| All patients | | | | | |
|--|----------------------|--------------------|----------------------|--------------------|---------------|
| | MRD- at EOI1 (n=544) | | MRD+ at EOI1 (n=240) | | MRD- vs. MRD+ |
| | Ν | % ± 2SE% | Ν | % ± 2SE% | Р |
| 5yr OS from end of induction 1 (Course 1) | 544 | $72.9\%\pm3.9\%$ | 240 | $48.2\%\pm6.6\%$ | <0.001 |
| | | | | | |
| 5yr DFS from end of induction 1 (Course 1) | 544 | 57.2% ± 4.3% | 240 | $30.5\% \pm 6.1\%$ | <0.001 |
| | | | | | |
| 5yr RR from end of induction 1 (Course 1) | 544 | $38.6\% \pm 4.2\%$ | 240 | $61.3\% \pm 6.4\%$ | <0.001 |
| | | | | | |
| 5yr TRM from end of induction 1 (Course 1) | 544 | $4.3\%\pm1.7\%$ | 240 | $8.2\%\pm3.6\%$ | 0.028 |

Supplemental Table 1. 5 year-outcomes of patients with or without MRD at EOI1 or at EOI2

All patients

| | MRD- at EOI2 (n=544) | | MRD+ at EOI2 (n=121) | | MRD- vs. MRD+ |
|--|----------------------|--------------------|----------------------|--------------------|---------------|
| | Ν | % ± 2SE% | Ν | % ± 2SE% | Р |
| 5yr OS from end of induction 1 (Course 2) | 544 | $72.2\%\pm3.9\%$ | 121 | $45.7\%\pm9.2\%$ | <0.001 |
| | | | | | |
| 5yr DFS from end of induction 1 (Course 2) | 544 | $59.4\% \pm 4.3\%$ | 121 | $23.1\%\pm7.8\%$ | <0.001 |
| | | | | | |
| 5yr RR from end of induction 1 (Course 2) | 544 | $36.5\% \pm 4.2\%$ | 121 | $71.9\% \pm 8.3\%$ | <0.001 |
| | | | | | |
| 5yr TRM from end of induction 1 (Course 2) | 544 | $4.1\%\pm1.7\%$ | 121 | $5.0\%\pm4.0\%$ | 0.614 |

Supplemental Table 2 Demographic, laboratory, and clinical characteristics of patients with or without MRD at the EOI2.

| | | End o | f Induction 2 MRI | Induction 2 MRD status | |
|--|----------|-----------------|-------------------|------------------------|------------------|
| | MRD- | at EOI2 (n=544) | MRD+ | at EOI2 (n=121) | |
| Characteristic | Ν | % | Ν | % | Р |
| Total | 544 | | 121 | | |
| Gender | | | | | |
| Male | 257 | 47.2% | 64 | 52.9% | 0.261 |
| Female | 287 | 52.8% | 57 | 47.1% | |
| Age Group | | | | | |
| 0-1 year | 101 | 18.6% | 22 | 18.2% | 0.922 |
| 2-10 year | 205 | 37.7% | 41 | 33.9% | 0.434 |
| >11 year | 238 | 43.8% | 58 | 47.9% | 0.402 |
| Cytogenetics | | | | | |
| Normal | 77 | 18.8% | 52 | 31.5% | 0.001 |
| t(8;21) | 75 | 18.3% | 14 | 8.5% | 0.003 |
| inv(16) | 62 | 15.2% | 7 | 4.2% | <0.001 |
| t(9;11)/11q23 | 94 | 23.0% | 24 | 14.5% | 0.024 |
| t(6;9) | 5 | 1.2% | 5 | 3.0% | 0.160 |
| Monosomy 7 | 3 | 0.7% | 6 | 3.6% | 0.020 |
| Del(7q) | 1 | 0.2% | 5 | 3.0% | 0.009 |
| -5/5q | 7 | 1.7% | 3 | 1.8% | 1.000 |
| +8 | 19 | 4.6% | 19 | 11.5% | 0.003 |
| Other | 66 | 16.1% | 30 | 18.2% | 0.552 |
| Unknown | 12 | | 3 | | |
| ITD status | | | 2 | | |
| Negative | 462 | 89.0% | 75 | 67.0% | <0.001 |
| Positive | 61 | 11.8% | 37 | 33.0% | |
| High Allelic Ratio | 31 | 5.9% | 22 | 19.6% | |
| Low Allelic Ratio | 30 | 5.7% | 15 | 13.4% | |
| Unknown | 21 | 5.170 | 9 | 13.170 | |
| CEBPA status | 21 | | , | | |
| Negative | 486 | 93.6% | 112 | 100.0% | 0.006 |
| Positive | 33 | 6.4% | 0 | 0.0% | 0.000 |
| Unknown | 25 | 0.470 | 9 | 0.070 | |
| NPM status | 25 | | , | | |
| Negative | 475 | 91.3% | 104 | 92.9% | 0.601 |
| Positive | 45 | 8.7% | 8 | 7.1% | 0.001 |
| Unknown | 24 | 0.770 | 9 | 7.170 | |
| Risk group (cyto/mutation) | 24 | | 7 | | |
| Standard | 258 | 48.4% | 78 | 65.5% | 0.001 |
| Low | 230 | 43.2% | 14 | 11.8% | <0.001 |
| High | 45 | 8.4% | 27 | 22.7% | <0.001 |
| Unknown | 43 | 0.470 | 27 | 22.170 | <0.001 |
| Risk group (cytogenetic only) | 11 | | 2 | | |
| Standard | 347 | 65.6% | 102 | 86.4% | <0.001 |
| Low | 167 | 31.6% | 102 | 9.3% | <0.001 |
| Low High | 167 | 2.8% | 5 | 4.2% | 0.387 |
| | | 2.0% | | 4.270 | 0.587 |
| Unknown Response by End of Course 1 | 15 | | 3 | | + |
| * * | 167 | 06 20/ | 17 | 20.00/ | <0.001 |
| CR Dortial remission (> 5% 15%) | 463 | 86.2% | 47 | 39.2% | |
| Partial remission (>5% - 15%) Persistent disease (>15%) | 46 28 | 8.6% 5.2% | 31 42 | 25.8% 35.0% | <0.001 <0.001 |
| | | | | | |
| CNS refractory (declared off therapy) | 0 | 0.0% | 0 | 0.0% | - |
| Death Not Evolucida | 0 7 | 0.0% | 0 | 0.0% | |
| Not Evaluable | / | | 1 | | |
| Response by End of Course 2 | 520 | 00.10/ | 70 | 50.20/ | .0.001 |
| CR | 530 | 98.1% | 70 | 59.3% | <0.001 |
| Not CR | 10 | 1.9% | 48 | 40.7% | |
| Not Evaluable | 4 | | 3 | | |
| Age Madian (magaz) | 0.5 | (0.005 | 10.0 | (0.002 20.4) | 0.107 |
| Median (range) | 9.5 | (0.005 - 23.5) | 10.6 | (0.003 - 29.4) | 0.186 |
| WBC x10 ³ µL | 22.7 | (0 5 007 0) | 27.5 | (1.0 | 0.170 |
| Median (range) | 23.7 | (0.5 - 827.2) | 27.6 | (1.0 - 446) | 0.170 |
| BM Blasts % | ~~ | (0.100) | | (2 | 0.001 |
| Median (range) | 65 | (0 - 100) | 71.1 | (3 - 99) | 0.206 |
| Protocol SCT received? | | | | | |
| Yes | 85 | 15.6% | 22 | 18.2% | 0.489 |
| No | 459 | 84.4% | 99 | 81.8% | 1 |

Supplemental Table 3. Information of patients who received bone marrow transplant with or without MRD at EOI1 and EOI2

| All patients | N (total patients) | N (patients who received BMT) | % of patients with BMT |
|----------------------------|--------------------|-------------------------------|------------------------|
| MRD- at EOI1, MRD- at EOI2 | 401 | 56 | 11 |
| MRD+ at EOI1, MRD- at EOI2 | 65 | 19 | 29 |
| MRD+ at EOI1, MRD+ at EOI2 | 57 | 18 | 32 |

| Standard risk only | N (total patients) | N (patients who received BMT) | % of patients with BMT |
|----------------------------|--------------------|-------------------------------|------------------------|
| MRD- at EOI1, MRD- at EOI2 | 182 | 29 | 16 |
| MRD+ at EOI1, MRD- at EOI2 | 35 | 10 | 29 |
| MRD+ at EOI1, MRD+ at EOI2 | 38 | 9 | 24 |

Supplemental Figure 1. 5-year TRM subdivided by EOI1 leukemia burden. The correlation of 5-year TRM and the amount of MRD at EOI1 was shown.

