## SUPPLEMENTARY APPENDIX

## Spleen tyrosine kinase inhibitors reduce CD40L-induced proliferation of chronic lymphocytic leukemia cells but not normal B cells

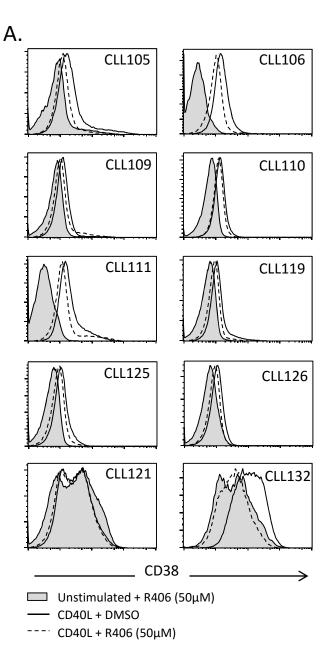
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B.	MFI: Unstim + R406 (50μM)	MFI: CD40L + DMSO	MFI: CD40L + R406 (50μM)
CLL105	18	62	22
CLL106	3.7	19	15
CLL109	11	19	19
CLL110	7,5	16	13
CLL111	4.4	32	33
CLL119	6.8	28	12
CLL125	6.4	17	11
CLL126	7	12	9.2
CLL121	63	56	48
CLL132	79	170	62

## **Supplementary Figure S1**

Expression of the activation marker CD38 on CLL cells after CD40L stimulation and Syk inhibition. CLL cells from 10 patients were stimulated by irradiated CD40L<sup>+</sup> or untransfected L cells (unstimulated) in the presence of R406 using relevant *in vivo* concentrations or DMSO vehicle control. See also Figure 1C.

- (A) Expression of the activation marker CD38 on CD19<sup>+</sup> CLL cells is shown in unstimulated condition, after CD40L stimulation with vehicle (DMSO) only, or after CD40L stimulation in the presence of R406.
- **(B)** Mean fluorescent intensity of CD38-expression in CD19<sup>+</sup>CLL cells as shown in (A).

## Supplemental Table SI. Patient information.

Patient identifier	Gender/age (years)*	Binet stage*	IgVH,% homology to germline <sup>1</sup>	CD38 (%) <sup>2</sup>	FISH	Observation time before treatment / Treatment (T) <sup>3</sup>
CLL105	F/56	A	VH3-23, 100 VH3-49, 94.8 (biallelic)	0	NA	96/Not T
CLL106	F/52	A	VH3-23, 95.2	1	NA	49/Not T
CLL107	M/38	A	VH3-30, 92.9	1	del(13q14)	84/Not T
CLL109	M/54	A	VH3, 93.8	1	NA	61/Not T
CLL110	M/49	A	VH6-01, 95.4	1	NA	89/Not T
CLL111	M/59	A	VH1-69, 91	2	NA	48/Not T
CLL114	M/55	A	VH3-23, 88.2	2	NA	78/Not T
CLL119	M/56	С	VH3-21, 98,7	NA	Normal karyotype	5/FCA
CLL121	M/64	С	VH4-39, 98,7	61	Normal karyotype	1/FCA
CLL125	M/44	A at diagnosis, progressed to C	VH1-18 98,1/ VH3-30 97,3 (biallelic)	0	Normal karyotype	10/FCA
CLL126	F/55	С	VH4-34, 95.4	1	del(13q14)	F-FC-FCR/A
CLL132	M/51	В	VH3-30, 100	94	del(13q14)	29/FC
CLL135	M/61	A at diagnosis, progressed to C	VH4-34, 100	50	Normal karyotype	36/FCR-A-BR
CLL136	F/66	A	VH3-30, 91,9	1	del(13q14)	52/Not T

<sup>\* :</sup>at diagnosis

NA: Not analyzed.

<sup>&</sup>lt;sup>1</sup> IgVH family, percent homology to germline.

<sup>&</sup>lt;sup>2</sup> Percentage of cells staining positive for CD38, above isotype control.

Observation time, treatment-free months since diagnosis. Non-treated patients are marked with Not T. Treated patients have received regimens with the following drugs marked with abbreviations: F= Fludarabine; C = Cyclophosphamide; A=Alemtuzumab; R=Rituximab; BR=bendamustine. Cells were gathered before treatment except for CLL126 and CLL135. Patient CLL126 was initially treated with F, thereafter with FC, then FCR (aborted due to toxicity), and replaced by A monotherapy. Samples for CLL126 were gathered prior to FCR treatment. Patient CLL135 was initially treated with FCR, then A and finally BR. Samples were gathered between 2nd and 3rd treatment.