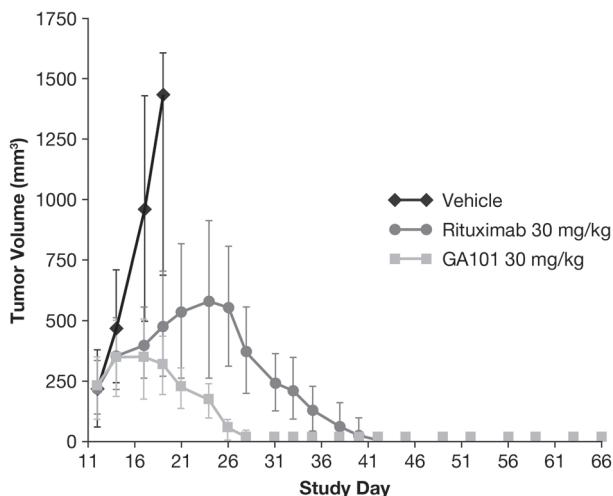
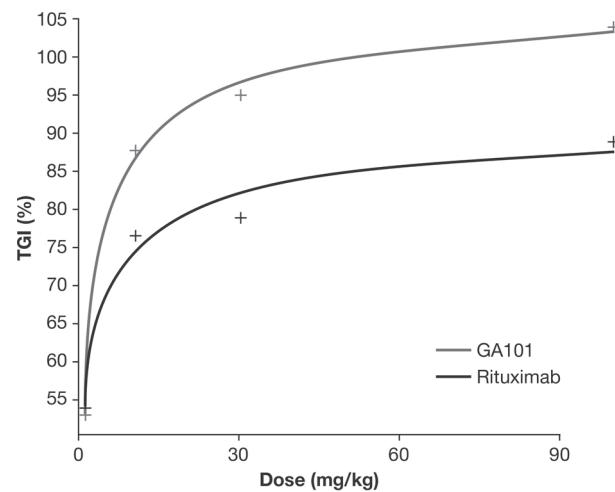


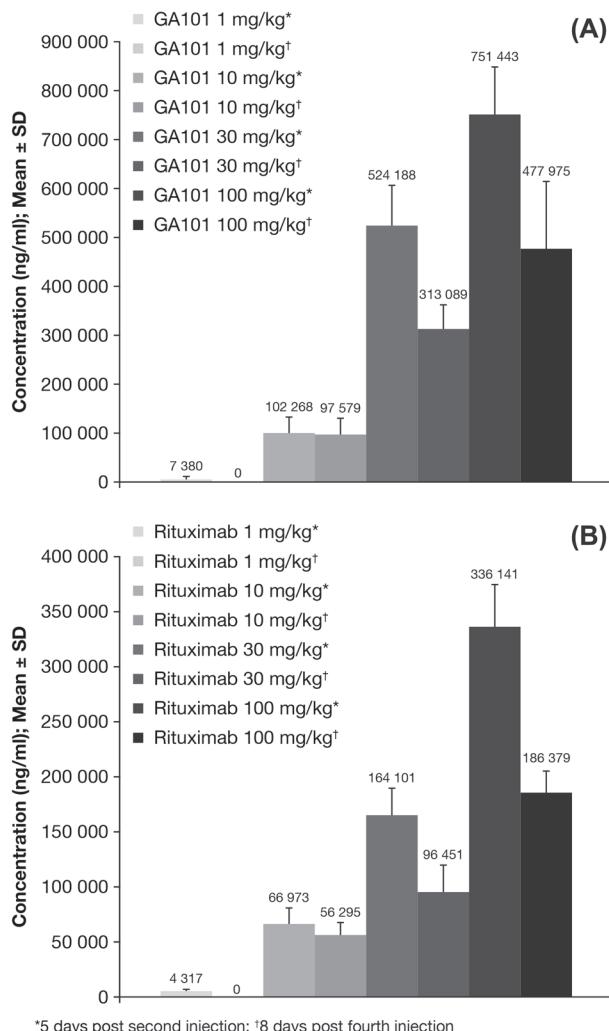
Supplementary material for Herting F, et al. Enhanced anti-tumor activity of the glycoengineered type II CD20 antibody obinutuzumab (GA101) in combination with chemotherapy in xenograft models of human lymphoma. Leuk Lymphoma 2014; 55:2151–2160.



Supplementary Figure 1. Tumor size during treatment with rituximab or obinutuzumab (GA101) (30 mg/kg i.p. x3) in single-agent Z138 when tumor load reached a volume of 220 mm³.



Supplementary Figure 3. Tumour growth inhibition (%) at Day 31 following treatment with obinutuzumab (GA101) and rituximab at various dose concentrations. TGI, tumor growth inhibition.



*5 days post second injection; †8 days post fourth injection

Supplementary Figure 2. Serum levels of (A) anti-CD20 antibody obinutuzumab (GA101) and (B) rituximab at various dose concentrations. SD, standard deviation.

Supplementary Table I. Treatment schedules by chemotherapy combination partner.

	n	Treatment schedule	Median tumor volume at staging (mm ³)
Single-agent studies in Z138 xenograft			
Single-agent obinutuzumab (GA101) or rituximab			
Z138 (mantle cell lymphoma) tumor xenograft	10	Vehicle	214
Treatment initiation 14 days post cell transplantation			
Study termination Day 35–78	10	Rituximab 10 mg/kg i.v. q7d × 4	207
	10	Obinutuzumab (GA101) 10 mg/kg i.v. q7d × 4	211
Single-agent obinutuzumab (GA101) or rituximab			
Z138 (mantle cell lymphoma) tumor xenograft	10	Vehicle	-
Treatment initiation 12 days post cell transplantation			
Study termination Day 21–66	10	Rituximab 30 mg/kg i.p. × 3	-
	10	Obinutuzumab (GA101) 30 mg/kg i.p. × 3	-
Single-agent obinutuzumab (GA101) or rituximab			
Z138 (mantle cell lymphoma) tumor xenograft	10	Vehicle	397
Treatment initiation 20 days post cell transplantation			
Study termination Day 49	10	Rituximab 1 mg/kg i.v. q7d × 4	396
	10	Rituximab 10 mg/kg i.v. q7d × 4	396
	10	Rituximab 30 mg/kg i.v. q7d × 4	401
	10	Rituximab 100 mg/kg i.v. q7d × 4	406
	10	Obinutuzumab (GA101) 1 mg/kg i.v. q7d × 4	402
	10	Obinutuzumab (GA101) 10 mg/kg i.v. q7d × 4	394
	10	Obinutuzumab (GA101) 30 mg/kg i.v. q7d × 4	410
	10	Obinutuzumab (GA101) 100 mg/kg i.v. q7d × 4	393
Single-agent obinutuzumab (GA101)			
Z138 (mantle cell lymphoma) tumor xenograft	10	Vehicle	201
Treatment initiation 14 days post cell transplantation			
Study termination Day 40	10	Obinutuzumab (GA101) 10 mg/kg i.v. q7d × 4	203
	10	Obinutuzumab (GA101) WT 10 mg/kg i.v. q7d × 4	202
Combination studies in Z138 xenograft			
Bendamustine combination			
Z138 (mantle cell lymphoma) tumor xenograft	10	Vehicle	442
Treatment initiation 19 days post cell transplantation			
Study termination Day 33	10	Obinutuzumab (GA101) 1 mg/kg i.p., Days 19, 26	433
	10	Rituximab 1 mg/kg i.p., Days 19, 26	455
	10	Bendamustine 3 mg/kg i.p., Days 19, 20, 21, 22	443
	10	Obinutuzumab (GA101) 1 mg/kg i.p. Days 19, 26 + bendamustine 3 mg/kg i.p., Days 19, 20, 21, 22	446
	10	Rituximab 1 mg/kg i.p., Days 19, 26 + bendamustine 3 mg/kg i.p., Days 19, 20, 21, 22	431
Fludarabine combination			
Z138 (mantle cell lymphoma) tumor xenograft	10	Vehicle	399
Treatment initiation 22 days post cell transplantation			
Study termination Day 43	10	Obinutuzumab (GA101) 1 mg/kg i.v., Days 22, 29	411
	10	Rituximab 1 mg/kg i.v., Days 22, 29	406
	10	Fludarabine 40 mg/kg i.p., Days 22, 23, 24	408
	10	Obinutuzumab (GA101) 1 mg/kg i.v., Days 22, 29 + fludarabine 40 mg/kg i.p., Days 22, 23, 24	403
	10	Rituximab 1 mg/kg i.v., Days 22, 29 + fludarabine 40 mg/kg i.p., Days 22, 23, 24	416
Chlorambucil combination			
Z138 (mantle cell lymphoma) tumor xenograft	10	Vehicle	272
Treatment initiation 27 days post cell transplantation			
Study termination Day 41			

(Continued)

Supplementary Table I. (Continued)

	n	Treatment schedule	Median tumor volume at staging (mm ³)
Single-agent studies in WSU-DLCL2 xenograft			
Single-agent obinutuzumab (GA101) or cyclophosphamide/vincristine/doxorubicin	10	Obinutuzumab (GA101) 1 mg/kg i.v., Days 27, 34	269
WSU-DLCL2 lymphoma xenograft	10	Rituximab 1 mg/kg i.v., Days 27, 34	281
Treatment initiation 9 days post cell transplantation	10	Chlorambucil 4 mg/kg i.p., Days 27, 28, 29	273
Study termination Day 30	10	Obinutuzumab (GA101) 1 mg/kg i.v., Days 27, 34 + chlorambucil 4 mg/kg i.p., Days 27, 28, 29	274
10	Rituximab 1 mg/kg i.v., Days 27, 34 + chlorambucil 4 mg/kg i.p., Days 27, 28, 29	274	
Combination studies in WSU-DLCL2 xenograft			
Cyclophosphamide/vincristine combination	10	Vehicle	143
WSU-DLCL2 lymphoma xenograft	10	Cyclophosphamide 25 mg/kg plus vincristine 0.25 mg/kg i.v. q7d × 3	140
Treatment initiation 9 days post cell transplantation	10	Doxorubicin 3 mg/kg i.v. q7d × 3	143
Study termination Day 35–64	10	Cyclophosphamide 25 mg/kg plus vincristine 0.25 mg/kg i.v. q7d × 3 plus doxorubicin 3 mg/kg i.v. q7d × 3	140
10	Obinutuzumab (GA101) 30 mg/kg i.v. q7d × 3	143	
10	Obinutuzumab (GA101) 100 mg/kg i.v. q7d × 3	144	
Combination studies in WSU-DLCL2 xenograft			
Cyclophosphamide/vincristine combination	10	Vehicle	115
WSU-DLCL2 lymphoma xenograft	10	Rituximab 30 mg/kg i.v., Days 9, 15, 23, 30, 37, 44, 51, 58	114
Treatment initiation 9 days post cell transplantation	10	Obinutuzumab (GA101) 30 mg/kg i.v., Days 9, 15, 23, 30, 37, 44, 51, 58	114
Study termination Day 35–64	10	CYC/VINC 25/0.25 mg/kg i.v., Days 9, 15, 23, 30, 37, 44, 51, 58	115
10	Obinutuzumab (GA101) 30 mg/kg i.v. + CYC/VINC 25/0.25 mg/kg i.v., Days 9, 15, 23, 30, 37, 44, 51, 58	112	
10	Rituximab 30 mg/kg i.v. + CYC/VINC 25/0.25 mg/kg i.v., Days 9, 15, 23, 30, 37, 44, 51, 58	112	
10	CYC/VINC 25/0.25 mg/kg i.v., Days 9, 15, 23 then obinutuzumab (GA101) 30 mg/kg i.v. Days 30, 37, 44, 51, 58	115	
10	CYC/VINC 25/0.25 mg/kg i.v., Days 9, 15, 23 then rituximab 30 mg/kg i.v., Days 30, 37, 44, 51, 58	113	
10	CYC/VINC 25/0.25 mg/kg i.v., Days 9, 15, 23 then CYC/VINC 25/0.25 mg/kg i.v. + obinutuzumab (GA101) 30 mg/kg i.v., Days 30, 37, 44, 51, 58	113	
10	CYC/VINC 25/0.25 mg/kg i.v., Days 9, 15, 23 then CYC/VINC 25/0.25 mg/kg i.v. + rituximab 30 mg/kg i.v., Days 30, 37, 44, 51, 58	113	

CYC, cyclophosphamide; i.p., intraperitoneal; i.v., intravenous; q7d, every 7 days; VINC, vincristine.