

| | | |
|-----------|--------|-----------|
| | | 9 |
| 1. | (| |
| |) | 10 |
| 1.1. |) | 10 |
| 1.2. | (| 10 |
| |) | 11 |
| 1.3. | (| 11 |
| 1.4. | (| 12 |
| |) | 12 |
| 1.5. | (| 14 |
| 1.5.1. | | 14 |
| 1.5.2. | | 15 |
| 1.6. | (| 23 |
| |) | 23 |
| 2. | (| |
| |), | 26 |
| 2.1. | | 28 |
| 2.2. | | 28 |
| 2.3. | | 30 |
| 2.4. | | 31 |
| 2.5. | | 34 |
| 3. | , | |
| | , | |
| | , | |
| | , | 37 |
| 3.1. | (I-II) | 37 |
| 3.1.1 | | 40 |
| 3.2. | III | 43 |
| 3.3. | | 45 |
| 3.4. | | |
| (III/C/D | - IV) | 49 |
| 3.4.1. | 1- | 49 |

| | | | |
|-----------|-----|-------------------|------------|
| 3.4.2. | 1- | <i>BRAF</i> | 50 |
| 3.4.3. | 1- | <i>c-Kit</i> | 57 |
| 3.4.4. | 1 - | <i>BRAF c-Kit</i> | 59 |
| 3.4.5. | - | PD1 CTLA4 | 60 |
| 3.4.6. | - | PD1 CTLA4 | 61 |
| 3.4.7. | | : | 62 |
| 3.4.8. | | | 64 |
| 3.4.9. | | | 65 |
| 3.5. | | | 66 |
| 3.6. | | | 66 |
| 3.7. | | | 68 |
| 4. | | | 69 |
| 4.1. | | | 69 |
| 4.2. | | | 69 |
| 4.2.1. | | | 69 |
| 4.2.2. | | | 71 |
| 4.2.3. | | | 72 |
| 4.3. | | | 72 |
| 4.4. | | | 74 |
| 4.5. | | | 75 |
| 5. | | | 77 |
| 6. | | | 80 |
| 7. | (| , | 85 |
| |) | | 86 |
| | | | 88 |
| | | | 116 |
| 1. | | | 116 |
| 2. | | | 120 |

| | | |
|----|--------------|------------|
| . | | 123 |
| , | | 124 |
| . | | 132 |
| 1. | BO3/ECOG 137 | |
| 2. | | 138 |
| . | (RECIST 1.1) | 139 |