

Table S1 Detailed information on the allelic fraction (AF) and maximum allelic fraction (maxAF) of all patient samples at various timepoints

| Patient ID | Timepoint | T790M status at baseline | ctDNA clearance status at F1 | Gene      | Mutation type        | Exon number | Variant        | AF     | maxAF status | Sequencing depth   | Chromosome number | Genomic position | dbnsnp_ID   | COSM_ID        | MatchReport    | Reference allele (REF) | Variant allele (ALT) | Population Frequency |   |
|------------|-----------|--------------------------|------------------------------|-----------|----------------------|-------------|----------------|--------|--------------|--------------------|-------------------|------------------|-------------|----------------|----------------|------------------------|----------------------|----------------------|---|
| P01        | baseline  | -                        | -                            | TP53      | missense_variant     | 7           | p.G244S        | 0.68%  | no           | 22576              | 17                | 7577551          | NA          | COSM10941,COSI | G244           | C                      | T                    | 0                    |   |
| P01        | baseline  | present                  | -                            | EGFR      | missense_variant     | 20          | p.T790M        | 2.39%  | no           | 29031              | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 2.84E-05             |   |
| P01        | baseline  | -                        | Yes                          | EGFR      | missense_variant     | 21          | p.L858R        | 3.23%  | maxAF        | 27205              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P01        | FU2,FU3   | -                        | -                            | EGFR      | missense_variant     | 18          | p.L718Q        | 0.90%  | no           | 32010              | 7                 | 55241705         | NA          | NA             | L718           | T                      | A                    | 0                    |   |
| P01        | FU2,FU3   | -                        | -                            | EGFR      | missense_variant     | 21          | p.L858R        | 1.47%  | maxAF        | 32035              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P01        | FU4       | -                        | -                            | EGFR      | missense_variant     | 18          | p.L718Q        | 1.05%  | no           | 28333              | 7                 | 55241705         | NA          | NA             | L718           | T                      | A                    | 0                    |   |
| P01        | FU4       | -                        | -                            | EGFR      | missense_variant     | 21          | p.L858R        | 1.57%  | maxAF        | 29720              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P02        | baseline  | -                        | -                            | TP53      | missense_variant     | 7           | p.M237I        | 2.30%  | no           | 20647              | 17                | 7577570          | rs587782664 | COSM301404,CO  | M237           | C                      | A                    | 0                    |   |
| P02        | baseline  | present                  | -                            | EGFR      | missense_variant     | 20          | p.T790M        | 1.36%  | no           | 30559              | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 2.84E-05             |   |
| P02        | baseline  | -                        | Yes                          | EGFR      | missense_variant     | 21          | p.L858R        | 2.80%  | maxAF        | 30575              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P02        | FU2       | -                        | -                            | EGFR      | missense_variant     | 21          | p.L858R        | 0.12%  | maxAF        | 24837              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P02        | FU3       | -                        | -                            | RB1       | splice_acceptor_vari | 20          | c.1961-1G>C    | 3.54%  | no           | 18631              | 13                | 49033823         | NA          | NA             | Exon20_splice_ | G                      | C                    | 0                    |   |
| P02        | FU3       | -                        | -                            | TP53      | missense_variant     | 7           | p.M237I        | 15.59% | maxAF        | 19934              | 17                | 7577570          | rs587782664 | COSM301404,CO  | M237           | C                      | A                    | 0                    |   |
| P02        | FU3       | -                        | -                            | POM121L12 | missense_variant     | 1           | p.P21H         | 3.75%  | no           | 33976              | 7                 | 53103426         | NA          | NA             | P21            | C                      | A                    | 0                    |   |
| P02        | FU3       | -                        | -                            | EGFR      | missense_variant     | 21          | p.L858R        | 14.52% | no           | 29857              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P02        | FU4       | -                        | -                            | MET       | cn_amp               | NA          | cn_amp         | 2.94   | no           | 0.2;4;13,18,7q31.2 | 7q31.2            | NA               | NA          | NA             | CN_gain        | A                      | 29                   | 29 NA                | 0 |
| P02        | FU4       | -                        | -                            | RB1       | splice_acceptor_vari | 20          | c.1961-1G>C    | 6.35%  | no           | 14868              | 13                | 49033823         | NA          | NA             | Exon20_splice_ | G                      | C                    | 0                    |   |
| P02        | FU4       | -                        | -                            | TP53      | missense_variant     | 7           | p.M237I        | 25.81% | maxAF        | 17421              | 17                | 7577570          | rs587782664 | COSM301404,CO  | M237           | C                      | A                    | 0                    |   |
| P02        | FU4       | -                        | -                            | BRCA1     | missense_variant     | 10          | p.E237K        | 1.16%  | no           | 14264              | 17                | 41246839         | NA          | NA             | E237           | C                      | T                    | 0                    |   |
| P02        | FU4       | -                        | -                            | KIT       | stop_gained          | 11          | p.W557X        | 0.54%  | no           | 16905              | 4                 | 55593605         | NA          | COSM1231       | LOF            | G                      | A                    | 0                    |   |
| P02        | FU4       | -                        | -                            | ROS1      | missense_variant     | 36          | p.L1937F       | 0.74%  | no           | 25984              | 6                 | 117641162        | NA          | NA             | L1937          | G                      | A                    | 0                    |   |
| P02        | FU4       | -                        | -                            | POM121L12 | missense_variant     | 1           | p.P21H         | 7.75%  | no           | 29700              | 7                 | 53103426         | NA          | NA             | P21            | C                      | A                    | 0                    |   |
| P02        | FU4       | -                        | -                            | EGFR      | missense_variant     | 21          | p.L858R        | 19.12% | no           | 24150              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P03        | baseline  | -                        | -                            | OR2T4     | missense_variant     | 1           | p.R171T        | 1.66%  | no           | 47872              | 1                 | 248525394        | NA          | NA             | R171           | G                      | C                    | 0                    |   |
| P03        | baseline  | -                        | -                            | TP53      | missense_variant     | 6           | p.L194R        | 4.93%  | no           | 27897              | 17                | 7578268          | NA          | COSM3403268,C  | L194           | A                      | C                    | 4.06E-06             |   |
| P03        | baseline  | -                        | Yes                          | EGFR      | disruptive_inframe_c | 19          | p.E746_A750del | 24.58% | maxAF        | 38390              | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram  | AGGAATTAAGA            | A                    | 0                    |   |
| P03        | baseline  | present                  | -                            | EGFR      | missense_variant     | 20          | p.T790M        | 16.14% | no           | 52269              | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 2.84E-05             |   |
| P03        | baseline  | -                        | -                            | EGFR      | cn_amp               | NA          | cn_amp         | 2.98   | no           | 0.327;16;16,7p11.2 | 7p11.2            | NA               | NA          | NA             | CN_gain        | C                      | 16                   | 16 NA                | 0 |
| P03        | FU3       | -                        | -                            | OR2T4     | missense_variant     | 1           | p.R171T        | 1.64%  | no           | 39909              | 1                 | 248525394        | NA          | NA             | R171           | G                      | C                    | 0                    |   |
| P03        | FU3       | -                        | -                            | TP53      | missense_variant     | 6           | p.L194R        | 4.90%  | no           | 26758              | 17                | 7578268          | NA          | COSM3403268,C  | L194           | A                      | C                    | 4.06E-06             |   |
| P03        | FU3       | -                        | -                            | EGFR      | disruptive_inframe_c | 19          | p.E746_A750del | 6.78%  | maxAF        | 25515              | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram  | AGGAATTAAGA            | A                    | 0                    |   |
| P03        | FU3       | -                        | -                            | EGFR      | missense_variant     | 20          | p.T790M        | 0.29%  | no           | 35120              | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 4.12E-05             |   |
| P03        | FU3       | -                        | -                            | EGFR      | cn_amp               | NA          | cn_amp         | 2.38   | no           | 0.431;2;42,7p11.2  | 7p11.2            | NA               | NA          | NA             | CN_gain        | C                      | 16                   | 15 NA                | 0 |
| P03        | FU3       | -                        | -                            | MET       | cn_amp               | NA          | cn_amp         | 3.11   | no           | 0.296;7;10,7q31.2  | 7q31.2            | NA               | NA          | NA             | CN_gain        | A                      | 29                   | 29 NA                | 0 |
| P04        | baseline  | -                        | -                            | TP53      | missense_variant     | 7           | p.L257P        | 18.02% | maxAF        | 23264              | 17                | 7577511          | NA          | COSM1386606,C  | L257           | A                      | G                    | 0                    |   |
| P04        | baseline  | -                        | -                            | CDH18     | missense_variant     | 15          | p.S734F        | 4.84%  | no           | 60100              | 5                 | 19473507         | NA          | NA             | S734           | G                      | A                    | 0                    |   |
| P04        | baseline  | -                        | -                            | EGFR      | disruptive_inframe_c | 19          | p.E746_A750del | 15.12% | no           | 30008              | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram  | AGGAATTAAGA            | A                    | 0                    |   |
| P04        | baseline  | present                  | -                            | EGFR      | missense_variant     | 20          | p.T790M        | 7.11%  | no           | 43111              | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 2.84E-05             |   |
| P04        | baseline  | -                        | -                            | EGFR      | cn_amp               | NA          | cn_amp         | 2.52   | no           | 0.151;2;15,7p11.2  | 7p11.2            | NA               | NA          | NA             | CN_gain        | C                      | 16                   | 15 NA                | 0 |
| P04        | FU4       | -                        | -                            | EGFR      | disruptive_inframe_c | 19          | p.E746_A750del | 0.04%  | maxAF        | 27602              | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram  | AGGAATTAAGA            | A                    | 0                    |   |
| P05        | baseline  | -                        | -                            | TP53      | missense_variant     | 7           | p.C242Y        | 1.00%  | no           | 18017              | 17                | 7577556          | rs121912655 | COSM2744612,C  | C242           | C                      | T                    | 0                    |   |
| P05        | baseline  | present                  | Yes                          | EGFR      | missense_variant     | 20          | p.T790M        | 6.05%  | maxAF        | 24327              | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 2.84E-05             |   |
| P05        | baseline  | -                        | -                            | EGFR      | missense_variant     | 21          | p.L858R        | 5.17%  | no           | 26087              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P05        | FU3       | -                        | -                            | EGFR      | missense_variant     | 20          | p.T790M        | 0.31%  | no           | 27326              | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 2.84E-05             |   |
| P05        | FU3       | -                        | -                            | EGFR      | missense_variant     | 21          | p.L858R        | 0.69%  | maxAF        | 26879              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P05        | FU5       | -                        | -                            | EGFR      | missense_variant     | 21          | p.L858R        | 0.14%  | maxAF        | 26420              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P06        | baseline  | -                        | -                            | CTNNB1    | missense_variant     | 3           | p.S37F         | 1.40%  | no           | 35311              | 3                 | 41266113         | rs121913403 | COSM5662       | S37            | C                      | T                    | 0                    |   |
| P06        | baseline  | present                  | -                            | EGFR      | missense_variant     | 20          | p.T790M        | 1.80%  | no           | 33652              | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 2.84E-05             |   |
| P06        | baseline  | -                        | -                            | EGFR      | missense_variant     | 21          | p.L858R        | 2.74%  | maxAF        | 32212              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P06        | baseline  | -                        | -                            | MET       | missense_variant     | 6           | p.D597H        | 1.74%  | no           | 34483              | 7                 | 116395496        | rs769073123 | NA             | D597           | G                      | C                    | 0                    |   |
| P06        | baseline  | -                        | -                            | TP53      | stop_gained          | 10          | p.R342X        | 5.72%  | no           | 17666              | 17                | 7574003          | rs730882029 | COSM99721,COSI | LOF            | G                      | A                    | 0                    |   |
| P08        | baseline  | -                        | -                            | EGFR      | disruptive_inframe_c | 19          | p.E746_A750del | 10.57% | maxAF        | 22956              | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram  | AGGAATTAAGA            | A                    | 0                    |   |
| P08        | baseline  | present                  | -                            | EGFR      | missense_variant     | 20          | p.T790M        | 4.99%  | no           | 32748              | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 2.84E-05             |   |
| P08        | baseline  | -                        | -                            | EGFR      | cn_amp               | NA          | cn_amp         | 2.33   | no           | 0.217;3;11,7p11.2  | 7p11.2            | NA               | NA          | NA             | CN_gain        | C                      | 16                   | 14 NA                | 0 |
| P09        | baseline  | -                        | -                            | APC       | missense_variant     | 16          | p.N827Y        | 1.61%  | no           | 21207              | 5                 | 112173770        | NA          | NA             | N827           | A                      | T                    | 0                    |   |
| P09        | baseline  | -                        | -                            | ERBB2     | cn_amp               | NA          | cn_amp         | 2.38   | no           | 0.424;2;40,17q12   | 17q12             | NA               | NA          | NA             | CN_gain        | A                      | 18                   | 17 NA                | 0 |
| P09        | baseline  | -                        | -                            | RB1       | frameshift_variant   | 2           | p.F57fs        | 25.24% | no           | 22146              | 13                | 48881444         | NA          | NA             | LOF            | G                      | ATTTT                | 0                    |   |
| P09        | baseline  | present                  | -                            | EGFR      | missense_variant     | 20          | p.T790M        | 21.82% | no           | 90465              | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 2.84E-05             |   |
| P09        | baseline  | -                        | Yes                          | EGFR      | missense_variant     | 21          | p.L858R        | 63.94% | maxAF        | 72434              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P09        | baseline  | -                        | -                            | TP53      | missense_variant     | 8           | p.R273L        | 23.65% | no           | 24752              | 17                | 7577120          | rs28934576  | COSM3675521,C  | R273           | C                      | A                    | 4.07E-06             |   |
| P09        | baseline  | -                        | -                            | EGFR      | cn_amp               | NA          | cn_amp         | 5.28   | no           | 0.234;19;14,7p11.2 | 7p11.2            | NA               | NA          | NA             | CN_gain        | C                      | 16                   | 16 NA                | 0 |
| P09        | baseline  | -                        | -                            | CDK6      | cn_amp               | NA          | cn_amp         | 3.99   | no           | 0.273;4;26,7q21.2  | 7q21.2            | NA               | NA          | NA             | CN_gain        | A                      | 7                    | 7 NA                 | 0 |
| P09        | baseline  | -                        | -                            | FGFR1     | cn_del               | NA          | cn_del         | 1.67   | no           | 0.406;2;38,8p11.22 | 8p11.22           | NA               | NA          | NA             | CN_loss        | G                      | 9                    | 7 NA                 | 0 |
| P09        | baseline  | -                        | -                            | MYC       | cn_amp               | NA          | cn_amp         | 2.37   | no           | 0.157;1;16,8q24.21 | 8q24.21           | NA               | NA          | NA             | CN_gain        | A                      | 8                    | 6 NA                 | 0 |
| P09        | FU2       | -                        | -                            | RB1       | frameshift_variant   | 2           | p.F57fs        | 1.41%  | no           | 24385              | 13                | 48881444         | NA          | NA             | LOF            | G                      | ATTTT                | 0                    |   |
| P09        | FU2       | -                        | -                            | TP53      | missense_variant     | 8           | p.R273L        | 1.99%  | no           | 31591              | 17                | 7577120          | rs28934576  | COSM3675521,C  | R273           | C                      | A                    | 4.07E-06             |   |
| P09        | FU2       | -                        | -                            | EGFR      | missense_variant     | 21          | p.L858R        | 7.67%  | maxAF        | 34879              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P09        | FU3       | -                        | -                            | RB1       | frameshift_variant   | 2           | p.F57fs        | 4.35%  | no           | 27756              | 13                | 48881444         | NA          | NA             | LOF            | G                      | ATTTT                | 0                    |   |
| P09        | FU3       | -                        | -                            | TP53      | missense_variant     | 8           | p.R273L        | 6.08%  | no           | 34185              | 17                | 7577120          | rs28934576  | COSM3675521,C  | R273           | C                      | A                    | 4.07E-06             |   |
| P09        | FU3       | -                        | -                            | PIK3CA    | missense_variant     | 10          | p.E545K        | 0.53%  | no           | 13021              | 3                 | 178936091        | rs104886003 | COSM125370,CO  | E545           | G                      | A                    | 4.09E-06             |   |
| P09        | FU3       | -                        | -                            | EGFR      | missense_variant     | 21          | p.L858R        | 21.63% | maxAF        | 43051              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P09        | FU3       | -                        | -                            | EGFR      | cn_amp               | NA          | cn_amp         | 2.55   | no           | 0.367;15;22,7p11.2 | 7p11.2            | NA               | NA          | NA             | CN_gain        | C                      | 16                   | 15 NA                | 0 |
| P09        | FU4       | -                        | -                            | RB1       | frameshift_variant   | 2           | p.F57fs        | 9.08%  | no           | 25021              | 13                | 48881444         | NA          | NA             | LOF            | G                      | ATTTT                | 0                    |   |
| P09        | FU4       | -                        | -                            | TP53      | missense_variant     | 8           | p.R273L        | 8.72%  | no           | 27718              | 17                | 7577120          | rs28934576  | COSM3675521,C  | R273           | C                      | A                    | 4.07E-06             |   |
| P09        | FU4       | -                        | -                            | PIK3CA    | missense_variant     | 10          | p.E545K        | 2.45%  | no           | 12553              | 3                 | 178936091        | rs104886003 | COSM125370,CO  | E545           | G                      | A                    | 4.09E-06             |   |
| P09        | FU4       | -                        | -                            | EGFR      | missense_variant     | 21          | p.L858R        | 22.84% | maxAF        | 36001              | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |   |
| P09        | FU4       | -                        | -                            | EGFR      | cn_amp               | NA          | cn_amp         | 2.51   | no           | 0.353;16;23,7p11.2 | 7p11.2            | NA               | NA          | NA             | CN_gain        | C                      | 16                   | 15 NA                | 0 |
| P09        | FU5       | -                        | -                            | RB1       | frameshift_variant   | 2           | p.F57fs        | 61.05% | no           | 27184              | 13                | 48881444         | NA          | NA             | LOF            | G                      | ATTTT                | 0                    |   |
| P09        | FU5       | -                        | -                            | TP53      | missense_variant     | 8           | p.R273L        | 56.94% | no           | 25288              | 17                |                  |             |                |                |                        |                      |                      |   |

| Patient ID | Timepoint   | T790M status at baseline | ctDNA clearance status at F1 | Gene   | Mutation type        | Exon number | Variant                | AF     | maxAF status | Sequencing depth | Chromosome number | Genomic position | dbSNP_ID    | COSM_ID        | MatchReport    | Reference allele (REF) | Variant allele (ALT) | Population Frequency |
|------------|-------------|--------------------------|------------------------------|--------|----------------------|-------------|------------------------|--------|--------------|------------------|-------------------|------------------|-------------|----------------|----------------|------------------------|----------------------|----------------------|
| P12        | FU1,FU2,FU3 | -                        | -                            | RB1    | splice_donor_variant | 13          | c.1332+1dup            | 0.52%  | no           | 24369            | 13                | 48951169         | NA          | NA             | Exon13_splice_ | A                      | AG                   | 0                    |
| P12        | FU1,FU2,FU3 | -                        | -                            | TP53   | missense_variant     | 6           | p.H214R                | 0.86%  | no           | 35738            | 17                | 7578208          | NA          | COSM3388198,C1 | H214           | T                      | C                    | 0                    |
| P12        | FU1,FU2,FU3 | -                        | -                            | APC    | missense_variant     | 16          | p.N1533S               | 0.63%  | no           | 19222            | 5                 | 112175889        | NA          | NA             | N1533          | A                      | G                    | 0                    |
| P12        | FU1,FU2,FU3 | -                        | No                           | EGFR   | missense_variant     | 21          | p.L858R                | 2.74%  | maxAF        | 36293            | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |
| P12        | FU4         | -                        | -                            | RB1    | splice_donor_variant | 13          | c.1332+1dup            | 3.24%  | no           | 28451            | 13                | 48951169         | NA          | NA             | Exon13_splice_ | A                      | AG                   | 0                    |
| P12        | FU4         | -                        | -                            | TP53   | missense_variant     | 6           | p.H214R                | 4.97%  | no           | 33603            | 17                | 7578208          | NA          | COSM3388198,C1 | H214           | T                      | C                    | 0                    |
| P12        | FU4         | -                        | -                            | APC    | missense_variant     | 16          | p.N1533S               | 3.05%  | no           | 19588            | 5                 | 112175889        | NA          | NA             | N1533          | A                      | G                    | 0                    |
| P12        | FU4         | -                        | -                            | EGFR   | missense_variant     | 21          | p.L858R                | 9.55%  | maxAF        | 40041            | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |
| P12        | FU4         | -                        | -                            | MET    | cn_amp               | NA          |                        | 2.56   | no           | 0.369;15;10      | 7q31.2            | 7q31.2           | NA          | NA             | cn_gain        |                        | 29                   | 29 NA                |
| P12        | FU5         | -                        | -                            | RB1    | splice_donor_variant | 13          | c.1332+1dup            | 11.11% | no           | 22012            | 13                | 48951169         | NA          | NA             | Exon13_splice_ | A                      | AG                   | 0                    |
| P12        | FU5         | -                        | -                            | TP53   | missense_variant     | 6           | p.H214R                | 20.02% | no           | 27661            | 17                | 7578208          | NA          | COSM3388198,C1 | H214           | T                      | C                    | 0                    |
| P12        | FU5         | -                        | -                            | APC    | missense_variant     | 16          | p.N1533S               | 11.27% | no           | 15977            | 5                 | 112175889        | NA          | NA             | N1533          | A                      | G                    | 0                    |
| P12        | FU5         | -                        | -                            | EGFR   | missense_variant     | 21          | p.L858R                | 25.73% | maxAF        | 35610            | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |
| P12        | FU5         | -                        | -                            | EGFR   | cn_amp               | NA          |                        | 2.37   | no           | 0.371;9;26;1     | 7p11.2            | 7p11.2           | NA          | NA             | cn_gain        |                        | 16                   | 15 NA                |
| P14        | baseline    | -                        | -                            | PIK3CA | missense_variant     | 10          | p.E542K                | 0.30%  | maxAF        | 11681            | 3                 | 178936082        | rs121913273 | COSM760,COSM1  | E542           | G                      | A                    | 0                    |
| P14        | baseline    | absent                   | -                            | EGFR   | disruptive_inframe_c | 19          | p.E746_A750del         | 0.25%  | no           | 22128            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram  | AGGAATTAAGA            | A                    | 0                    |
| P14        | FU1,FU2,FU3 | -                        | No                           | PIK3CA | missense_variant     | 10          | p.E542K                | 0.12%  | maxAF        | 11720            | 3                 | 178936082        | rs121913273 | COSM760,COSM1  | E542           | G                      | A                    | 0                    |
| P14        | FU1,FU2,FU3 | -                        | -                            | EGFR   | disruptive_inframe_c | 19          | p.E746_A750del         | 0.07%  | no           | 24936            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram  | AGGAATTAAGA            | A                    | 0                    |
| P14        | FU4         | -                        | -                            | TP53   | missense_variant     | 8           | p.E286K                | 0.65%  | no           | 30634            | 17                | 7577082          | rs786201059 | COSM99924,COSI | E286           | C                      | T                    | 0                    |
| P14        | FU4         | -                        | -                            | PIK3CA | missense_variant     | 10          | p.E542K                | 1.24%  | maxAF        | 10319            | 3                 | 178936082        | rs121913273 | COSM760,COSM1  | E542           | G                      | A                    | 0                    |
| P14        | FU4         | -                        | -                            | EGFR   | disruptive_inframe_c | 19          | p.E746_A750del         | 0.24%  | no           | 20313            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram  | AGGAATTAAGA            | A                    | 0                    |
| P14        | FU5         | -                        | -                            | TP53   | missense_variant     | 8           | p.E286K                | 4.10%  | no           | 33303            | 17                | 7577082          | rs786201059 | COSM99924,COSI | E286           | C                      | T                    | 0                    |
| P14        | FU5         | -                        | -                            | TP53   | missense_variant     | 8           | p.R273C                | 0.15%  | no           | 24216            | 17                | 7577121          | rs121913343 | COSM1645518,C1 | R273           | G                      | A                    | 5.81E-05             |
| P14        | FU5         | -                        | -                            | PIK3CA | missense_variant     | 10          | p.E542K                | 5.93%  | maxAF        | 12277            | 3                 | 178936082        | rs121913273 | COSM760,COSM1  | E542           | G                      | A                    | 0                    |
| P14        | FU5         | -                        | -                            | EGFR   | disruptive_inframe_c | 19          | p.E746_A750del         | 1.07%  | no           | 20405            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram  | AGGAATTAAGA            | A                    | 0                    |
| P15        | baseline    | -                        | -                            | TP53   | stop_gained          | 8           | p.R306X                | 2.48%  | no           | 35095            | 17                | 7577022          | rs121913344 | COSM1640820,C1 | LOF            | G                      | A                    | 0                    |
| P15        | baseline    | -                        | -                            | EGFR   | conservative_infram  | 19          | p.E746_A750del         | 11.00% | maxAF        | 28628            | 7                 | 55242465         | rs727504233 | COSM6225       | Exon19_infram  | GGAATTAAGAG            | G                    | 0                    |
| P15        | baseline    | present                  | -                            | EGFR   | missense_variant     | 20          | p.T790M                | 4.23%  | no           | 38279            | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 2.84E-05             |
| P15        | baseline    | -                        | -                            | EGFR   | cn_amp               | NA          |                        | 2.42   | no           | 0.353;11;11      | 7p11.2            | 7p11.2           | NA          | NA             | cn_gain        |                        | 16                   | 15 NA                |
| P15        | baseline    | -                        | -                            | CSMD3  | cn_amp               | NA          |                        | 2.51   | no           | NA;0;NA          | 8q23.3            | 8q23.3           | NA          | NA             | cn_gain        |                        | 5                    | 5 NA                 |
| P16        | baseline    | -                        | -                            | CND1   | cn_amp               | NA          |                        | 2.36   | no           | 0.434;4;42;1     | 11q13.3           | 11q13.3          | NA          | NA             | cn_gain        |                        | 6                    | 3 NA                 |
| P16        | baseline    | -                        | -                            | FGFR19 | cn_amp               | NA          |                        | 2.39   | no           | 0.421;2;42;1     | 11q13.3           | 11q13.3          | NA          | NA             | cn_gain        |                        | 5                    | 4 NA                 |
| P16        | baseline    | -                        | -                            | TP53   | missense_variant     | 7           | p.R249S                | 13.38% | no           | 18106            | 17                | 7577534          | NA          | COSM1679495,C1 | R249           | C                      | G                    | 0                    |
| P16        | baseline    | -                        | -                            | EGFR   | disruptive_inframe_c | 19          | p.L747_A750delir       | 43.02% | maxAF        | 38315            | 7                 | 55242465         | rs121913436 | NA             | Exon19_infram  | GGAATTAAGAG            | GGAAC                | 0                    |
| P16        | baseline    | present                  | -                            | EGFR   | missense_variant     | 20          | p.T790M                | 20.20% | no           | 54903            | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 2.84E-05             |
| P16        | baseline    | -                        | -                            | EGFR   | cn_amp               | NA          |                        | 3.25   | no           | 0.306;17;20      | 7p11.2            | 7p11.2           | NA          | NA             | cn_gain        |                        | 16                   | 16 NA                |
| P16        | FU1         | -                        | -                            | TP53   | missense_variant     | 7           | p.R249S                | 1.74%  | no           | 21058            | 17                | 7577534          | NA          | COSM1679495,C1 | R249           | C                      | G                    | 0                    |
| P16        | FU1         | -                        | No                           | EGFR   | disruptive_inframe_c | 19          | p.L747_A750delir       | 7.06%  | maxAF        | 24323            | 7                 | 55242465         | rs121913436 | NA             | Exon19_infram  | GGAATTAAGAG            | GGAAC                | 0                    |
| P16        | FU2,FU3     | -                        | -                            | TP53   | missense_variant     | 7           | p.R249S                | 9.87%  | no           | 18809            | 17                | 7577534          | NA          | COSM1679495,C1 | R249           | C                      | G                    | 0                    |
| P16        | FU2,FU3     | -                        | -                            | BRAF   | missense_variant     | 15          | p.V600E                | 1.82%  | no           | 23717            | 7                 | 140453136        | rs113488022 | COSM476        | V600           | A                      | T                    | 4.07E-06             |
| P16        | FU2,FU3     | -                        | -                            | EGFR   | disruptive_inframe_c | 19          | p.L747_A750delir       | 22.18% | maxAF        | 25558            | 7                 | 55242465         | rs121913436 | NA             | Exon19_infram  | GGAATTAAGAG            | GGAAC                | 0                    |
| P16        | FU2,FU3     | -                        | -                            | EGFR   | cn_amp               | NA          |                        | 2.51   | no           | 0.361;14;22      | 7p11.2            | 7p11.2           | NA          | NA             | cn_gain        |                        | 16                   | 15 NA                |
| P16        | FU2,FU3     | -                        | -                            | BRAF   | cn_amp               | NA          |                        | 2.71   | no           | NA;0;NA          | 7q34              | 7q34             | NA          | NA             | cn_gain        |                        | 5                    | 5 NA                 |
| P16        | FU4         | -                        | -                            | TP53   | missense_variant     | 7           | p.R249S                | 17.48% | no           | 19724            | 17                | 7577534          | NA          | COSM1679495,C1 | R249           | C                      | G                    | 0                    |
| P16        | FU4         | -                        | -                            | BRAF   | missense_variant     | 15          | p.V600E                | 0.34%  | no           | 29687            | 7                 | 140453136        | rs113488022 | COSM476        | V600           | A                      | T                    | 4.07E-06             |
| P16        | FU4         | -                        | -                            | EGFR   | disruptive_inframe_c | 19          | p.L747_A750delir       | 30.59% | maxAF        | 26853            | 7                 | 55242465         | rs121913436 | NA             | Exon19_infram  | GGAATTAAGAG            | GGAAC                | 0                    |
| P16        | FU4         | -                        | -                            | EGFR   | cn_amp               | NA          |                        | 2.64   | no           | 0.357;15;31      | 7p11.2            | 7p11.2           | NA          | NA             | cn_gain        |                        | 16                   | 16 NA                |
| P16        | FU4         | -                        | -                            | BRAF   | cn_amp               | NA          |                        | 3.34   | no           | NA;0;NA          | 7q34              | 7q34             | NA          | NA             | cn_gain        |                        | 5                    | 5 NA                 |
| P17        | baseline    | -                        | -                            | EGFR   | conservative_infram  | 19          | p.E746_A750del         | 0.15%  | maxAF        | 30589            | 7                 | 55242465         | rs727504233 | COSM6225       | Exon19_infram  | GGAATTAAGAG            | G                    | 0                    |
| P18        | baseline    | present                  | -                            | EGFR   | missense_variant     | 20          | p.T790M                | 0.20%  | no           | 29204            | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 2.84E-05             |
| P18        | baseline    | -                        | -                            | EGFR   | missense_variant     | 21          | p.L858R                | 0.44%  | maxAF        | 30412            | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |
| P19        | baseline    | -                        | No                           | EGFR   | disruptive_inframe_c | 19          | p.L747_T751del         | 0.07%  | maxAF        | 27378            | 7                 | 55242467         | rs121913442 | COSM23571      | Exon19_infram  | AATTAAGAGAA            | A                    | 0                    |
| P19        | FU1,FU2     | -                        | -                            | EGFR   | disruptive_inframe_c | 19          | p.L747_T751del         | 0.02%  | maxAF        | 30136            | 7                 | 55242467         | rs121913442 | COSM23571      | Exon19_infram  | AATTAAGAGAA            | A                    | 0                    |
| P20        | baseline    | -                        | -                            | PIK3CA | missense_variant     | 21          | p.H1047L               | 1.10%  | no           | 16705            | 3                 | 178952085        | rs121913279 | COSM94987      | H1047          | A                      | T                    | 0                    |
| P20        | baseline    | present                  | -                            | EGFR   | missense_variant     | 20          | p.T790M                | 1.21%  | no           | 5706             | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 0                    |
| P20        | baseline    | -                        | -                            | EGFR   | missense_variant     | 21          | p.L858R                | 3.88%  | maxAF        | 12315            | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |
| P20        | baseline    | -                        | -                            | CTNNB1 | missense_variant     | 3           | p.A13S                 | 0.97%  | no           | 12770            | 3                 | 41266040         | NA          | NA             | A13            | G                      | T                    | 0                    |
| P20        | FU1,FU2     | -                        | No                           | EGFR   | missense_variant     | 21          | p.L858R                | 0.19%  | maxAF        | 13890            | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |
| P20        | FU4         | -                        | -                            | EGFR   | missense_variant     | 21          | p.L858R                | 0.28%  | maxAF        | 39605            | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |
| P22        | baseline    | absent                   | -                            | EGFR   | missense_variant     | 21          | p.L858R                | 36.07% | maxAF        | 17421            | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |
| P22        | baseline    | -                        | -                            | TP53   | missense_variant     | 8           | p.R273L                | 13.86% | no           | 12594            | 17                | 7577120          | NA          | COSM3675521    | R273           | C                      | A                    | 0                    |
| P22        | baseline    | -                        | -                            | BRCA1  | splice_region_varian | 11          | NA                     | 1.75%  | no           | 11446            | 17                | 41242938         | NA          | NA             | Exon11_splice_ | GCACA                  | GCA                  | 0                    |
| P22        | baseline    | -                        | -                            | FGFR1  | intergenic_in        | 19          | LINC00662-FGFR1 fusion | 3.06%  | no           | 15851            | 19:2773210        | 19:27732102_8:   | NA          | NA             | Intron3_fusion | 11;13;33;34            | 2;7;2;12             | NA                   |
| P23        | baseline    | present                  | -                            | EGFR   | missense_variant     | 20          | p.T790M                | 1.29%  | no           | 6434             | 7                 | 55249071         | rs121434569 | COSM6240       | T790           | C                      | T                    | 0                    |
| P23        | baseline    | -                        | -                            | EGFR   | missense_variant     | 21          | p.L858R                | 2.95%  | maxAF        | 15314            | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |
| P24        | baseline    | absent                   | -                            | EGFR   | missense_variant     | 21          | p.L858R                | 0.36%  | maxAF        | 14968            | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |
| P24        | FU1,FU2     | -                        | -                            | PIK3CA | missense_variant     | 10          | p.E545K                | 0.24%  | no           | 6713             | 3                 | 178936091        | rs104886003 | COSM763        | E545           | G                      | A                    | 0                    |
| P24        | FU1,FU2     | -                        | No                           | EGFR   | missense_variant     | 21          | p.L858R                | 0.25%  | maxAF        | 10673            | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |
| P24        | FU3         | -                        | -                            | TP53   | missense_variant     | 6           | p.H193R                | 2.55%  | no           | 31811            | 17                | 7578271          | rs786201838 | COSM1740322,C1 | H193           | T                      | C                    | 0                    |
| P24        | FU3         | -                        | -                            | PIK3CA | missense_variant     | 10          | p.E545K                | 0.56%  | no           | 10623            | 3                 | 178936091        | rs104886003 | COSM125370,CO  | E545           | G                      | A                    | 8.34E-06             |
| P24        | FU3         | -                        | -                            | PIK3CA | missense_variant     | 10          | p.Q546P                | 0.58%  | no           | 10778            | 3                 | 178936095        | NA          | COSM767,COSM1  | Q546           | A                      | C                    | 0                    |
| P24        | FU3         | -                        | -                            | EGFR   | missense_variant     | 21          | p.L858R                | 4.23%  | maxAF        | 35753            | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |
| P24        | FU4         | -                        | -                            | TBX3   | missense_variant     | 7           | p.T446P                | 2.21%  | no           | 2260             | 12                | 115112404        | NA          | NA             | H446           | T                      | G                    | 0                    |
| P24        | FU4         | -                        | -                            | TP53   | missense_variant     | 6           | p.H193R                | 3.28%  | no           | 7017             | 17                | 7578271          | rs786201838 | COSM1740322,C1 | H193           | T                      | C                    | 0                    |
| P24        | FU4         | -                        | -                            | PIK3CA | missense_variant     | 10          | p.E545K                | 0.66%  | no           | 7745             | 3                 | 178936091        | rs104886003 | COSM125370,CO  | E545           | G                      | A                    | 4.09E-06             |
| P24        | FU4         | -                        | -                            | PIK3CA | missense_variant     | 10          | p.Q546P                | 0.57%  | no           | 7907             | 3                 | 178936095        | NA          | COSM767,COSM1  | Q546           | A                      | C                    | 0                    |
| P24        | FU4         | -                        | -                            | EGFR   | missense_variant     | 21          | p.L858R                | 5.70%  | maxAF        | 8353             | 7                 | 55259515         | rs121434568 | COSM6224       | L858           | T                      | G                    | 0                    |
| P25        | baseline    | -                        | Yes                          | EGFR   | conservative_infram  | 19          | p.E746_A750del         | 9.36%  | maxAF        | 17187            | 7                 | 55242465         | NA          | COSM6225       | Exon19_infram  | GGAATTAAGAG            | G                    | 0                    |
| P25        | baseline    | present                  | -                            | EGFR   | missense_variant     | 20          | p.T79                  |        |              |                  |                   |                  |             |                |                |                        |                      |                      |

| Patient ID | Timepoint   | T790M status at baseline | ctDNA clearance status at F1 | Gene     | Mutation type        | Exon number  | Variant          | AF     | maxAF status | Sequencing depth | Chromosome number | Genomic position | dbsnp_ID    | COSM_ID         | MatchReport     | Reference allele (REF)          | Variant allele (ALT) | Population Frequency |
|------------|-------------|--------------------------|------------------------------|----------|----------------------|--------------|------------------|--------|--------------|------------------|-------------------|------------------|-------------|-----------------|-----------------|---------------------------------|----------------------|----------------------|
| P26        | FU4         | -                        | -                            | EGFR     | conservative_infram  | 19           | p.E746_A750del   | 5.74%  | no           | 11686            | 7                 | 55242465         | NA          | COSM6225        | Exon19_infram   | GGAATTAAGAG AAGCA               | G                    | 0                    |
| P26        | FU4         | -                        | -                            | RB1      | missense_variant     | 6            | p.V190M          | 24.25% | no           | 8411             | 13                | 48923120         | NA          | NA              | V190            | C                               | A                    | 0                    |
| P26        | FU4         | -                        | -                            | TP53     | missense_variant     | 6            | p.Y220C          | 24.74% | maxAF        | 13081            | 17                | 7578190          | NA          | COSM99720       | Y220            | T                               | C                    | 0                    |
| P26        | FU4         | -                        | -                            | PTEN     | frameshift_variant   | 8            | p.T319fs         | 14.55% | no           | 10165            | 10                | 89720798         | rs146650273 | COSM4898        | LOF             | G                               | T                    | 0                    |
| P26        | FU4         | -                        | -                            | RB1      | frameshift_variant   | 6            | p.V193fs         | 23.65% | no           | 8848             | 13                | 48923124         | NA          | NA              | LOF             | TA                              | G                    | 0                    |
| P26        | FU4         | -                        | -                            | MYC      | cn_amp               | NA           | cn_amp           | 2.73   | no           | NA               | 8q24.21           | 8q24.21          | NA          | NA              | CN_gain         |                                 |                      | 20 NA                |
| P27        | baseline    | absent                   | -                            | EGFR     | missense_variant     | 21           | p.L858R          | 0.69%  | maxAF        | 13674            | 7                 | 55259515         | rs121434568 | COSM6224        | L858            | T                               | G                    | 0                    |
| P28        | baseline    | -                        | -                            | EGFR     | disruptive_inframe_c | 19           | p.E746_A750del   | 1.13%  | maxAF        | 17487            | 7                 | 55242464         | rs121913421 | COSM6223        | Exon19_infram   | AGGAATTAAGA GAAGC               | A                    | 0                    |
| P28        | baseline    | present                  | -                            | EGFR     | missense_variant     | 20           | p.T790M          | 0.18%  | no           | 7743             | 7                 | 55249071         | rs121434569 | COSM6240        | T790            | C                               | T                    | 0                    |
| P28        | baseline    | -                        | -                            | TP53     | missense_variant     | 7            | p.R248Q          | 0.31%  | no           | 12989            | 17                | 7577538          | rs11540652  | COSM99602       | R248            | C                               | T                    | 2.00E-04             |
| P29        | baseline    | present                  | -                            | EGFR     | missense_variant     | 20           | p.T790M          | 0.34%  | no           | 7727             | 7                 | 55249071         | rs121434569 | COSM6240        | T790            | C                               | T                    | 0                    |
| P29        | baseline    | -                        | -                            | EGFR     | missense_variant     | 21           | p.L858R          | 1.48%  | maxAF        | 15237            | 7                 | 55259515         | rs121434568 | COSM6224        | L858            | T                               | G                    | 0                    |
| P29        | baseline    | -                        | -                            | TP53     | missense_variant     | 8            | p.R273H          | 0.32%  | no           | 15475            | 17                | 7577120          | rs28934576  | COSM99729       | R273            | C                               | T                    | 2.00E-04             |
| P29        | baseline    | -                        | -                            | CDKN2A   | missense_variant     | 2            | p.H83Y           | 0.52%  | no           | 8660             | 9                 | 21971111         | rs121913385 | COSM99724       | H83             | G                               | A                    | 0                    |
| P29        | FU1,FU2     | -                        | No                           | TP53     | missense_variant     | 8            | p.R273H          | 0.32%  | maxAF        | 13935            | 17                | 7577120          | rs28934576  | COSM99729       | R273            | C                               | T                    | 2.00E-04             |
| P29        | FU3         | -                        | -                            | TP53     | missense_variant     | 8            | p.R273H          | 0.46%  | maxAF        | 32289            | 17                | 7577120          | rs28934576  | COSM10660, COSI | R273            | C                               | T                    | 2.00E-04             |
| P29        | FU3         | -                        | -                            | EGFR     | missense_variant     | 21           | p.L858R          | 0.20%  | no           | 35364            | 7                 | 55259515         | rs121434568 | COSM6224        | L858            | T                               | G                    | 0                    |
| P29        | FU4         | -                        | -                            | TP53     | missense_variant     | 8            | p.R273H          | 0.37%  | no           | 21072            | 17                | 7577120          | rs28934576  | COSM10660, COSI | R273            | C                               | T                    | 2.00E-04             |
| P29        | FU4         | -                        | -                            | TP53     | missense_variant     | 7            | p.L257V          | 2.72%  | no           | 16503            | 17                | 7577512          | NA          | COSM3742462, CI | L257            | G                               | C                    | 0                    |
| P29        | FU4         | -                        | -                            | EGFR     | missense_variant     | 21           | p.L858R          | 6.39%  | maxAF        | 27370            | 7                 | 55259515         | rs121434568 | COSM6224        | L858            | T                               | G                    | 0                    |
| P29        | FU4         | -                        | -                            | CDKN2A   | missense_variant     | 2            | p.H83Y           | 1.60%  | no           | 10872            | 9                 | 21971111         | rs121913385 | COSM1650884, CI | H83             | G                               | A                    | 0                    |
| P30        | baseline    | -                        | -                            | EGFR     | disruptive_inframe_c | 19           | p.L747_T751del   | 10.02% | no           | 15221            | 7                 | 55242467         | rs121913442 | COSM392194      | Exon19_infram   | AATTAAGAGAA GCAAC               | A                    | 0                    |
| P30        | baseline    | present                  | -                            | EGFR     | missense_variant     | 20           | p.T790M          | 4.10%  | no           | 7740             | 7                 | 55249071         | rs121434569 | COSM6240        | T790            | C                               | T                    | 0                    |
| P30        | baseline    | -                        | -                            | TP53     | missense_variant     | 8            | p.R273H          | 10.35% | maxAF        | 14014            | 17                | 7577120          | rs28934576  | COSM99729       | R273            | C                               | T                    | 2.00E-04             |
| P30        | baseline    | -                        | -                            | LRP1B    | missense_variant     | 22           | p.L1156P         | 5.83%  | no           | 19490            | 2                 | 141665499        | NA          | NA              | L1156           | A                               | G                    | 0                    |
| P30        | baseline    | -                        | -                            | ROS1     | missense_variant     | 34           | p.I1866S         | 6.66%  | no           | 12441            | 6                 | 117645539        | NA          | NA              | I1866           | A                               | C                    | 0                    |
| P30        | baseline    | -                        | -                            | NOTCH1   | conservative_infram  | 34           | p.P2415del       | 2.13%  | no           | 12114            | 9                 | 139390944        | NA          | NA              | Exon34_infram   | TGTGGTG                         | TGTG                 | 0                    |
| P31        | baseline    | -                        | Yes                          | EGFR     | disruptive_inframe_c | 19           | p.E746_A750del   | 2.62%  | maxAF        | 20199            | 7                 | 55242464         | rs121913421 | COSM6223        | Exon19_infram   | AGGAATTAAGA GAAGC               | A                    | 0                    |
| P31        | baseline    | present                  | -                            | EGFR     | missense_variant     | 20           | p.T790M          | 1.70%  | no           | 9085             | 7                 | 55249071         | rs121434569 | COSM6240        | T790            | C                               | T                    | 0                    |
| P31        | baseline    | -                        | -                            | TP53     | splice_donor_variant | 9            | c.993>2T>C       | 1.74%  | no           | 25362            | 17                | 7576851          | NA          | COSM45552       | Exon9_splice_si | A                               | G                    | 0                    |
| P31        | baseline    | -                        | -                            | STK11    | missense_variant     | 4            | p.Q159H          | 0.79%  | no           | 13392            | 19                | 1220384          | NA          | NA              | Q159            | G                               | C                    | 0                    |
| P31        | FU3         | -                        | -                            | EGFR     | disruptive_inframe_c | 19           | p.E746_A750del   | 0.22%  | maxAF        | 20663            | 7                 | 55242464         | rs121913421 | COSM6223        | Exon19_infram   | AGGAATTAAGA GAAGC               | A                    | 0                    |
| P31        | FU3         | -                        | -                            | EGFR     | missense_variant     | 20           | p.T790M          | 0.18%  | no           | 27568            | 7                 | 55249071         | rs121434569 | COSM6240        | T790            | C                               | T                    | 4.12E-05             |
| P31        | FU4         | -                        | -                            | TP53     | splice_donor_variant | 9            | c.993>2T>C       | 0.57%  | maxAF        | 8666             | 17                | 7576851          | NA          | COSM4170831, CI | Exon9_splice_si | A                               | G                    | 0                    |
| P31        | FU4         | -                        | -                            | EGFR     | disruptive_inframe_c | 19           | p.E746_A750del   | 0.28%  | no           | 11782            | 7                 | 55242464         | rs121913421 | COSM6223        | Exon19_infram   | AGGAATTAAGA GAAGC               | A                    | 0                    |
| P32        | baseline    | -                        | -                            | EGFR     | disruptive_inframe_c | 19           | p.E746_A750del   | 3.96%  | no           | 12505            | 7                 | 55242464         | rs121913421 | COSM6223        | Exon19_infram   | AGGAATTAAGA GAAGC               | A                    | 0                    |
| P32        | baseline    | present                  | -                            | EGFR     | missense_variant     | 20           | p.T790M          | 2.86%  | no           | 6714             | 7                 | 55249071         | rs121434569 | COSM6240        | T790            | C                               | T                    | 0                    |
| P32        | baseline    | -                        | -                            | TP53     | splice_region_varian | 10           | p.I332S          | 4.18%  | no           | 6835             | 17                | 7574032          | NA          | NA              | Exon10_splice_  | A                               | C                    | 0                    |
| P32        | baseline    | -                        | -                            | CTNNB1   | missense_variant     | 3            | p.S33F           | 4.36%  | maxAF        | 15476            | 3                 | 41266101         | rs121913400 | COSM5669        | S33             | C                               | T                    | 0                    |
| P32        | baseline    | -                        | -                            | CDKN2A   | frameshift_variant   | 1            | p.Y44fs          | 3.04%  | no           | 10878            | 9                 | 21974694         | NA          | COSM4448825     | LOF             | CG                              | C                    | 0                    |
| P32        | baseline    | -                        | -                            | KRAS     | cn_amp               | NA           | cn_amp           | 2.46   | no           | NA               | 12p12.1           | 12p12.1          | NA          | NA              | CN_gain         |                                 |                      | 5 NA                 |
| P32        | FU1,FU2     | -                        | -                            | EGFR     | disruptive_inframe_c | 19           | p.E746_A750del   | 0.22%  | maxAF        | 17713            | 7                 | 55242464         | rs121913421 | COSM6223        | Exon19_infram   | AGGAATTAAGA GAAGC               | A                    | 0                    |
| P32        | FU3         | -                        | -                            | EGFR     | disruptive_inframe_c | 19           | p.E746_A750del   | 0.16%  | maxAF        | 26196            | 7                 | 55242464         | rs121913421 | COSM6223        | Exon19_infram   | AGGAATTAAGA GAAGC               | A                    | 0                    |
| P32        | FU4         | -                        | -                            | EGFR     | disruptive_inframe_c | 19           | p.E746_A750del   | 0.23%  | maxAF        | 11276            | 7                 | 55242464         | rs121913421 | COSM6223        | Exon19_infram   | AGGAATTAAGA GAAGC               | A                    | 0                    |
| P33        | baseline    | present                  | -                            | EGFR     | missense_variant     | 20           | p.T790M          | 11.65% | no           | 11174            | 7                 | 55249071         | rs121434569 | COSM6240        | T790            | C                               | T                    | 0                    |
| P33        | baseline    | -                        | -                            | TP53     | missense_variant     | 21           | p.L858R          | 43.02% | no           | 17728            | 7                 | 55259515         | rs121434568 | COSM6224        | L858            | T                               | G                    | 0                    |
| P33        | baseline    | -                        | -                            | RB1      | stop_gained          | 13           | p.K420X          | 46.74% | maxAF        | 13958            | 13                | 48951096         | NA          | NA              | LOF             | A                               | T                    | 0                    |
| P33        | baseline    | -                        | -                            | TP53     | splice_region_varian | 8            | NA               | 11.46% | no           | 14890            | 17                | 7576943          | NA          | NA              | Exon8_splice_si | GAGGCATAACT C                   | G                    | 0                    |
| P33        | baseline    | -                        | -                            | EGFR     | cn_amp               | NA           | cn_amp           | 2.32   | no           | NA               | 7p11.2            | 7p11.2           | NA          | NA              | CN_gain         |                                 |                      | 20                   |
| P33        | FU1,FU2     | -                        | No                           | RB1      | stop_gained          | 13           | p.K420X          | 0.98%  | maxAF        | 17364            | 13                | 48951096         | NA          | NA              | LOF             | A                               | T                    | 0                    |
| P33        | FU1,FU2     | -                        | -                            | EGFR     | missense_variant     | 21           | p.L858R          | 0.70%  | no           | 30904            | 7                 | 55259515         | rs121434568 | COSM6224        | L858            | T                               | G                    | 0                    |
| P33        | FU3         | -                        | -                            | KRTAP5-5 | missense_variant     | 1            | p.S151C          | 0.66%  | no           | 14552            | 11                | 1651522          | NA          | NA              | S151            | C                               | G                    | 0                    |
| P33        | FU3         | -                        | -                            | ERBB2    | cn_amp               | NA           | cn_amp           | 2.34   | no           | NA               | 0,0;NA            | 17q12            | 17q12       | NA              | NA              | CN_gain                         |                      | 18                   |
| P33        | FU3         | -                        | -                            | ALK      | fusion               | Intron13_int | EML4-ALK         | 0.08%  | no           | 44040            | 2:42523448        | 2:42523448_2_2   | NA          | NA              | fusion          | 1;0;1,2                         | 1;0;0,1              | 0                    |
| P33        | FU3         | -                        | -                            | RB1      | stop_gained          | 13           | p.K420X          | 35.42% | maxAF        | 19473            | 13                | 48951096         | NA          | NA              | LOF             | A                               | T                    | 0                    |
| P33        | FU3         | -                        | -                            | EGFR     | missense_variant     | 20           | p.T790M          | 5.20%  | no           | 37292            | 7                 | 55249071         | rs121434569 | COSM6240        | T790            | C                               | T                    | 4.12E-05             |
| P33        | FU3         | -                        | -                            | EGFR     | missense_variant     | 20           | p.C797S          | 0.21%  | no           | 37960            | 7                 | 55249092         | NA          | COSM5945664     | C797(in-cis_T79 | G                               | C                    | 0                    |
| P33        | FU3         | -                        | -                            | EGFR     | missense_variant     | 21           | p.L858R          | 25.15% | no           | 35795            | 7                 | 55259515         | rs121434568 | COSM6224        | L858            | T                               | G                    | 0                    |
| P33        | FU4         | -                        | -                            | PTEN     | cn_del               | NA           | cn_del           | 1.49   | no           | 0.179;3,17;1     | 10q23.31          | 10q23.31         | NA          | NA              | CN_loss         |                                 |                      | 7 NA                 |
| P33        | FU4         | -                        | -                            | KRTAP5-5 | missense_variant     | 1            | p.S151C          | 0.52%  | no           | 18127            | 11                | 1651522          | NA          | NA              | S151            | C                               | G                    | 0                    |
| P33        | FU4         | -                        | -                            | ERBB2    | cn_amp               | NA           | cn_amp           | 2.59   | no           | NA               | 0,0;NA            | 17q12            | 17q12       | NA              | NA              | CN_gain                         |                      | 18                   |
| P33        | FU4         | -                        | -                            | ALK      | fusion               | Intron13_int | EML4-ALK         | 0.06%  | no           | 37326            | 2:42523448        | 2:42523448_2_2   | NA          | NA              | fusion          | 1;0;0,0                         | 1;0;0,0              | 0                    |
| P33        | FU4         | -                        | -                            | RB1      | stop_gained          | 13           | p.K420X          | 78.89% | maxAF        | 20194            | 13                | 48951096         | NA          | NA              | LOF             | A                               | T                    | 0                    |
| P33        | FU4         | -                        | -                            | EGFR     | missense_variant     | 20           | p.T790M          | 15.74% | no           | 45299            | 7                 | 55249071         | rs121434569 | COSM6240        | T790            | C                               | T                    | 4.12E-05             |
| P33        | FU4         | -                        | -                            | EGFR     | missense_variant     | 20           | p.C797S          | 0.75%  | no           | 45674            | 7                 | 55249092         | NA          | COSM5945664     | C797(in-cis_T79 | G                               | C                    | 0                    |
| P33        | FU4         | -                        | -                            | EGFR     | missense_variant     | 21           | p.L858R          | 56.41% | no           | 38569            | 7                 | 55259515         | rs121434568 | COSM6224        | L858            | T                               | G                    | 0                    |
| P33        | FU4         | -                        | -                            | EGFR     | cn_amp               | NA           | cn_amp           | 2.29   | no           | 0.298;6,16;1     | 7p11.2            | 7p11.2           | NA          | NA              | CN_gain         |                                 | 16                   |                      |
| P33        | FU4         | -                        | -                            | FGFR1    | cn_del               | NA           | cn_del           | 1.48   | no           | 0.217;2,21;1     | 8p11.22           | 8p11.22          | NA          | NA              | CN_loss         |                                 | 9                    |                      |
| P34        | baseline    | absent                   | -                            | EGFR     | disruptive_inframe_c | 19           | p.E746_A750del   | 0.40%  | maxAF        | 13519            | 7                 | 55242464         | rs121913421 | COSM6223        | Exon19_infram   | AGGAATTAAGA GAAGC               | A                    | 0                    |
| P34        | FU1,FU2,FU3 | -                        | No                           | EGFR     | disruptive_inframe_c | 19           | p.E746_A750del   | 0.49%  | maxAF        | 27748            | 7                 | 55242464         | rs121913421 | COSM6223        | Exon19_infram   | AGGAATTAAGA GAAGC               | A                    | 0                    |
| P34        | FU4         | -                        | -                            | EGFR     | disruptive_inframe_c | 19           | p.E746_A750del   | 0.73%  | no           | 11372            | 7                 | 55242464         | rs121913421 | COSM6223        | Exon19_infram   | AGGAATTAAGA GAAGC               | A                    | 0                    |
| P34        | FU4         | -                        | -                            | TP53     | missense_variant     | 7            | p.G245S          | 1.06%  | maxAF        | 10362            | 17                | 7577548          | rs28934575  | COSM6932        | G245            | C                               | T                    | 1.00E-04             |
| P34        | FU4         | -                        | -                            | ERBB2    | cn_amp               | NA           | cn_amp           | 2.32   | no           | NA               | 17q12             | 17q12            | NA          | NA              | CN_gain         |                                 | 29                   |                      |
| P35        | baseline    | -                        | -                            | EGFR     | conservative_infram  | 19           | p.T751_1759delin | 2.04%  | no           | 13915            | 7                 | 55242479         | NA          | COSM24270       | Exon19_infram   | CAACATCTCCG AAAGCCAACAA GGAAATC | CAAT                 | 0                    |
| P35        | baseline    | present                  | -                            | EGFR     | missense_variant     | 20           | p.T790M          | 2.68%  | maxAF        | 10098            | 7                 | 55249071         | rs121434569 | COSM6240        | T790            | C                               | T                    | 0                    |
| P35        | baseline    | -                        | -                            | TP53     | stop_gained          | 4            | p.W91X           | 0.91%  | no           | 12079            | 17                | 7579415          | NA          | COSM44192       | LOF             | C                               | T                    | 0                    |
| P36        | baseline    | present                  | -                            | EGFR     | missense_variant     | 20           | p.T790M          | 2.16%  | no           | 14778            | 7                 | 55249071         | rs121434569 | COSM6240        | T790            | C                               | T                    | 0                    |
| P36        | baseline    | -                        | Yes                          | EGFR     | missense_variant     | 21           | p.L858R          | 2.75%  | maxAF        | 16580            | 7                 | 55259515         | rs121434568 | COSM6224        | L858            | T                               | G                    | 0                    |
| P36        | FU3         | -                        | -                            | EGFR     | missense_variant     | 21           | p.L858R          | 0.26%  | maxAF        | 27409            | 7                 | 55259515         | rs121434568 | COSM6224        | L858            | T                               | G                    | 0                    |

| Patient ID | Timepoint   | T790M status at baseline | ctDNA clearance status at F1 | Gene      | Mutation type         | Exon number | Variant           | AF     | maxAF status | Sequencing depth | Chromosome number | Genomic position | dbsnp_ID    | COSM_ID        | MatchReport     | Reference allele (REF) | Variant allele (ALT) | Population Frequency |   |
|------------|-------------|--------------------------|------------------------------|-----------|-----------------------|-------------|-------------------|--------|--------------|------------------|-------------------|------------------|-------------|----------------|-----------------|------------------------|----------------------|----------------------|---|
| P39        | FU4         | -                        | -                            | EGFR      | missense_variant      | 20          | p.T790M           | 10.39% | no           | 26757            | 7                 | 55249071         | rs121434569 | COSM6240       | T790            | C                      | T                    | 2.84E-05             |   |
| P39        | FU4         | -                        | -                            | EGFR      | missense_variant      | 20          | p.C797S           | 6.65%  | no           | 24779            | 7                 | 55249091         | NA          | NA             | C797(in-cis_T79 | T                      | A                    | 0                    |   |
| P39        | FU4         | -                        | -                            | TP53      | splice_region_variant | 9           | c.920-12_920-3del | 5.08%  | no           | 23543            | 17                | 7576928          | NA          | NA             | Exon9_splice_si | TAGGAAAGAGG            | T                    | 0                    |   |
| P39        | FU4         | -                        | -                            | EGFR      | cn_amp                | NA          | cn_amp            | 2.73   | no           | 0.344;16;10      | 7p11.2            | 7p11.2           | NA          | NA             | CN_gain         | 16                     | 15 NA                | 0                    |   |
| P40        | baseline    | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.E746_A750del    | 1.01%  | maxAF        | 31228            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram   | AGGAATTAAGA            | GAAGC                | A                    | 0 |
| P40        | baseline    | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M           | 0.50%  | no           | 11202            | 7                 | 55249071         | rs121434569 | COSM6240       | T790            | C                      | T                    | 0                    |   |
| P40        | FU1,FU2     | -                        | No                           | EGFR      | disruptive_inframe_c  | 19          | p.E746_A750del    | 0.05%  | maxAF        | 28535            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram   | AGGAATTAAGA            | GAAGC                | A                    | 0 |
| P40        | FU4         | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.E746_A750del    | 0.06%  | maxAF        | 30317            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram   | AGGAATTAAGA            | GAAGC                | A                    | 0 |
| P41        | baseline    | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.L747_T751del    | 4.76%  | no           | 17426            | 7                 | 55242467         | rs121913442 | COSM392194     | Exon19_infram   | AATTAAGAGAA            | GCAAC                | A                    | 0 |
| P41        | baseline    | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M           | 4.98%  | maxAF        | 18397            | 7                 | 55249071         | rs121434569 | COSM6240       | T790            | C                      | T                    | 0                    |   |
| P41        | baseline    | -                        | -                            | TP53      | stop_gained           | 9           | p.Q317X           | 1.32%  | no           | 25495            | 17                | 7576897          | NA          | COSM3388166    | LOF             | G                      | A                    | 0                    |   |
| P41        | FU1         | -                        | No                           | EGFR      | disruptive_inframe_c  | 19          | p.L747_T751del    | 0.07%  | maxAF        | 20859            | 7                 | 55242467         | rs121913442 | COSM23571      | Exon19_infram   | AATTAAGAGAA            | GCAAC                | A                    | 0 |
| P41        | FU2         | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.L747_T751del    | 0.03%  | maxAF        | 27737            | 7                 | 55242467         | rs121913442 | COSM23571      | Exon19_infram   | AATTAAGAGAA            | GCAAC                | A                    | 0 |
| P41        | FU3         | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.L747_T751del    | 0.17%  | maxAF        | 26464            | 7                 | 55242467         | rs121913442 | COSM23571      | Exon19_infram   | AATTAAGAGAA            | GCAAC                | A                    | 0 |
| P41        | FU4         | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.L747_T751del    | 0.19%  | maxAF        | 28249            | 7                 | 55242467         | rs121913442 | COSM23571      | Exon19_infram   | AATTAAGAGAA            | GCAAC                | A                    | 0 |
| P42        | baseline    | absent                   | -                            | EGFR      | missense_variant      | 21          | p.L858R           | 0.27%  | maxAF        | 31595            | 7                 | 55259515         | rs121434568 | COSM6224       | L858            | T                      | G                    | 0                    |   |
| P42        | FU1,FU2     | -                        | acquired at FU               | KRAS      | missense_variant      | 2           | p.G12V            | 0.16%  | no           | 13678            | 12                | 25398284         | rs121913529 | COSM520,COSM1  | G12             | C                      | A                    | 0                    |   |
| P42        | FU1,FU2     | -                        | No                           | EGFR      | missense_variant      | 21          | p.L858R           | 0.34%  | maxAF        | 33553            | 7                 | 55259515         | rs121434568 | COSM6224       | L858            | T                      | G                    | 0                    |   |
| P42        | FU3         | -                        | -                            | MTOR      | splice_region_variant | 43          | c.6033+5G>T       | 1.04%  | no           | 33128            | 1                 | 11188056         | NA          | NA             | Exon43_splice_c | C                      | A                    | 0                    |   |
| P42        | FU3         | -                        | -                            | KRAS      | missense_variant      | 2           | p.G12V            | 2.01%  | no           | 14955            | 12                | 25398284         | rs121913529 | COSM520,COSM1  | G12             | C                      | A                    | 0                    |   |
| P42        | FU3         | -                        | -                            | TP53      | stop_gained           | 6           | p.R196X           | 1.19%  | no           | 32596            | 17                | 7578263          | rs397516435 | COSM99668,COSI | LOF             | G                      | A                    | 8.24E-06             |   |
| P42        | FU3         | -                        | -                            | KEAP1     | missense_variant      | 2           | p.I128F           | 2.57%  | maxAF        | 26417            | 19                | 10610328         | NA          | NA             | I128            | T                      | A                    | 0                    |   |
| P42        | FU3         | -                        | -                            | EGFR      | missense_variant      | 21          | p.L858R           | 1.73%  | no           | 34884            | 7                 | 55259515         | rs121434568 | COSM6224       | L858            | T                      | G                    | 0                    |   |
| P42        | FU3         | -                        | -                            | CDKN2A    | stop_gained           | 2           | p.E69X            | 0.94%  | no           | 16935            | 9                 | 21971153         | rs121913383 | COSM3092316,CI | LOF             | C                      | A                    | 0                    |   |
| P42        | FU4         | -                        | -                            | MTOR      | splice_region_variant | 43          | c.6033+5G>T       | 0.64%  | no           | 12342            | 1                 | 11188056         | NA          | NA             | Exon43_splice_c | C                      | A                    | 0                    |   |
| P42        | FU4         | -                        | -                            | KRAS      | missense_variant      | 2           | p.G12V            | 2.04%  | no           | 10276            | 12                | 25398284         | rs121913529 | COSM520,COSM1  | G12             | C                      | A                    | 0                    |   |
| P42        | FU4         | -                        | -                            | TP53      | stop_gained           | 6           | p.R196X           | 2.66%  | no           | 12296            | 17                | 7578263          | rs397516435 | COSM99668,COSI | LOF             | G                      | A                    | 4.06E-06             |   |
| P42        | FU4         | -                        | -                            | KEAP1     | missense_variant      | 2           | p.I128F           | 3.76%  | maxAF        | 9869             | 19                | 10610328         | NA          | NA             | I128            | T                      | A                    | 0                    |   |
| P42        | FU4         | -                        | -                            | EGFR      | missense_variant      | 21          | p.L858R           | 3.19%  | no           | 11955            | 7                 | 55259515         | rs121434568 | COSM6224       | L858            | T                      | G                    | 0                    |   |
| P42        | FU4         | -                        | -                            | CDKN2A    | stop_gained           | 2           | p.E69X            | 2.19%  | no           | 5032             | 9                 | 21971153         | rs121913383 | COSM3092316,CI | LOF             | C                      | A                    | 0                    |   |
| P43        | baseline    | -                        | -                            | BRCA2     | missense_variant      | 10          | p.A565S           | 5.85%  | no           | 18903            | 13                | 32907308         | NA          | NA             | A565            | G                      | T                    | 0                    |   |
| P43        | baseline    | -                        | -                            | TP53      | missense_variant      | 8           | p.C275Y           | 17.57% | no           | 29226            | 17                | 7577114          | rs863224451 | COSM165084,CO  | C275            | C                      | T                    | 0                    |   |
| P43        | baseline    | -                        | -                            | ERBB2     | cn_amp                | NA          | cn_amp            | 2.54   | no           | 0.396;240;17     | 17q12             | 17q12            | NA          | NA             | CN_gain         | 18                     | 18 NA                | 0                    |   |
| P43        | baseline    | -                        | -                            | NRXN1     | missense_variant      | 19          | p.V1204L          | 4.51%  | no           | 42952            | 2                 | 50463983         | NA          | COSM5494954,CI | V1204           | C                      | A                    | 4.07E-06             |   |
| P43        | baseline    | -                        | -                            | ALK       | cn_amp                | NA          | cn_amp            | 2.32   | no           | NA;0;NA          | 2p23.1            | 2p23.1           | NA          | NA             | CN_gain         | 10                     | 9 NA                 | 0                    |   |
| P43        | baseline    | -                        | -                            | POM121L12 | missense_variant      | 1           | p.S149I           | 7.12%  | no           | 46162            | 7                 | 53103810         | NA          | NA             | S149            | G                      | T                    | 0                    |   |
| P43        | baseline    | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M           | 6.46%  | no           | 55544            | 7                 | 55249071         | rs121434569 | COSM6240       | T790            | C                      | T                    | 4.12E-05             |   |
| P43        | baseline    | -                        | -                            | EGFR      | missense_variant      | 21          | p.L858R           | 24.42% | maxAF        | 48548            | 7                 | 55259515         | rs121434568 | COSM6224       | L858            | T                      | G                    | 0                    |   |
| P43        | baseline    | -                        | -                            | EGFR      | cn_amp                | NA          | cn_amp            | 3.09   | no           | 0.345;4;24;7     | 7p11.2            | 7p11.2           | NA          | NA             | CN_gain         | 16                     | 16 NA                | 0                    |   |
| P43        | baseline    | -                        | -                            | CDK6      | cn_amp                | NA          | cn_amp            | 2.82   | no           | 0.426;2;43;7     | 7q21.2            | 7q21.2           | NA          | NA             | CN_gain         | 7                      | 7 NA                 | 0                    |   |
| P43        | baseline    | -                        | -                            | MET       | cn_amp                | NA          | cn_amp            | 2.28   | no           | 0.298;5;15;7     | 7q31.2            | 7q31.2           | NA          | NA             | CN_gain         | 29                     | 21 NA                | 0                    |   |
| P43        | baseline    | -                        | -                            | FGFR1     | cn_amp                | NA          | cn_amp            | 2.34   | no           | 0.337;10;30      | 8p11.22           | 8p11.22          | NA          | NA             | CN_gain         | 9                      | 6 NA                 | 0                    |   |
| P43        | FU1,FU2     | -                        | -                            | TP53      | missense_variant      | 8           | p.C275Y           | 0.59%  | no           | 25616            | 17                | 7577114          | rs863224451 | COSM165084,CO  | C275            | C                      | T                    | 0                    |   |
| P43        | FU1,FU2     | -                        | No                           | EGFR      | missense_variant      | 21          | p.L858R           | 1.44%  | maxAF        | 28268            | 7                 | 55259515         | rs121434568 | COSM6224       | L858            | T                      | G                    | 0                    |   |
| P43        | FU1,FU2     | -                        | -                            | MET       | cn_amp                | NA          | cn_amp            | 2.28   | no           | 0.296;5;12;17    | 7q31.2            | 7q31.2           | NA          | NA             | CN_gain         | 29                     | 20 NA                | 0                    |   |
| P44        | baseline    | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M           | 0.12%  | no           | 12728            | 7                 | 55249071         | rs121434569 | COSM6240       | T790            | C                      | T                    | 0                    |   |
| P44        | baseline    | -                        | -                            | EGFR      | missense_variant      | 21          | p.L858R           | 0.50%  | no           | 13903            | 7                 | 55259515         | rs121434568 | COSM6224       | L858            | T                      | G                    | 0                    |   |
| P44        | baseline    | -                        | -                            | TP53      | missense_variant      | 5           | p.V143M           | 1.53%  | maxAF        | 13101            | 17                | 7578503          | NA          | COSM43878      | V143            | C                      | T                    | 0                    |   |
| P44        | baseline    | -                        | -                            | RB1       | frameshift_variant    | 19          | p.A628fs          | 0.55%  | no           | 18116            | 13                | 49030405         | NA          | NA             | LOF             | A                      | AT                   | 0                    |   |
| P45        | baseline    | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.E746_A750del    | 0.25%  | no           | 25269            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram   | AGGAATTAAGA            | GAAGC                | A                    | 0 |
| P45        | baseline    | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M           | 0.29%  | maxAF        | 32967            | 7                 | 55249071         | rs121434569 | COSM6240       | T790            | C                      | T                    | 4.12E-05             |   |
| P45        | FU1,FU2,FU3 | -                        | No                           | EGFR      | disruptive_inframe_c  | 19          | p.E746_A750del    | 0.02%  | maxAF        | 23275            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram   | AGGAATTAAGA            | GAAGC                | A                    | 0 |
| P45        | FU4         | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.E746_A750del    | 0.08%  | maxAF        | 13032            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram   | AGGAATTAAGA            | GAAGC                | A                    | 0 |
| P46        | baseline    | -                        | -                            | PIK3CA    | missense_variant      | 10          | p.E545K           | 0.16%  | no           | 7504             | 3                 | 178936091        | rs104886003 | COSM763        | E545            | G                      | A                    | 0                    |   |
| P46        | baseline    | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.E746_A750del    | 0.46%  | maxAF        | 12418            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram   | AGGAATTAAGA            | GAAGC                | A                    | 0 |
| P46        | baseline    | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M           | 0.26%  | no           | 8715             | 7                 | 55249071         | rs121434569 | COSM6240       | T790            | C                      | T                    | 0                    |   |
| P47        | baseline    | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M           | 1.00%  | no           | 7703             | 7                 | 55249071         | rs121434569 | COSM6240       | T790            | C                      | T                    | 0                    |   |
| P47        | baseline    | -                        | -                            | EGFR      | missense_variant      | 21          | p.L858R           | 1.90%  | maxAF        | 14320            | 7                 | 55259515         | rs121434568 | COSM6224       | L858            | T                      | G                    | 0                    |   |
| P47        | baseline    | -                        | -                            | TP53      | missense_variant      | 8           | p.R273H           | 0.26%  | no           | 12866            | 17                | 7577120          | rs28934576  | COSM99729      | R273            | C                      | T                    | 2.00E-04             |   |
| P47        | baseline    | -                        | -                            | BRINP3    | missense_variant      | 8           | p.T504M           | 0.59%  | no           | 17077            | 1                 | 190067938        | NA          | NA             | T504            | G                      | A                    | 0                    |   |
| P47        | baseline    | -                        | -                            | PTEN      | stop_gained           | 2           | p.Y189X           | 1.42%  | no           | 9538             | 10                | 89624274         | NA          | COSM28916      | LOF             | T                      | A                    | 0                    |   |
| P47        | baseline    | -                        | -                            | TP53      | missense_variant      | 5           | p.S127F           | 0.63%  | no           | 12966            | 17                | 7578550          | NA          | COSM44226      | S127            | G                      | A                    | 0                    |   |
| P47        | baseline    | -                        | -                            | HGF       | missense_variant      | 4           | p.H158Q           | 0.84%  | no           | 17790            | 7                 | 81386513         | NA          | NA             | H158            | G                      | T                    | 0                    |   |
| P48        | baseline    | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.E746_A750del    | 1.12%  | maxAF        | 14071            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram   | AGGAATTAAGA            | GAAGC                | A                    | 0 |
| P48        | baseline    | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M           | 0.49%  | no           | 7612             | 7                 | 55249071         | rs121434569 | COSM6240       | T790            | C                      | T                    | 0                    |   |
| P49        | baseline    | -                        | -                            | BRCA2     | missense_variant      | 18          | p.A2730V          | 1.00%  | no           | 33435            | 13                | 32937528         | rs80359067  | NA             | A2730           | C                      | T                    | 4.08E-06             |   |
| P49        | baseline    | -                        | -                            | EGFR      | conservative_infram   | 19          | p.E746_A750del    | 2.08%  | maxAF        | 23797            | 7                 | 55242465         | rs277504233 | COSM6225       | Exon19_infram   | AGGAATTAAGAG           | AAGCA                | G                    | 0 |
| P49        | baseline    | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M           | 1.47%  | no           | 29401            | 7                 | 55249071         | rs121434569 | COSM6240       | T790            | C                      | T                    | 4.12E-05             |   |
| P50        | baseline    | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M           | 22.13% | no           | 25182            | 7                 | 55249071         | rs121434569 | COSM6240       | T790            | C                      | T                    | 0                    |   |
| P50        | baseline    | -                        | -                            | EGFR      | missense_variant      | 21          | p.L858R           | 52.34% | maxAF        | 30265            | 7                 | 55259515         | rs121434568 | COSM6224       | L858            | T                      | G                    | 0                    |   |
| P50        | baseline    | -                        | -                            | MTOR      | missense_variant      | 35          | p.G1661S          | 48.27% | no           | 14515            | 1                 | 11199607         | NA          | NA             | G1661           | C                      | G                    | 0                    |   |
| P50        | baseline    | -                        | -                            | TP53      | missense_variant      | 5           | p.R156P           | 2.96%  | no           | 15825            | 17                | 7578463          | NA          | COSM3712457    | R156            | C                      | G                    | 0                    |   |
| P50        | baseline    | -                        | -                            | BRCA1     | splice_region_variant | 11          | NA                | 2.17%  | no           | 12284            | 17                | 41242938         | NA          | NA             | Exon11_splice_  | GCACA                  | GCA                  | 0                    |   |
| P50        | baseline    | -                        | -                            | EGFR      | cn_amp                | NA          | cn_amp            | 4      | no           | NA               | 7p11.2            | 7p11.2           | NA          | NA             | CN_gain         | 20                     | 20 NA                | 0                    |   |
| P51        | baseline    | absent                   | -                            | EGFR      | missense_variant      | 21          | p.L858R           | 12.45% | maxAF        | 14983            | 7                 | 55259515         | rs121434568 | COSM6224       | L858            | T                      | G                    | 0                    |   |
| P51        | baseline    | -                        | -                            | CNGB3     | missense_variant      | 3           | p.G96A            | 0.90%  | no           | 17153            | 8                 | 87738810         | NA          | NA             | G96             | C                      | G                    | 0                    |   |
| P51        | baseline    | -                        | -                            | RET       | splice_region_variant | 11          | NA                | 5.67%  | no           | 6001             | 10                | 43610199         | NA          | NA             | Exon11_splice_  | G                      | A                    | 4.00E-04             |   |
| P51        | baseline    | -                        | -                            | EXOC5     | missense_variant      | 15          | p.M530I           | 2.90%  | no           | 10341            | 14                | 57684723         | NA          | NA             | M530            | C                      | T                    | 1.00E-04             |   |
| P51        | baseline    | -                        | -                            | TP        |                       |             |                   |        |              |                  |                   |                  |             |                |                 |                        |                      |                      |   |

| Patient ID | Timepoint | T790M status at baseline | ctDNA clearance status at F1 | Gene      | Mutation type         | Exon number | Variant          | AF     | maxAF status | Sequencing depth | Chromosome number | Genomic position | dbsnp_ID    | COSM_ID        | MatchReport       | Reference allele (REF)                     | Variant allele (ALT) | Population Frequency |
|------------|-----------|--------------------------|------------------------------|-----------|-----------------------|-------------|------------------|--------|--------------|------------------|-------------------|------------------|-------------|----------------|-------------------|--------------------------------------------|----------------------|----------------------|
| P58        | baseline  | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.L747_P753delin | 7.28%  | maxAF        | 19771            | 7                 | 55242469         | rs121913438 | COSM133197     | Exon19_infram     | TTAAGAGAAGC<br>AACATCTC                    | T                    | 0                    |
| P58        | baseline  | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M          | 2.48%  | no           | 7392             | 7                 | 55249071         | rs121434569 | COSM6240       | T790              | C                                          | T                    | 0                    |
| P58        | baseline  | -                        | -                            | TP53      | frameshift_variant    | 3           | p.V31fs          | 4.32%  | no           | 9812             | 17                | 7579702          | NA          | NA             | LOF               | GAA                                        | G                    | 0                    |
| P59        | baseline  | -                        | -                            | EGFR      | conservative_infram   | 19          | p.E746_A750del   | 12.20% | maxAF        | 20047            | 7                 | 55242465         | NA          | COSM6225       | Exon19_infram     | GGAAATTAAGAG<br>AAGCA                      | G                    | 0                    |
| P59        | baseline  | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M          | 4.20%  | no           | 6635             | 7                 | 55249071         | rs121434569 | COSM6240       | T790              | C                                          | T                    | 0                    |
| P59        | baseline  | -                        | -                            | SI        | missense_variant      | 10          | p.S358R          | 3.01%  | no           | 10928            | 3                 | 16477764         | NA          | NA             | S358              | T                                          | G                    | 0                    |
| P59        | baseline  | -                        | -                            | PIK3CA    | missense_variant      | 21          | p.N1044K         | 3.14%  | no           | 21443            | 3                 | 178952077        | NA          | COSM27504      | N1044             | T                                          | G                    | 0                    |
| P59        | baseline  | -                        | -                            | TP53      | frameshift_variant    | 3           | p.N29fs          | 6.78%  | no           | 8022             | 17                | 7579707          | NA          | NA             | LOF               | TTG                                        | T                    | 0                    |
| P60        | baseline  | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.E746_A750del   | 21.82% | maxAF        | 19558            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram     | AGGAATTAAGA<br>GAAGC                       | A                    | 0                    |
| P60        | baseline  | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M          | 16.17% | no           | 9240             | 7                 | 55249071         | rs121434569 | COSM6240       | T790              | C                                          | T                    | 0                    |
| P60        | baseline  | -                        | -                            | TP53      | stop_gained           | 5           | p.W146X          | 13.26% | no           | 14747            | 17                | 7578492          | NA          | COSM3970379    | LOF               | C                                          | T                    | 0                    |
| P61        | baseline  | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M          | 1.98%  | no           | 16128            | 7                 | 55249071         | rs121434569 | COSM6240       | T790              | C                                          | T                    | 0                    |
| P61        | baseline  | -                        | -                            | EGFR      | missense_variant      | 21          | p.L858R          | 4.63%  | maxAF        | 19096            | 7                 | 55259515         | rs121434568 | COSM6224       | L858              | T                                          | G                    | 0                    |
| P61        | baseline  | -                        | -                            | TP53      | frameshift_variant    | 7           | p.C242fs         | 1.95%  | no           | 16504            | 17                | 7577554          | NA          | NA             | LOF               | TGCAG                                      | T                    | 0                    |
| P62        | baseline  | -                        | -                            | TP53      | missense_variant      | 8           | p.R283H          | 9.12%  | no           | 36017            | 17                | 7577090          | rs371409680 | COSM11483      | R283              | C                                          | T                    | 1.00E-04             |
| P62        | baseline  | -                        | -                            | TP53      | missense_variant      | 6           | p.Y205C          | 21.28% | maxAF        | 31168            | 17                | 7578235          | NA          | COSM99630,COSI | Y205              | T                                          | C                    | 0                    |
| P62        | baseline  | -                        | -                            | BRAF      | missense_variant      | 15          | p.V600E          | 0.37%  | no           | 21502            | 7                 | 140453136        | rs113488022 | COSM476        | V600              | A                                          | T                    | 1.65E-05             |
| P62        | baseline  | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M          | 11.31% | no           | 37018            | 7                 | 55249071         | rs121434569 | COSM6240       | T790              | C                                          | T                    | 4.12E-05             |
| P62        | baseline  | -                        | -                            | EGFR      | missense_variant      | 21          | p.L858R          | 19.88% | no           | 35470            | 7                 | 55259515         | rs121434568 | COSM6224       | L858              | T                                          | G                    | 0                    |
| P62        | baseline  | -                        | -                            | EGFR      | cn_amp                | NA          | cn_amp           | 2.35   | no           | 0.252;6;11;1     | 7p11.2            | 7p11.2           | NA          | NA             | CN_gain           |                                            | 16                   | 13 NA                |
| P62        | baseline  | -                        | -                            | FGFR1     | cn_del                | NA          | cn_del           | 1.62   | no           | NA;0;NA          | 8p11.22           | 8p11.22          | NA          | NA             | CN_loss           |                                            | 9                    | 9 NA                 |
| P62        | baseline  | -                        | -                            | MYC       | cn_del                | NA          | cn_del           | 1.75   | no           | 0.327;3;16;4     | 8q24.21           | 8q24.21          | NA          | NA             | CN_loss           |                                            | 8                    | 4 NA                 |
| P62        | FU1       | -                        | No                           | TP53      | missense_variant      | 6           | p.Y205C          | 0.81%  | maxAF        | 31644            | 17                | 7578235          | NA          | COSM99630,COSI | Y205              | T                                          | C                    | 0                    |
| P62        | FU1       | -                        | -                            | EGFR      | missense_variant      | 21          | p.L858R          | 0.56%  | no           | 35192            | 7                 | 55259515         | rs121434568 | COSM6224       | L858              | T                                          | G                    | 0                    |
| P62        | FU1       | -                        | -                            | BRAF      | missense_variant      | 15          | p.V600E          | 0.23%  | no           | 22132            | 7                 | 140453136        | rs113488022 | COSM476        | V600              | A                                          | T                    | 1.65E-05             |
| P62        | FU3       | -                        | -                            | TP53      | missense_variant      | 8           | p.R283H          | 0.81%  | no           | 34608            | 17                | 7577090          | rs371409680 | COSM11483      | R283              | C                                          | T                    | 1.00E-04             |
| P62        | FU3       | -                        | -                            | TP53      | missense_variant      | 6           | p.Y205C          | 1.35%  | no           | 26919            | 17                | 7578235          | NA          | COSM99630,COSI | Y205              | T                                          | C                    | 0                    |
| P62        | FU3       | -                        | -                            | EGFR      | missense_variant      | 21          | p.L858R          | 1.45%  | maxAF        | 25442            | 7                 | 55259515         | rs121434568 | COSM6224       | L858              | T                                          | G                    | 0                    |
| P62        | FU4       | -                        | -                            | TP53      | missense_variant      | 8           | p.R283H          | 0.90%  | no           | 29528            | 17                | 7577090          | rs371409680 | COSM11483      | R283              | C                                          | T                    | 1.00E-04             |
| P62        | FU4       | -                        | -                            | TP53      | missense_variant      | 6           | p.Y205C          | 2.41%  | no           | 23928            | 17                | 7578235          | NA          | COSM99630,COSI | Y205              | T                                          | C                    | 0                    |
| P62        | FU4       | -                        | -                            | EGFR      | missense_variant      | 21          | p.L858R          | 2.57%  | maxAF        | 23446            | 7                 | 55259515         | rs121434568 | COSM6224       | L858              | T                                          | G                    | 0                    |
| P63        | baseline  | -                        | -                            | TP53      | splice_region_variant | 6           | p.E224r          | 32.39% | no           | 33890            | 17                | 7578177          | rs267605076 | COSM707897,CO  | Exon6_splice_si   | C                                          | T                    | 0                    |
| P63        | baseline  | -                        | -                            | TP53      | missense_variant      | 9           | p.C176Y          | 1.86%  | no           | 34789            | 17                | 7578403          | rs786202962 | COSM1649384,C  | C176              | C                                          | T                    | 0                    |
| P63        | baseline  | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.E746_A750del   | 43.42% | maxAF        | 36238            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram     | AGGAATTAAGA<br>GAAGC                       | A                    | 0                    |
| P63        | baseline  | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M          | 13.93% | no           | 65684            | 7                 | 55249071         | rs121434569 | COSM6240       | T790              | C                                          | T                    | 4.12E-05             |
| P63        | baseline  | -                        | -                            | EGFR      | cn_amp                | NA          | cn_amp           | 2.8    | no           | 0.282;10;14      | 7p11.2            | 7p11.2           | NA          | NA             | CN_gain           |                                            | 16                   | 16 NA                |
| P64        | baseline  | -                        | -                            | APC       | missense_variant      | 16          | p.K2052T         | 18.62% | no           | 22469            | 5                 | 112177446        | NA          | NA             | K2052             | A                                          | C                    | 0                    |
| P64        | baseline  | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M          | 12.88% | no           | 14305            | 7                 | 55249071         | rs121434569 | COSM6240       | T790              | C                                          | T                    | 0                    |
| P64        | baseline  | -                        | -                            | OR4A15    | missense_variant      | 1           | p.K53N           | 1.05%  | no           | 26354            | 11                | 55135518         | NA          | NA             | K53               | G                                          | T                    | 0                    |
| P64        | baseline  | -                        | -                            | CTNNB1    | inframe_deletion      | 3           | p.S23_S33del     | 8.58%  | no           | 15587            | 3                 | 41266068         | NA          | NA             | Exon3_inframe     | TTAGTCACTGG<br>CAGCAACAGTC<br>TTACTTGGACTC | T                    | 0                    |
| P64        | baseline  | -                        | -                            | EGFR      | inframe_deletion      | 19          | p.E746_A750del   | 34.29% | maxAF        | 23159            | 7                 | 55242465         | NA          | COSM6225       | Exon19_infram     | GGAAATTAAGAG<br>AAGCA                      | G                    | 0                    |
| P64        | baseline  | -                        | -                            | BRAF      | cn_amp                | NA          | cn_amp           | 2.62   | no           | NA               | 7q34              | 7q34             | NA          | NA             | CN_gain           |                                            | 6                    | 6 NA                 |
| P64        | baseline  | -                        | -                            | CDKN2A    | cn_del                | NA          | cn_del           | 1.23   | no           | NA               | 9p21.3            | 9p21.3           | NA          | NA             | CN_loss           |                                            | 12                   | 11 NA                |
| P64        | baseline  | -                        | -                            | EGFR      | cn_amp                | NA          | cn_amp           | 2.49   | no           | NA               | 7p11.2            | 7p11.2           | NA          | NA             | CN_gain           |                                            | 20                   | 18 NA                |
| P64        | baseline  | -                        | -                            | MET       | cn_amp                | NA          | cn_amp           | 2.58   | no           | NA               | 7q31.2            | 7q31.2           | NA          | NA             | CN_gain           |                                            | 45                   | 43 NA                |
| P64        | baseline  | -                        | -                            | POM121L12 | cn_amp                | NA          | cn_amp           | 2.77   | no           | NA               | 7p12.1            | 7p12.1           | NA          | NA             | CN_gain           |                                            | 8                    | 8 NA                 |
| P64        | FU4       | -                        | -                            | EGFR      | missense_variant      | 20          | p.C797S          | 4.20%  | no           | 30193            | 7                 | 55249092         | NA          | COSM5945664    | C797(in-cis_T79 G | C                                          | C                    | 0                    |
| P64        | FU4       | -                        | -                            | EGFR      | conservative_infram   | 19          | p.E746_A750del   | 9.93%  | maxAF        | 31103            | 7                 | 55242465         | rs727504233 | COSM6225       | Exon19_infram     | GGAAATTAAGAG<br>AAGCA                      | G                    | 0                    |
| P64        | FU4       | -                        | -                            | APC       | missense_variant      | 16          | p.K2052T         | 5.44%  | no           | 28328            | 5                 | 112177446        | NA          | NA             | K2052             | A                                          | C                    | 0                    |
| P64        | FU4       | -                        | -                            | CTNNB1    | conservative_infram   | 3           | p.S23_S33del     | 1.39%  | no           | 36303            | 3                 | 41266068         | NA          | NA             | Exon3_inframe     | TTAGTCACTGG<br>CAGCAACAGTC<br>TTACTTGGACTC | T                    | 0                    |
| P64        | FU4       | -                        | -                            | EGFR      | missense_variant      | 20          | p.T790M          | 5.05%  | no           | 31174            | 7                 | 55249071         | rs121434569 | COSM6240       | T790              | C                                          | T                    | 2.84E-05             |
| P65        | baseline  | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M          | 20.08% | no           | 22144            | 7                 | 55249071         | rs121434569 | COSM6240       | T790              | C                                          | T                    | 0                    |
| P65        | baseline  | -                        | -                            | EGFR      | missense_variant      | 21          | p.L858R          | 58.42% | maxAF        | 32248            | 7                 | 55259515         | rs121434568 | COSM6224       | L858              | T                                          | G                    | 0                    |
| P65        | baseline  | -                        | -                            | EGFR      | cn_amp                | NA          | cn_amp           | 4.04   | no           | NA               | 7p11.2            | 7p11.2           | NA          | NA             | CN_gain           |                                            | 20                   | 20 NA                |
| P67        | baseline  | -                        | -                            | EGFR      | disruptive_inframe_c  | 19          | p.E746_A750del   | 5.27%  | maxAF        | 27832            | 7                 | 55242464         | rs121913421 | COSM6223       | Exon19_infram     | AGGAATTAAGA<br>GAAGC                       | A                    | 0                    |
| P67        | baseline  | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M          | 1.63%  | no           | 11804            | 7                 | 55249071         | rs121434569 | COSM6240       | T790              | C                                          | T                    | 0                    |
| P67        | baseline  | -                        | -                            | TP53      | missense_variant      | 8           | p.R273C          | 0.25%  | no           | 20616            | 17                | 7577121          | rs121913343 | COSM99933      | R273              | G                                          | A                    | 0                    |
| P68        | baseline  | -                        | -                            | EGFR      | conservative_infram   | 19          | p.E746_A750del   | 3.49%  | no           | 19630            | 7                 | 55242465         | NA          | COSM6225       | Exon19_infram     | GGAAATTAAGAG<br>AAGCA                      | G                    | 0                    |
| P68        | baseline  | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M          | 6.45%  | maxAF        | 7243             | 7                 | 55249071         | rs121434569 | COSM6240       | T790              | C                                          | T                    | 0                    |
| P68        | baseline  | -                        | -                            | TP53      | missense_variant      | 5           | p.A159P          | 6.41%  | no           | 17917            | 17                | 7578455          | NA          | COSM562628     | A159              | C                                          | G                    | 0                    |
| P68        | baseline  | -                        | -                            | CTNNB1    | missense_variant      | 3           | p.S37C           | 5.74%  | no           | 19590            | 3                 | 41266113         | rs121913403 | COSM5679       | S37               | C                                          | G                    | 0                    |
| P68        | baseline  | -                        | -                            | CSMD1     | stop_gained           | 25          | p.R1288X         | 4.43%  | no           | 19524            | 8                 | 3165305          | NA          | COSM4782185    | LOF               | G                                          | A                    | 0                    |
| P69        | baseline  | -                        | -                            | TP53      | frameshift_variant    | 4           | p.G112fs         | 5.93%  | no           | 10003            | 17                | 7579351          | NA          | NA             | LOF               | GC                                         | G                    | 0                    |
| P69        | baseline  | -                        | -                            | MET       | missense_variant      | 19          | p.Y1235C         | 1.89%  | no           | 15795            | 7                 | 116423429        | NA          | NA             | Y1235             | A                                          | G                    | 0                    |
| P69        | baseline  | present                  | -                            | EGFR      | missense_variant      | 20          | p.T790M          | 28.10% | no           | 52170            | 7                 | 55249071         | rs121434569 | COSM6240       | T790              | C                                          | T                    | 4.12E-05             |
| P69        | baseline  | -                        | -                            | EGFR      | missense_variant      | 21          | p.L858R          | 49.55% | maxAF        | 41700            | 7                 | 55259515         | rs121434568 | COSM6224       | L858              | T                                          | G                    | 0                    |
| P69        | baseline  | -                        | -                            | EGFR      | cn_amp                | NA          | cn_amp           | 4.23   | no           | 0.266;10;23      | 7p11.2            | 7p11.2           | NA          | NA             | CN_gain           |                                            | 16                   | 16 NA                |
| P81        | baseline  | -                        | -                            | TP53      | missense_variant      | 6           | p.I195T          | 0.54%  | no           | 35857            | 17                | 7578265          | rs760043106 | COSM116923,CO  | I195              | A                                          | G                    | 0                    |
| P81        | baseline  | -                        | -                            | EGFR      | conservative_infram   | 19          | p.E746_A750del   | 2.66%  | maxAF        | 30410            | 7                 | 55242465         | rs727504233 | COSM6225       | Exon19_infram     | GGAAATTAAGAG<br>AAGCAAG                    |                      |                      |

Abbreviations: FU, follow-up; cn\_amp, copy number amplification; cn\_del, copy number deletion; CN\_gain, copy number gain; NA, not applicable; -, no data or not applicable