

**Supplementary table 1. Additional information on patients' characteristics and transplantation procedure (n = 180).**

<b>Body weight (range)</b>	[kg]	73 (47 - 130)
<b>Phenotype (%)</b>	proB- / C- / preB- / mature B-ALL	16 / 83 / 23 / 9 (9 / 46 / 13 / 5)
	early / thymic / mature T-ALL	23 / 13 / 8 (13 / 7 / 4)
	biphenotypic ALL	5 (3)
	tALL	6 (3)
<b>Therapy prior to allo-SCT (%)</b>	initial treatment in prospective GMALL trial yes / no	42 / 138 (23 / 77)
	<b>auto-SCT</b> yes / no	2 / 178 (1 / 99)
	<b>TKI</b> yes / no	29 / 21 (58 / 42) <sup>1</sup>
	<b>rituximab</b> yes / no	118 / 62 (66 / 34)
	<b>WBI</b> yes / no	86 / 94 (48 / 52)
	<b>mediastinal irradiation</b> yes / no	8 / 172 (4 / 96)
	<b>Salvage therapy in case of relapse (%)</b>	re-induction
	high-dose cytosine arabinoside +/- mitoxantrone	31 (49) <sup>2</sup>
	TKI	4 (6) <sup>2</sup>
	radiation in case of extramedullary recurrence	4 (6) <sup>2</sup>
	blinatumomab	2 (3) <sup>2</sup>
	no / not known	3 (5) <sup>2</sup>
<b>Time between ID and allo-SCT (range) [months]</b>		6.2 (2.0 - 87.7)
<b>Time between relapse and allo-SCT (range) [months]</b>		3.6 (0.9 - 30.4)
<b>Myeloablative conditioning regimen (%)</b>	TBI	47 / 2 / 67 / 3 (39 / 2 / 56 / 3) <sup>3</sup>
	+ etoposide 50mg/kg / + cyclophosphamide 60mg/kg / + both / + other	
	fludarabine 3x30 [25] mg/m <sup>2</sup> + thiotepa 3 [2] x5mg/kg + carmustin 300mg/m <sup>2</sup> / treosulfan 3x10g/m <sup>2</sup>	34 [1] / 1 (56 [2] / 1) <sup>4</sup>
	busulfan only 4x3.2mg/kg / + cyclophosphamide 2x60mg/kg / + fludarabine 4x30mg/m <sup>2</sup>	1 / 9 / 1 (1 / 15 / 1) <sup>4</sup>
	[toxicity-reduced] fludarabine 4x30mg/m <sup>2</sup> + thiotepa 2x5mg/m <sup>2</sup> / + melphalan 140 [110] mg/m <sup>2</sup>	4 / 7 [3] (7 / 12 [5]) <sup>4</sup>
	fludarabine 4x30mg/m <sup>2</sup> + carmustine 2x150mg/m <sup>2</sup> + melphalan 140 [110] mg/m <sup>2</sup>	4 [2] (7 [3]) <sup>4</sup>
<b>Donor sex (%)</b>	male / female	106 / 74 (59 / 41)
<b>CMV-status (%)</b>	donor pos / neg	77 / 103 (43 / 57)
	patient pos / neg	99 / 81 (55 / 45)
<b>Median engraftment (range)</b>	ANC > 500/nl	17 (9 - 32)
	[day after allo-SCT] PLT > 20/nl	13 (7 - 150)
<b>DLI (%)</b>	yes / no	38 / 142 (21 / 79)
<b>Time between allo-SCT and DLI (range) [months]</b>		7.3 (1.0 - 102.4)
<b>Subsequent allo-SCT (%)</b>	yes / no	23 / 157 (13 / 87)

**Abbreviations:** ALL = acute lymphoblastic leukemia; t = therapy-related; GMALL = German ALL study group; auto-SCT = autologous stem cell transplantation; TKI = tyrosine kinase inhibitor; WBI = whole-brain irradiation; ID = initial diagnosis; allo-SCT = allogeneic stem cell transplantation; TBI = total body irradiation; pos = positive; neg = negative; CMV = cytomegalovirus, ANC = absolute neutrophil count; PLT = platelet count; DLI = donor lymphocyte infusion.

<sup>1</sup> as proportion of 50 patients with B-ALL Ph pos; <sup>2</sup> as proportion of 64 relapses; <sup>3</sup> as proportion of 119 patients with TBI; <sup>4</sup> as proportion of 61 patients without TBI.

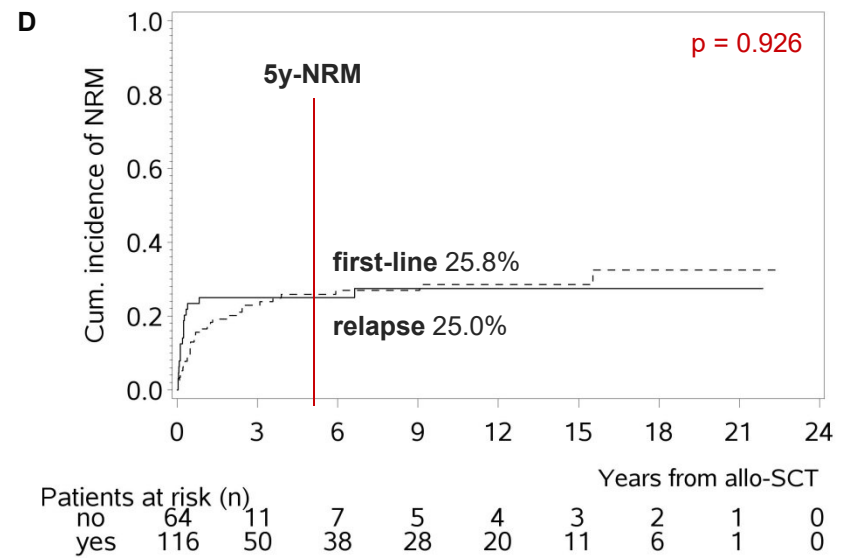
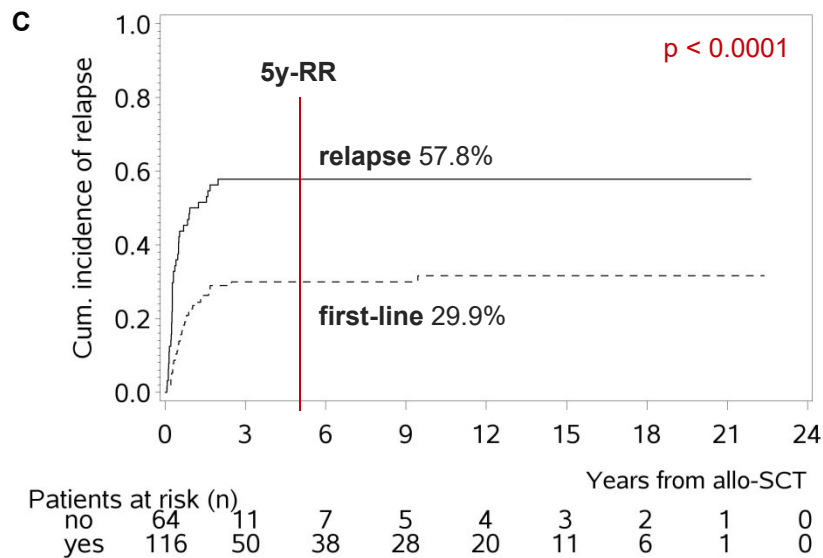
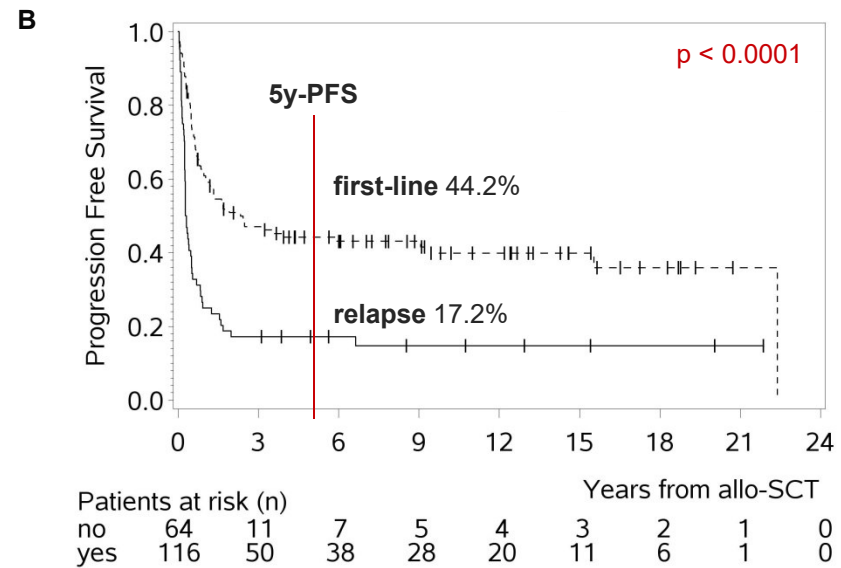
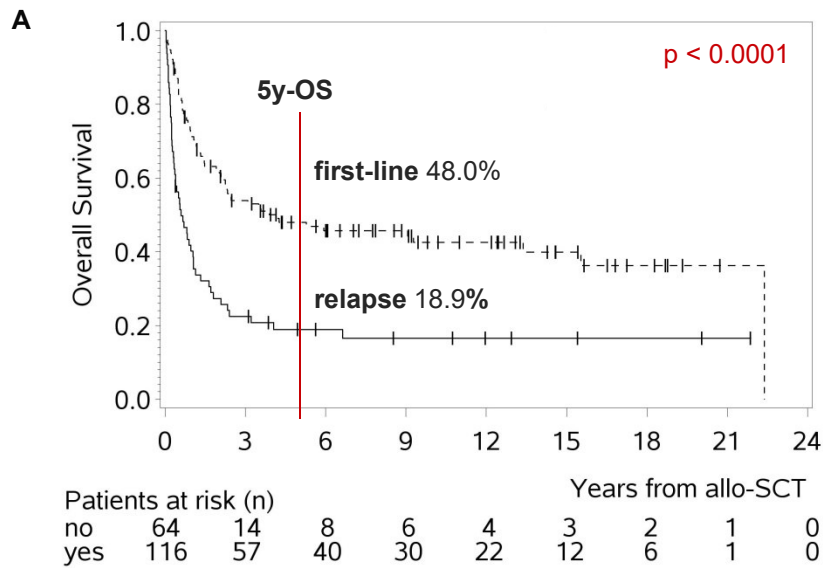
**Supplementary table 2. Patients' characteristics with regard to the timepoint of allo-SCT.**

	<b>1995 - 2000</b> n = 55	<b>2001 - 2010</b> n = 82	<b>2011 - 2018</b> n = 43	
<b>Median age (range) [years]</b>	33 (16 - 57)	45 ( 19 - 69)	35 (18 - 79)	
<b>Phenotype (%)</b>	B-ALL Ph neg	26 (47)	39 (49)	21 (49)
	B-ALL Ph pos	15 (27)	25 (29)	10 (23)
	T-ALL	14 (26)	18 (22)	12 (28)
<b>Remission state at allo-SCT (%)</b>	CR1 MRD neg PCR <sup>1</sup> / FACS <sup>2</sup>	7 / 8 (13 / 14)	16 / 21 (19 / 26)	16 / 1 (37 / 2)
	CR1 MRD pos <sup>1+2</sup>	6 (11)	13 (16)	14 (32)
	primary refractory	3 (5)	8 (10)	3 (7)
	CR2 MRD neg PCR <sup>1</sup> / FACS <sup>2</sup>	0 / 7 (0 / 13)	1 / 5 (1 / 6)	3 / 2 (7 / 4)
	CR2 MRD pos <sup>1+2</sup>	2 (4)	3 (4)	0 (0)
	relapsed refractory	22 (40)	15 (18)	4 (9)
<b>Donor (%) related / unrelated</b>	28 / 27 (51 / 49)	24 / 58 (29 / 71)	7 / 36 (16 / 84)	
<b>Stem cell source (%) BM / PB</b>	23 / 32 (42 / 58)	3 / 79 (4 / 96)	1 / 42 (2 / 98)	
<b>TBI-based conditioning (%) yes / no</b>	43 / 12 (78 / 22)	50 / 32 (61 / 39)	24 / 19 (56 / 44)	
<b>GvHD prophylaxis (%)</b>	CyA	54 (98)	81 (99)	43 (100)
	+ ATG	23 (42)	23 (28)	34 (79)
	+ Alemtuzumab	0 (0)	41 (50)	6 (14)

**Abbreviations:** allo-SCT = allogeneic stem cell transplantation; ALL = acute lymphoblastic leukemia; Ph = Philadelphia chromosome; pos = positive; neg = negative; CR = complete remission; MRD = minimal residual disease; TBI = total body irradiation; BM = bone marrow; PB = peripheral blood; GvHD = graft-versus-host disease; CyA = cyclosporin A; ATG = Anti-thymocyte globulin.

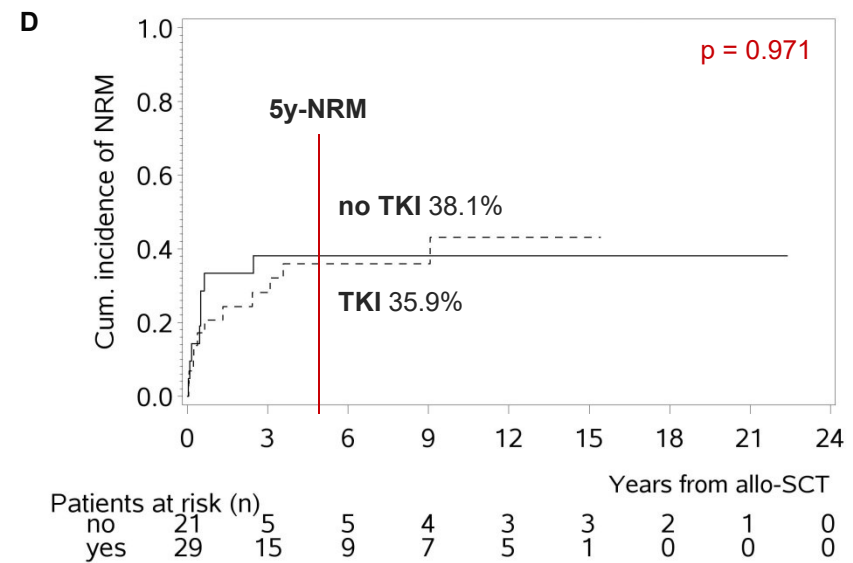
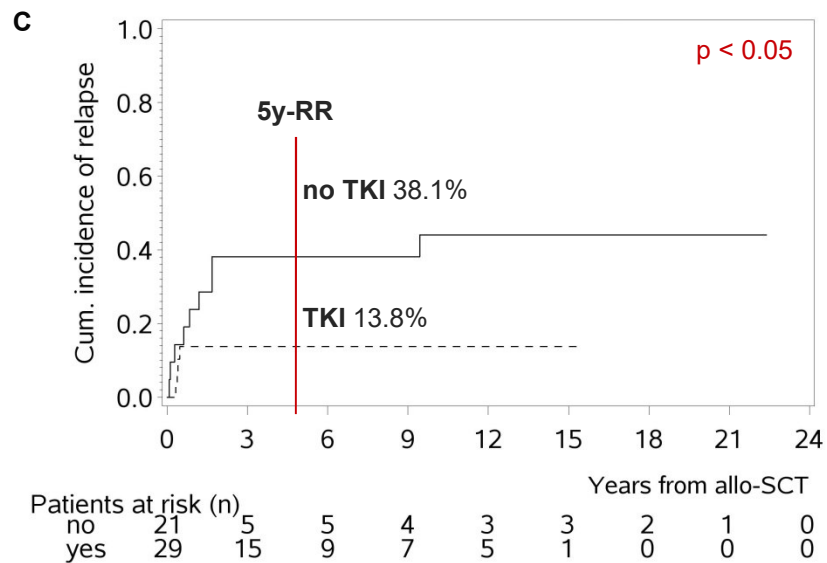
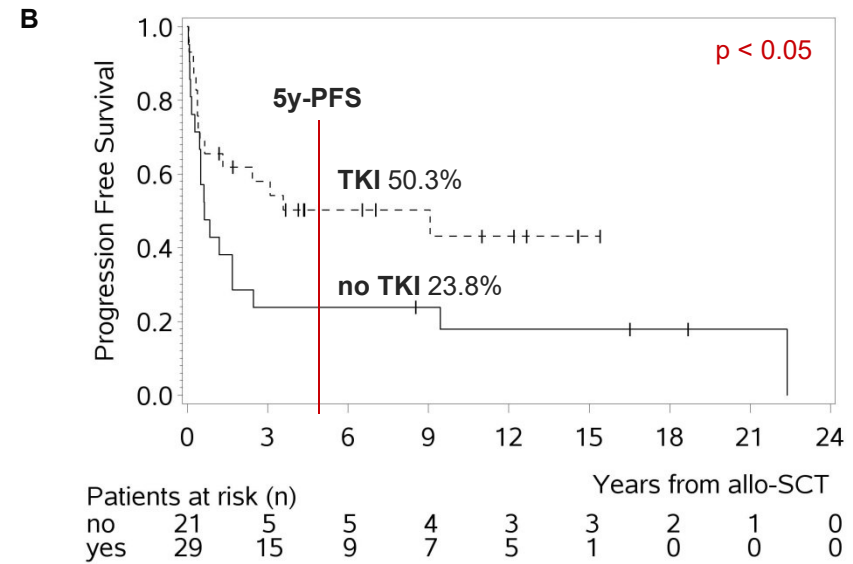
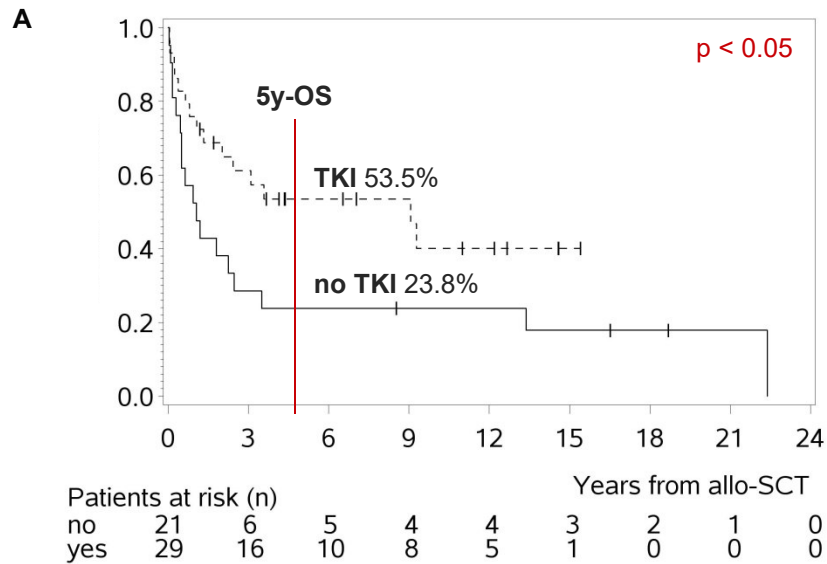
<sup>1</sup> analysis of Ig-/TR-gene rearrangements via rtPCR in 11 (20%), 35 (43%) and 39 (91%) patients, respectively;

<sup>2</sup> analysis via surface marker analysis by flow cytometry in 44 (80%), 47 (57%) and 4 (9%) patients, respectively.



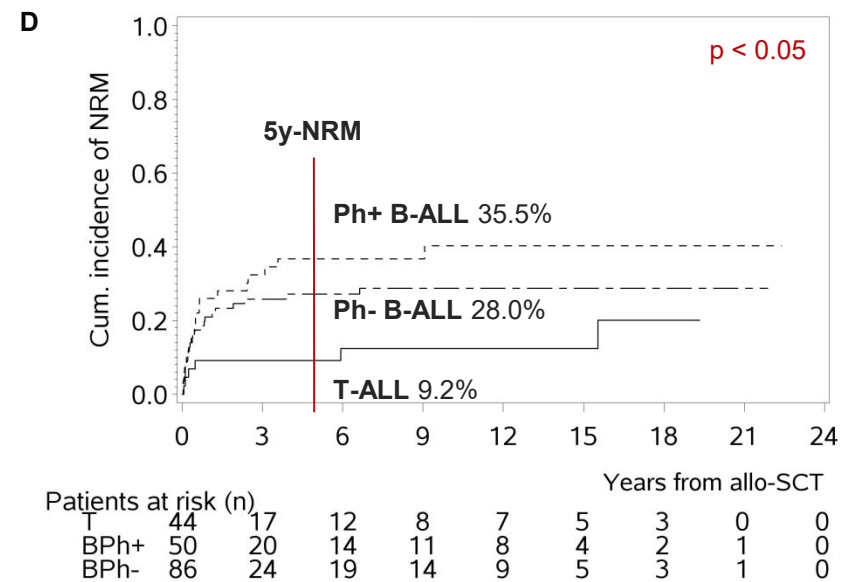
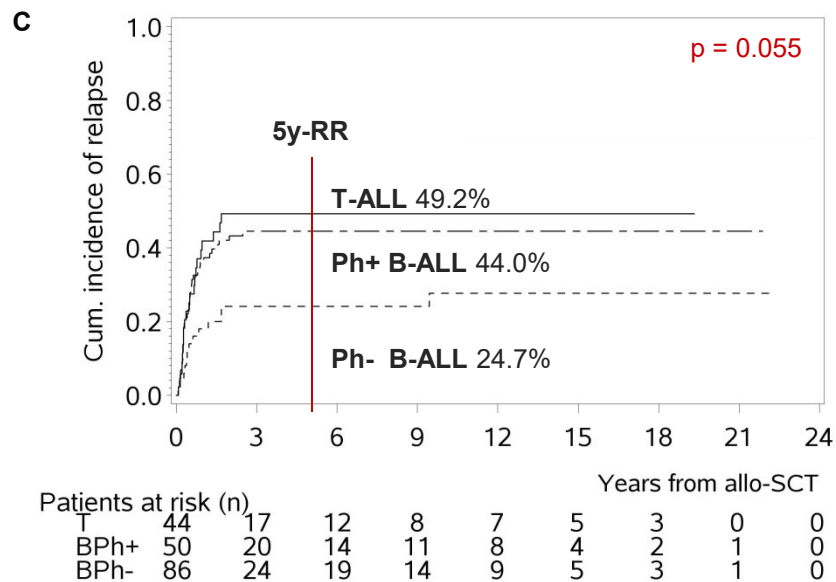
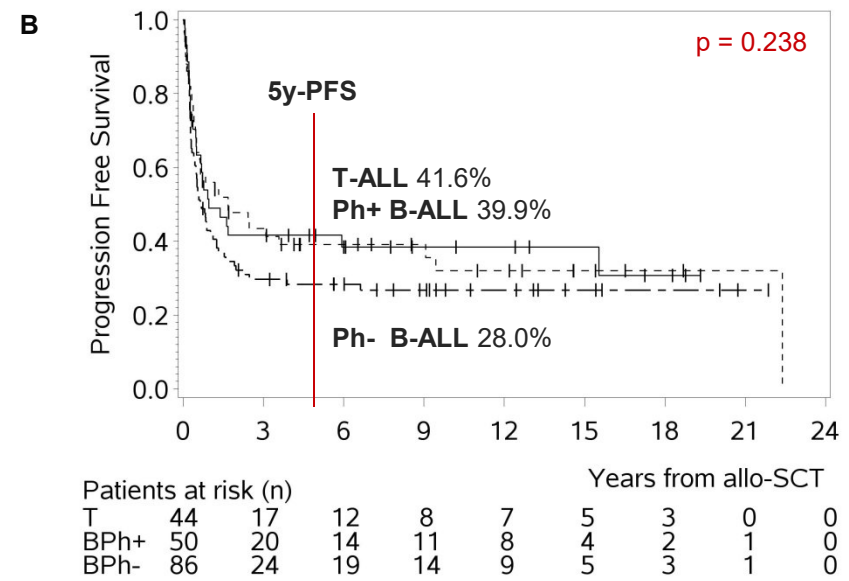
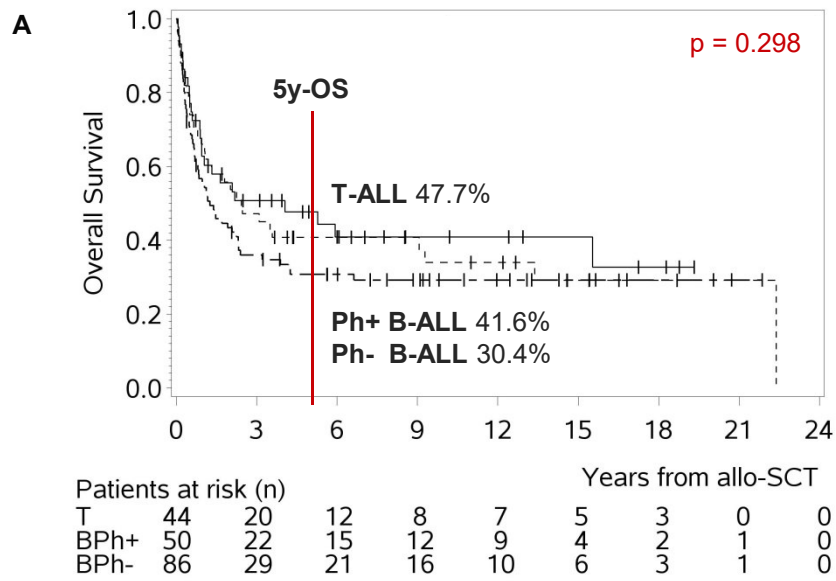
**Supplementary figure 1.** Impact of therapy line on outcome: Patients who received allo-SCT upfront in the course of their first-line therapy (1. CR and primary refractory;  $n = 116$ ) vs. those transplanted in case of relapsed disease (2. CR and relapsed refractory;  $n = 64$ ). **(A)** Kaplan-Meier estimates for OS, **(B)** Kaplan-Meier estimates for PFS, **(C)** cumulative incidence of relapse, **(D)** cumulative incidence of NRM.

**Abbreviations:** allo-SCT = allogeneic transplantation; OS = overall survival; PFS = progression free survival; RR = relapse rate; NRM = non-relapse-related mortality; y = year; CR = complete remission.



**Supplementary figure 2.** Differentiation of patients diagnosed with Philadelphia-positive B-ALL ( $n = 50$ ) treated in the pre-TKI era ( $n = 21$ ) vs. those who received additional therapy with TKI before allo-SCT ( $n = 29$ ). **(A)** Kaplan-Meier estimates for OS, **(B)** Kaplan-Meier estimates for PFS, **(C)** cumulative incidence of relapse, **(D)** cumulative incidence of NRM.

**Abbreviations:** OS = overall survival; PFS = progression free survival; RR = relapse rate; NRM = non-relapse-related mortality; y = year; ALL = acute lymphoblastic leukemia; allo-SCT = allogeneic transplantation; TKI = tyrosine kinase inhibitor.



**Supplementary figure 3.** Impact of phenotype on outcome: Patients diagnosed with T-ALL (n = 44) vs. Philadelphia-positive B-ALL (n = 50) vs. Philadelphia-negative B-ALL (n = 86). **(A)** Kaplan-Meier estimates for OS, **(B)** Kaplan-Meier estimates for PFS, **(C)** cumulative incidence of relapse, **(D)** cumulative incidence of NRM.

**Abbreviations:** OS = overall survival; PFS = progression free survival; RR = relapse rate; NRM = non-relapse-related mortality; y = year; ALL = acute lymphoblastic leukemia; Ph = Philadelphia chromosome.