

## Phase I dose-escalation study of the mTOR inhibitor sirolimus and the HDAC inhibitor vorinostat in patients with advanced malignancy

### Supplementary Material

**Supplementary Table 1. Detected molecular aberrations, affected pathways and treatment outcomes**

| Molecular Aberration                            | Affected pathway | PR | SD > 12 months | SD < 12 months | PD | Not evaluable for response |
|---|------------------|----|----------------|----------------|----|----------------------------|
| APC A2122_C2123insA                             | Wnt              |    | 1              |                |    |                            |
| BRAF V600E                                      | MAPK             |    |                |                | 1  |                            |
| GNAQ Q209P                                      | MAPK             |    |                |                | 1  |                            |
| KIT M541L <sup>1</sup> , KDR C482R <sup>1</sup> | KIT, VEGFR       |    |                | 1              |    |                            |
| KIT M541L <sup>1</sup>                          | KIT              | 1  | 1              |                |    |                            |
| KRAS G12C                                       | MAPK             |    |                | 1              |    |                            |
| KRAS G12D                                       | MAPK             |    |                |                | 1  |                            |
| KRAS G12S                                       | MAPK             |    |                |                |    | 1                          |
| KRAS G13D                                       | MAPK             |    |                | 1              |    |                            |
| MET N375S                                       | MET              |    |                | 1              |    |                            |
| NF1 R1241*, BAP1 truncation, CTNNB1 N387K       | MAPK, BAP, Wnt   |    |                |                | 1  |                            |
| NRAS Q61K                                       | MAPK             |    |                |                | 1  |                            |
| NRAS Q61R                                       | MAPK             |    |                | 1              | 1  | 1                          |
| PIK3CA H1047L                                   | PI3K             |    |                |                |    | 1                          |
| PIK3CA H1047R, CTNNB1 S37C, PTPRD S1845fs*2     | PI3K, Wnt, STAT3 |    |                | 1              |    |                            |
| PTEN R335*, CDKN2A/B loss                       | PI3K, CDK        |    |                | 1              |    |                            |
| TP53 G245S                                      | TP53             |    |                |                |    | 1                          |
| TP53 L194F                                      | TP53             |    |                | 1              |    |                            |
| TP53 R175H                                      | TP53             |    |                |                | 2  |                            |
| TSC2 loss, XPO1 E571K                           | mTOR, XPO1       | 1  |                |                |    |                            |
| No aberration                                   |                  |    |                | 8              | 13 | 3                          |
| Not done  |                  |    |                | 7              | 14 | 2                          |

Abbreviations: PR, partial response; SD, stable disease; PD, progressive disease

<sup>1</sup>Possible germline polymorphism