemia, without mention of having achieved remission
204.91	Unspecified lymphoid leukemia in remission
204.92	Unspecified lymphoid leukemia, in relapse
205	Myeloid leukemia
205.0	Acute myeloid leukemia
205.00	Acute myeloid leukemia, without mention of having achieved remission
205.01	Acute myeloid leukemia in remission
205.02	Acute myeloid leukemia, in relapse
205.1	Chronic myeloid leukemia
205.10	Chronic myeloid leukemia, without mention of having achieved remission
205.11	Chronic myeloid leukemia in remission
205.12	Chronic myeloid leukemia, in relapse
205.2	Subacute myeloid leukemia
205.20	Subacute myeloid leukemia, without mention of having achieved remission
205.21	Subacute myeloid leukemia in remission
205.22	Subacute myeloid leukemia, in relapse
205.8	Other myeloid leukemia
205.80	Other myeloid leukemia, without mention of having achieved remission
205.81	Other myeloid leukemia in remission
205.82	Other myeloid leukemia, in relapse
205.9	Unspecified myeloid leukemia
205.90	Unspecified myeloid leukemia, without mention of having achieved remission
205.91	Unspecified myeloid leukemia in remission
205.92	Unspecified myeloid leukemia, in relapse
206	Monocytic leukemia
206.0	Acute monocytic leukemia
206.00	Acute monocytic leukemia, without mention of having achieved remission
206.01	Acute monocytic leukemia in remission
206.02	Acute monocytic leukemia, in relapse
206.1	Chronic monocytic leukemia
206.10	Chronic monocytic leukemia, without mention of having achieved remission
206.11	Chronic monocytic leukemia in remission
206.12	Chronic monocytic leukemia, in relapse
206.2	Subacute monocytic leukemia
206.20	Subacute monocytic leukemia, without mention of having achieved remission
206.21	Subacute monocytic leukemia in remission
206.22	Subacute monocytic leukemia, in relapse
206.8	Other monocytic leukemia
206.80	Other monocytic leukemia, without mention of having achieved remission
206.81	Other monocytic leukemia in remission
206.82	Other monocytic leukemia, in relapse
206.9	Unspecified monocytic leukemia
206.90	Unspecified monocytic leukemia, without mention of having achieved remission
206.91	Unspecified monocytic leukemia in remission
206.92	Unspecified monocytic leukemia, in relapse
207	Other specified leukemia

207.0	Acute erythremia and erythroleukemia
207.00	Acute erythremia and erythroleukemia, without mention of having achieved remission
207.01	Acute erythremia and erythroleukemia in remission
207.02	Acute erythremia and erythroleukemia, in relapse
207.2	Megakaryocytic leukemia
207.20	Megakaryocytic leukemia, without mention of having achieved remission
207.21	Megakaryocytic leukemia in remission
207.22	Megakaryocytic leukemia, in relapse
207.8	Other specified leukemia
207.80	Other specified leukemia, without mention of having achieved remission
207.81	Other specified leukemia in remission
207.82	Other specified leukemia, in relapse
208	Leukemia of unspecified cell type
208.0	Acute leukemia of unspecified cell type
208.00	Acute leukemia of unspecified cell type, without mention of having achieved remission
208.01	Acute leukemia of unspecified cell type in remission
208.02	Acute leukemia of unspecified cell type, in relapse
208.1	Chronic leukemia of unspecified cell type
208.10	Chronic leukemia of unspecified cell type, without mention of having achieved remission
208.11	Chronic leukemia of unspecified cell type in remission
208.12	Chronic leukemia of unspecified cell type, in relapse
208.2	Subacute leukemia of unspecified cell type
208.20	Subacute leukemia of unspecified cell type, without mention of having achieved remission
208.21	Subactue leukemia of unspecified cell type in remission
208.22	Subacute leukemia of unspecified cell type, in relapse
208.8	Other leukemia of unspecified cell type
208.80	Other leukemia of unspecified cell type, without mention of having achieved remission
208.81	Other leukemia of unspecified cell type in remission
208.82	Other leukemia of unspecified cell type, in relapse
208.9	Unspecified leukemia
208.90	Unspecified leukemia, without mention of having achieved remission
208.91	Unspecified leukemia in remission
208.92	Unspecified leukemia, in relapse
Ovarian Cancer	
183	Malignant neoplasm of ovary and other uterine adnexa
183.0	Malignant neoplasm of ovary

Appendix E: List of Procedure Codes used to Define Genetic Tests Inclusion Criteria in this Request

Code	Description	Code Type
KRAS		
81275	KRAS (v-Ki-ras2 Kirsten rat sarcoma viral oncogene) (eg, carcinoma) gene analysis, variants in codons 12 and 13	CPT-4 Procedure
S3713	Kras mutation analysis testing	HCPCS Procedure
BRAF		
81210	BRAF (v-raf murine sarcoma viral oncogene homolog B1) (eg, colon cancer), gene analysis, V600E variant	CPT-4 Procedure
EGFR		
81235	EGFR (epidermal growth factor receptor) (eg, non-small cell lung cancer) gene analysis, common variants (eg, exon 19 LREA deletion, L858R, T790M, G719A, G719S, L861Q)	CPT-4 Procedure
BCR-ABL		
81207	BCR/ABL1 (t(9;22)) (eg, chronic myelogenous leukemia) translocation analysis; minor breakpoint, qualitative or quantitative	CPT-4 Procedure
81206	BCR/ABL1 (t(9;22)) (eg, chronic myelogenous leukemia) translocation analysis; major breakpoint, qualitative or quantitative	CPT-4 Procedure
81208	BCR/ABL1 (t(9;22)) (eg, chronic myelogenous leukemia) translocation analysis; other breakpoint, qualitative or quantitative	CPT-4 Procedure
BRCA		
81211	BRCA1, BRCA2 (breast cancer 1 and 2) (eg, hereditary breast and ovarian cancer) gene analysis; full sequence analysis and common duplication/deletion variants in BRCA1 (ie, exon 13 del 3.835kb, exon 13 dup 6kb, exon 14-20 del 26kb, exon 22 del 510bp, exon 8-9 del 7.1kb)	CPT-4 Procedure
81212	BRCA1, BRCA2 (breast cancer 1 and 2) (eg, hereditary breast and ovarian cancer) gene analysis; 185delAG, 5385insC, 6174delT variants	CPT-4 Procedure
81213	BRCA1, BRCA2 (breast cancer 1 and 2) (eg, hereditary breast and ovarian cancer) gene analysis; uncommon duplication/deletion variants	CPT-4 Procedure
81214	BRCA1 (breast cancer 1) (eg, hereditary breast and ovarian cancer) gene analysis; full sequence analysis and common duplication/deletion variants (ie, exon 13 del 3.835kb, exon 13 dup 6kb, exon 14-20 del 26kb, exon 22 del 510bp, exon 8-9 del 7.1kb)	CPT-4 Procedure
81215	BRCA1 (breast cancer 1) (eg, hereditary breast and ovarian cancer) gene analysis; known familial variant	CPT-4 Procedure
81216	BRCA2 (breast cancer 2) (eg, hereditary breast and ovarian cancer) gene analysis; full sequence analysis	CPT-4 Procedure
81217	BRCA2 (breast cancer 2) (eg, hereditary breast and ovarian cancer) gene analysis; known familial variant	CPT-4 Procedure
S3818	Complete gene sequence analysis; BRCA1 gene	HCPCS Procedure
S3819	Complete gene sequence analysis; BRCA2 gene	HCPCS Procedure
S3820	Complete BRCA1 and BRCA2 gene sequence analysis for susceptibility to breast and ovarian cancer	HCPCS Procedure
S3822	Single mutation analysis (in individual with a known BRCA1 or BRCA2 mutation in the family) for susceptibility to breast and ovarian cancer	HCPCS Procedure
S3823	Three-mutation BRCA1 and BRCA2 analysis for susceptibility to breast and ovarian cancer in Ashkenazi individuals	HCPCS Procedure

Appendix F: Modular Program Specifications for cder_mpl1r_wp021_nsdp_v01

The Cohort Identification and Descriptive Analysis (CIDA) tool, version 2.2.1, will be used to investigate drug initiation among those with a genetic test and/or cancer diagnosis within the 183 days prior to drug initiation. The query period was from January 1, 2013 - current, and the enrollment gap was set at 45 days. Age groups were split as follows: 0-21, 22-44, 45-64, 65+. In total, 20 scenarios were examined in this report.

Enrollment Gap: 45 Days
Age Groups: 0-21, 22-44, 45-64, 65+
Query Period: January 1, 2013 - Current
Coverage Requirement: Medical and Drug

		Drug/Exposure				Inclusion/Exclusion				
Scenario	Enrollment Requirement (days)	Incident exposure	Incident w/ respect to:	Washout (days)	Cohort Definition	Criteria	Include/Exclude	Lookback Start	Lookback End	Caresetting
1	183	Cetuximab, Panitumumab	Cetuximab, Panitumumab	183	01	N/A	N/A	N/A	N/A	N/A
2	183	Cetuximab, Panitumumab	Cetuximab, Panitumumab	183	01	KRAS	Include	-183	0	Any
3	183	Cetuximab, Panitumumab	Cetuximab, Panitumumab	183	01	KRAS + Colorectal Cancer	Include	-183	0	Any
4	183	Cetuximab, Panitumumab	Cetuximab, Panitumumab	183	01	Colorectal Cancer	Include	-183	0	Any
5	183	Trametinib, Dabrafenib, Vemurafenib, Cobimetinib	Trametinib, Dabrafenib, Vemurafenib, Cobimetinib	183	01	N/A	N/A	N/A	N/A	N/A
6	183	Trametinib, Dabrafenib, Vemurafenib, Cobimetinib	Trametinib, Dabrafenib, Vemurafenib, Cobimetinib	183	01	BRAF	Include	-183	0	Any
7	183	Trametinib, Dabrafenib, Vemurafenib, Cobimetinib	Trametinib, Dabrafenib, Vemurafenib, Cobimetinib	183	01	BRAF + Melanoma	Include	-183	0	Any
8	183	Trametinib, Dabrafenib, Vemurafenib, Cobimetinib	Trametinib, Dabrafenib, Vemurafenib, Cobimetinib	183	01	Melanoma	Include	-183	0	Any
9	183	Cetuximab, Panitumumab, Afatinib, Erlotinib, Tagrisso, Gefitinib	Cetuximab, Panitumumab, Afatinib, Erlotinib, Tagrisso, Gefitinib	183	01	N/A	N/A	N/A	N/A	N/A
10	183	Cetuximab, Panitumumab, Afatinib, Erlotinib, Tagrisso, Gefitinib	Cetuximab, Panitumumab, Afatinib, Erlotinib, Tagrisso, Gefitinib	183	01	EGFR	Include	-183	0	Any
11	183	Cetuximab, Panitumumab, Afatinib, Erlotinib, Tagrisso, Gefitinib	Cetuximab, Panitumumab, Afatinib, Erlotinib, Tagrisso, Gefitinib	183	01	EGFR + Colorectal Cancer OR Non-Small Cell Lung Cancer	Include	-183	0	Any
12	183	Cetuximab, Panitumumab, Afatinib, Erlotinib, Tagrisso, Gefitinib	Cetuximab, Panitumumab, Afatinib, Erlotinib, Tagrisso, Gefitinib	183	01	Colorectal Cancer OR Non-Small Cell Lung Cancer	Include	-183	0	Any
13	183	Dasatinib, Imatinib, Bosutinib, Nilotinib, Ponatinib	Dasatinib, Imatinib, Bosutinib, Nilotinib, Ponatinib	183	01	N/A	N/A	-183	0	N/A

14	183	Dasatinib, Imatinib, Bosutinib, Nilotinib, Ponatinib	Dasatinib, Imatinib, Bosutinib, Nilotinib, Ponatinib	183	01	BCR-ABL	Include	-183	0	Any
15	183	Dasatinib, Imatinib, Bosutinib, Nilotinib, Ponatinib	Dasatinib, Imatinib, Bosutinib, Nilotinib, Ponatinib	183	01	BCR-ABL + Leukemia	Include	-183	0	Any
16	183	Dasatinib, Imatinib, Bosutinib, Nilotinib, Ponatinib	Dasatinib, Imatinib, Bosutinib, Nilotinib, Ponatinib	183	01	Leukemia	Include	-183	0	Any
17	183	Olaparib	Olaparib	183	01	N/A	N/A	-183	0	N/A
18	183	Olaparib	Olaparib	183	01	BRCA	Include	-183	0	Any
19	183	Olaparib	Olaparib	183	01	BRCA + Ovarian Cancer	Include	-183	0	Any
20	183	Olaparib	Olaparib	183	01	Ovarian Cancer	Include	-183	0	Any

Note: ICD-9, ICD-10, HCPCS, and CPT codes are provided by Optum360. NDC codes are checked against First Data Bank's “National Drug Data File (NDDF®) Plus”