

TABLE S1 Characterisation of B-ALL patients according to *CRLF2* expression

	<i>CRLF2</i> -low n (%)	<i>CRLF2</i> -high n (%)	<i>p</i> value
Paediatric (<18 years-old)			
Median age (range)	6.2 (0.2-17)	6.1 (2-16)	
Adult (≥18 years-old)			
Median age (range)	31 (18-67)	40.5 (18-82)	
Sex			0.887
Female	36 (36.7)	11 (33.3)	
Male	62 (63.3)	22 (66.7)	
WBC count (x10⁹/L)			0.846
Median (range)	16.5 (0.9-335.4)	16.7 (1.1-493.5)	
<50	72 (73.5)	23 (69.7)	
≥50	26 (26.5)	10 (30.3)	
CNS involvement			0.503
Yes	9 (9.6)	4 (13.8)	
No	85 (90.4)	25 (86.2)	
Glucocorticoid response*			0.333
Good	47 (73.4)	14 (87.5)	
Poor	17 (26.6)	2 (12.5)	
Total	98 (100)	33 (100)	

CNS, Central Nervous System; WBC, white blood cell count;

Paediatric patients were treated with BFM (n=50), EsPhALL (n=2) or RELLA05 (n=32) protocols. Adult patients were treated with BFM (n=20), Hyper-CVAD (n=13), GMALL (n=2), CALB (n=1) or EWALL (n=1) protocols (n=20).

*Prednisone or dexamethasone response, however 45 cases were excluded from this analysis due to treatment strategy (Hyper-CVAD and RELLA05); Good: <1,000 leukaemic blasts/μL on treatment day 7/ 8; Poor: ≥1,000 leukaemic blasts/μL on treatment day 7/ 8. Statistical significance was accessed with Fisher's exact test using the RStudio software version 1.2. *p* values<0.05 were interpreted as significant.

TABLE S2 Overall survival analysis of paediatric B-ALL patients.

RELLA05 cohort					Paediatric cohort				
Variable	Univariate analysis		Multivariate analysis		Variable	Univariate analysis		Multivariate analysis	
	OS (95% CI)*	<i>p</i> value [†]	HR (95% CI)	<i>p</i> value [†]		OS (95% CI)*	<i>p</i> value [†]	HR (95% CI)	<i>p</i> value [†]
<i>KMT2A</i>-rearranged		0.064	1.0 (0.0-128.5)	1.000	Glucocorticoid response		0.015	1.8 (0.8-4.3)	0.163
Negative	67.3 (60.9-73.6)				Poor	53.0 (41.1-64.9)			
Positive	39.6 (0.0-84.5)				Good	68.2 (63.1-73.3)			
<i>CRLF2</i> status		0.067	1.0 (0.0-352.4)	1.000	WBC count (x10⁹/L)		<0.001	2.7 (1.2-6.2)	0.017
Low	69.3 (64.1-74.5)				<50	67.7 (63.0-72.3)			
High	54.9 (34.3-75.5)				≥50	42.3 (25.6-59.0)			
Short isoform		<0.001	1.0 (0.0-22391.3)	1.000	<i>BCR-ABL1</i>		0.065	0.8 (0.3-2.0)	0.619
Low load	6.7 (6.7-6.7)				Negative	64.0 (58.1-69.9)			
High load	69.0 (63.4-74.7)				Positive	45.8 (36.4-55.3)			
<i>CRLF2/IKZF1</i> status		0.067	1.0 (0.0-128.6)	1.000					
<i>CRLF2</i> -low/ <i>IKZF1</i> wt	68.9 (63.0-74.8)								
<i>CRLF2</i> -low/ <i>IKZF1</i> del [‡]	-								
<i>CRLF2</i> -high/ <i>IKZF1</i> wt	61.1 (41.7-80.6)								
<i>CRLF2</i> -high/ <i>IKZF1</i> del	36.1 (0.0-85.9)								

CI, confidence interval. del, deleted, HR, hazard ratio; OS, overall survival, wt, wild-type.

*Mean for survival time. †Log-rank test.

‡No statistics are computed for this group because all cases are censored.

TABLE S3 Overall survival analysis of adult B-ALL patients.

Variable	Univariate analysis		Multivariate analysis	
	OS (95% CI)*	<i>p</i> value [†]	HR (95% CI)	<i>p</i> value [†]
<i>KMT2A</i>-rearranged		0.007	61.3 (2.1-1778.8)	0.017
Negative	38.1 (27.2-49.1)			
Positive	4.2 (0.0-8.5)			
<i>CRLF2</i> status		0.072	0.1 (0.0-5.0)	0.266
Low	40.8 (27.9-53.7)			
High	26.5 (11.5-41.6)			
<i>IKZF1</i> status		0.042	34.6 (1.7-724.6)	0.022
Wild-type	52.0 (35.7-68.4)			
Deleted	29.6 (16.4-42.7)			
<i>IKZF1</i> plus		0.007	1.2 (0.1-9.7)	0.858
Wild-type	52.0 (35.7-68.4)			
Non-plus	8.9 (1.9-16.0)			
Plus	30.5 (12.4-48.5)			
<i>CRLF2/IKZF1</i> status		0.094	0.7 (0.1-4.9)	0.724
<i>CRLF2</i> -low/ <i>IKZF1</i> wt	47.6 (32.4-62.7)			
<i>CRLF2</i> -low/ <i>IKZF1</i> del	18.6 (9.9-27.3)			
<i>CRLF2</i> -high/ <i>IKZF1</i> wt	9.7 (0.0-20.7)			
<i>CRLF2</i> -high/ <i>IKZF1</i> del	29.7 (11.8-47.5)			
Short isoform		0.006	0.9 (0.0-33.1)	0.960
Low load	21.0 (9.0-33.1)			
High load	49.3 (35.6-63.1)			
<i>CRLF2</i>/Short isoform		0.009	68.1 (0.4-11176.7)	0.105
<i>CRLF2</i> -low/Low load	24.8 (9.1-40.5)			
<i>CRLF2</i> -low/High load	50.6 (31.6-69.7)			
<i>CRLF2</i> -high/Low load	8.6 (1.2-15.9)			
<i>CRLF2</i> -high/High load	46.9 (26.8-67.1)			

CI, confidence interval. del, deleted, HR, hazard ratio; OS, overall survival, wt, wild-type.
 *Mean for survival time. †Log-rank test