

Supplementary Figs and legends

Figure S1

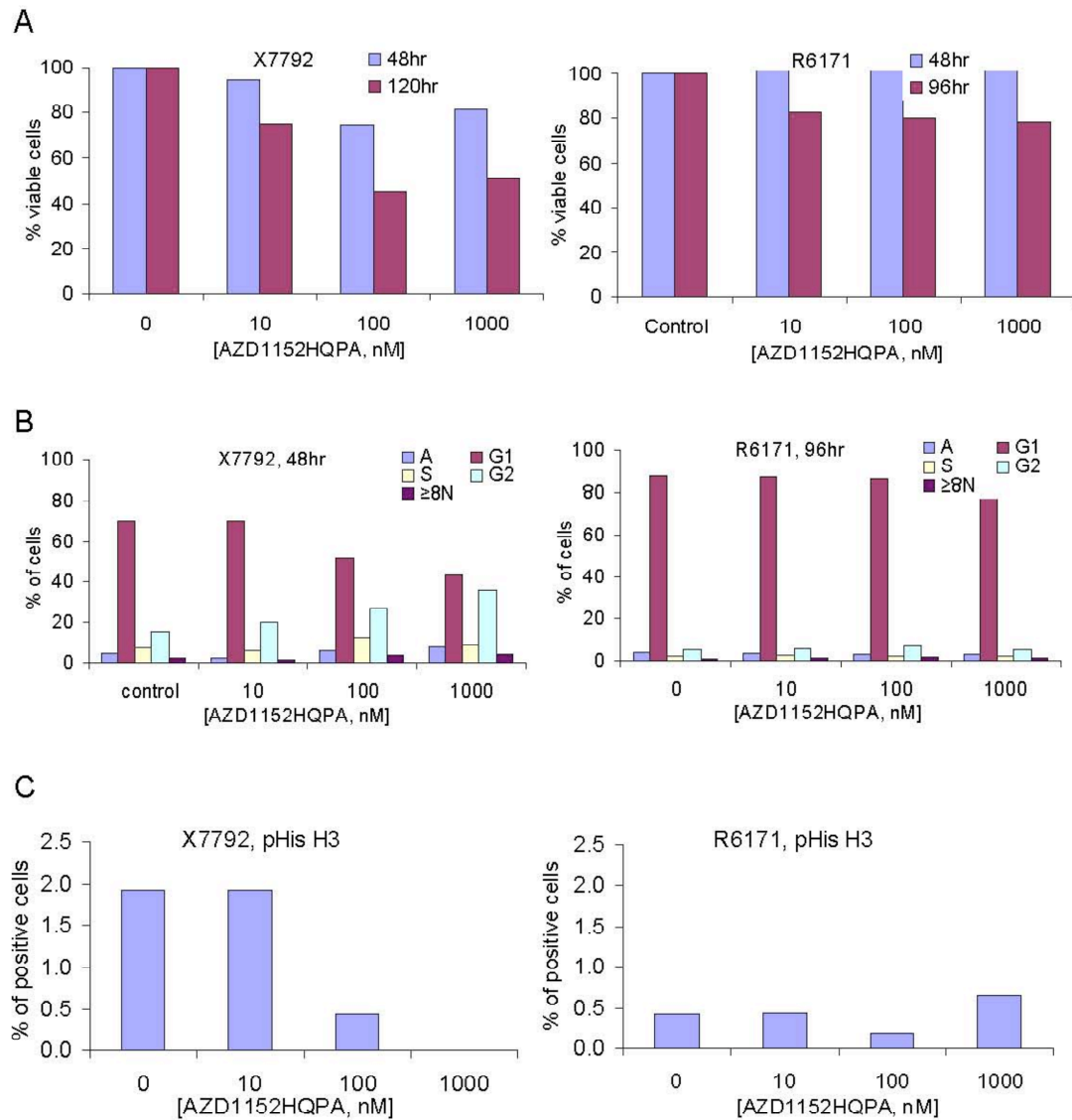


Figure S1. AZD1152-HQPA activity in 2 primary AML samples showing effect on viable cell number (Fig S1A), cell cycle distribution (Fig S1B, including apoptotic (sub-G1) and polyploid cells) and pHis H3 (Fig S1C).

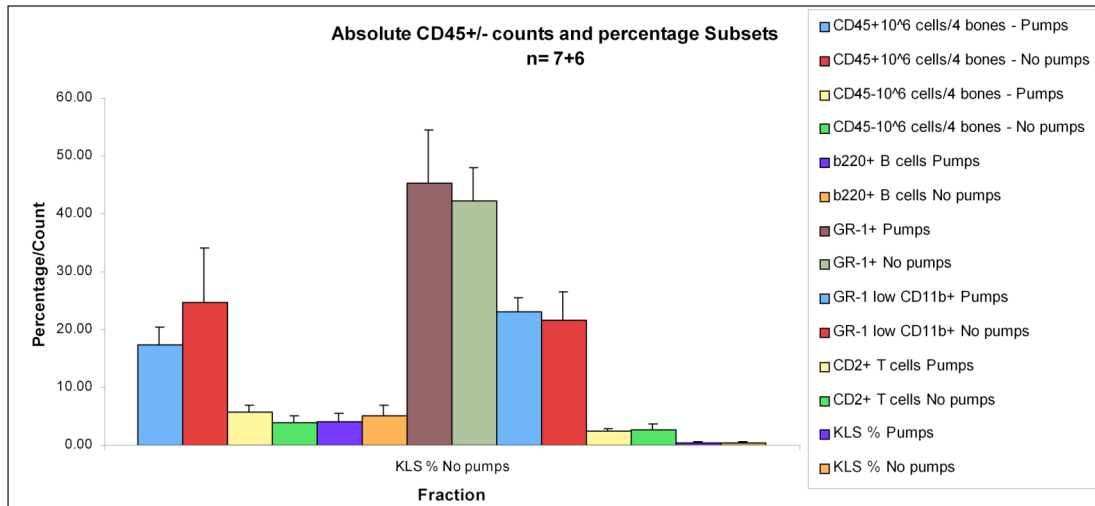


Figure S2: Effect of 1 week of AZD1152 treatment of the frequency of NOD/SCID haematopoietic cell subsets. CD45⁺ cell numbers are expressed as absolute numbers obtained from the four rear murine long bones. All other subsets are expressed as a percentage of live, debris-free CD45⁺ cells. Although there was an overall reduction in CD45⁺ cells, there did not seem to be any particular subset that was preferentially targeted and the mice tolerated this treatment.

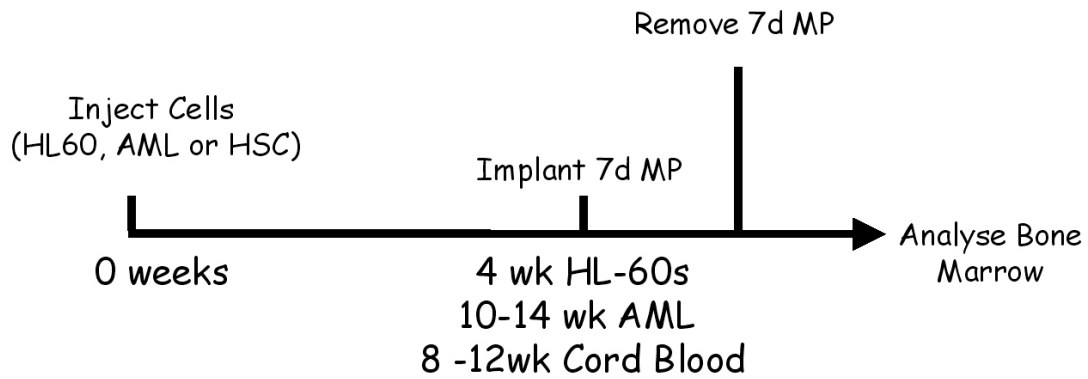


Figure S3: Summary of *in vivo* AZD1152 treatment regimens. A flow chart summary of the AZD1152 treatment timing during *in vivo* experiments. Cells (10^6 HL-60s, 10^7 primary AML cells or 5×10^4 Lin⁻ CB cells) were injected i.v. into sub-lethally irradiated NOD/SCID mice. Mice were then left for differing periods (depending on the cell source) of time before AZD1152 treatment for the xenograft to reach appropriate levels. This was 4 weeks for HL-60s, 10 weeks for primary AML cells and 8 weeks for CB Lin⁻ cells. At these time points, AZD1152 was administered via osmotic mini-pumps at 25mg/Kg/day for one week. At the end of this treatment week, pumps were removed and mice were either left for follow-up studies or sacrificed and analysed for human cell content.

Table S1

Sample ID	FAB type	Karyotype	WBC	% of Blast cells in BM
AML-1	AML M1	NK	151	90
AML-2	AML M2	+11+13	27	89
AML-3	AML M4	NK	43	95
AML-4	AML M5	ND	212	95
AML-5	AML M1	FK	64	91
R 6171	AML M5v	XY, ins(10;11)(p12;q23q21)	162	92
R 6187	AML M5v	XY, ins(10;11)(p12;q23q21)	162	92
R 6290	Biphenotypic AML	NK	200	91
R 3967	AML M4	NK	43	87
R 3921	AML M4	Inv 16	113	83
R 2110	AML M4	NK	127	90
X 7792	AML M4	NK	108	90
R 5940	AML M1	NK	206	91
F 7862	AML M1	NK	79	87
R 4801	AML M3v	XX, t(15;17)(q22;q21)	83	95
R 6298	AML M3v	XX, t(15;17)(q22;q21)	83	95
R 6302	AML M3v	XX, t(15;17)(q22;q21)	83	95

Table S1: Summary of patient details used for *in vivo* experiments. AZD1152 treatment was tested on samples from five different AML cases. Listed are the AML sample identifiers, the AML subtype according to the French-American-British (FAB) classification, the results of the karyotype assessment performed at diagnosis and the white blood cell count (WBC). NK – normal karyotype, ND – not done, FK – failed karyotype - v: represents samples from the same patients used at different time.