

Supplementary Table 1. Correlation between the mutational status of the EGFR gene detected by liquid biopsy techniques and alterations in the *ALK*, *ROS1* and *BRAF* genes.

| Molecular Characteristics | Number of patients [n (%)] | | Total of cases (n=38) | <i>P</i> |
|---------------------------|----------------------------|-----------------|--------------------------|----------|
| | No (n = 11) | Yes (n = 27) | | |
| <i>ALK</i> * | | | | |
| Negative | 8 (100) | 24 (100) | 32 (100) | NS |
| Positive | 0 (0) | 0 (0) | 0 (0) | |
| <i>ROS1</i> * | | | | |
| Negative | 1 (100) | 5 (100) | 6 (100) | NS |
| Positive | 0 (0) | 0 (0) | 0 (0) | |
| <i>BRAF</i> * | | | | |
| Negative | 2 (100) | 3 (100) | 5 (100) | NS |
| Positive | 0 (0) | 0 (0) | 0 (0) | |

The study of driver mutations in biopsy sample has been carried out according to current evidence at the time of diagnosis, but due to the scarcity of tumor tissue, alterations in the *ALK*, *ROS1* and *BRAF* genes were performed in 32, 16 and 5 patients; respectively.

Supplementary Table 2. Type of mutations detected in the *EGFR* gene in plasma samples from *EGFR* mutated non-small cell lung cancer (NSCLC) patients (*n*=38) during disease follow-up.

| ID | Biopsy mutation | Number of liquid biopsies per patient | | | | | | |
|----|-----------------|---------------------------------------|---------------------|---------------|-------------|---------------|---------------|---------------|
| | | 1 | 2 | 3 | 4 | 5 | 6 | 7 |
| 1 | del(19) | WT | WT | WT | WT | del(19) | - | - |
| 2 | L858R | L858R | - | - | - | - | - | - |
| 3 | G719X+S578I | WT | WT | WT | WT | WT | WT | - |
| 4 | ins(20) | WT | - | - | - | - | - | - |
| 5 | L858R | L858R | - | - | - | - | - | - |
| 6 | ins(20) | ins(20) | - | - | - | - | - | - |
| 7 | L858R | WT | WT | WT | L858R | L858R | L858R | - |
| 8 | L858R | WT | WT | WT | WT | - | - | - |
| 9 | ins(20) | WT | - | - | - | - | - | - |
| 10 | del(19) | WT | del(19) | WT | - | - | - | - |
| 11 | L858R | L858R | - | - | - | - | - | - |
| 12 | del(19) | del(19) | WT | - | - | - | - | - |
| 13 | del(19) | del(19) | - | - | - | - | - | - |
| 14 | del(19) | WT | del(19) | WT | del(19) | del(19) | - | - |
| 15 | del(19) | WT | WT | WT | - | - | - | - |
| 16 | del(19) | L858R | - | - | - | - | - | - |
| 17 | L858R | WT | - | - | - | - | - | - |
| 18 | L858R | WT | WT | WT | WT | WT | WT | WT |
| 19 | L858R | L858R | L858R | - | - | - | - | - |
| 20 | del(19) | WT | - | - | - | - | - | - |
| 21 | L858R | WT | WT | WT | WT | - | - | - |
| 22 | L858R | L858R | WT | WT | L858R | - | - | - |
| 23 | L858R | WT | WT | WT | - | - | - | - |
| 24 | L858R | L858R | L858R | - | - | - | - | - |
| 25 | L858R | WT | - | - | - | - | - | - |
| 26 | L858R | L858R | L858R | - | - | - | - | - |
| 27 | del(19) | del(19) | del(19)+T790M | del(19) | del(19) | - | - | - |
| 28 | del(19) | del(19) | del(19)+T790M | - | - | - | - | - |
| 29 | L858R | L858R | L858R | WT | L858R+T790M | WT | WT | WT |
| 30 | del(19) | WT | del(19)+T790 M | - | - | - | - | - |
| 31 | del(19)+L861Q | del(19)+L861Q | del(19)+L861Q+T790M | - | - | - | - | - |
| 32 | del(19)+T790M | del(19)+T790M | del(19)+T790M | del(19)+T790M | - | - | - | - |
| 33 | del(19) | del(19) | del(19) | del(19)+T790M | - | - | - | - |
| 34 | L858R | L858R | WT | L858R+T790M | - | - | - | - |
| 35 | del(19) | del(19) | del(19)+T790M | del(19) | - | - | - | - |
| 36 | del(19) | del(19) | del(19)+T790M | WT | - | - | - | - |
| 37 | del(19) | WT | del(19)+T790M | del(19) | del(19) | del(19)+T790M | del(19)+T790M | del(19)+T790M |
| 38 | del(19) | del(19)+T790M | - | - | - | - | - | - |

WT: Wild Type; -: not done.