

Table S1. List of biologics already approved by the FDA for the treatment of SLE, SSc and RA.

Agent	Action cells	Disease	Date of FDA approval	Clinical trials code / Reference number
<i>Belimumab</i>	Anti-BAFF, IgG1 moAb that specifically binds to the soluble form of human B-cell-stimulating protein	SLE	2011	NCT04956484 NCT01597492 NCT01632241
<i>Anifrolumab</i>	MoAb, Anti-IFN- α receptor	SLE	2021	NCT02794285 NCT02446899 NCT02446912
<i>Rituximab</i>	Chimeric anti-CD20 moAb. Eliminates B cells, and a subset of T cells	RA	2006	NCT02731560 NCT05116228
<i>Abatacept</i>	Protein, CTLA4-Ig construct. CD80/86 costimulation inhibitor, selectively modulates a key co-stimulatory signal for the full activation of T lymphocytes expressing CD28. Inhibits activated T cells, and antigen-presenting cells.	RA	2005	NCT03882008 NCT01758198 NCT03492658
<i>Tocilizumab</i>	moAb against IL-6R. Inhibits B cell differentiation, M2 macrophages, TH17 polarization, and myofibroblast activation	RA	2010	NCT02659150 NCT02010216 NCT01120366
<i>Sarilumab</i>	human anti-IL-6R moAb	RA	2017	NCT03449758
<i>Anakinra</i>	IL-1 inhibitor, it's a recombinant and slightly modified version of the human interleukin 1 receptor antagonist protein	RA	2001	NCT02236481

<i>Adalimumab</i>	moAb, works by inactivating tumor necrosis factor-alpha (TNF α)	RA	2002	NCT01270087 NCT00234897
<i>Infliximab</i>	chimeric human-mouse anti-(TNF α) moAb	RA	1998	NCT02770794 NCT00246064
<i>Certolizumab pegol</i>	Fragment of moAb, anti-(TNF α)	RA	2009	NCT00580840
<i>Etanercept</i>	It's a fusion protein produced by recombinant DNA, TNF inhibitor	RA	1998	NCT00913458 NCT00768053 NCT00099554 NCT01783015
<i>Golimumab</i>	Anti-TNF α human moAb	RA	2009	NCT01295151 NCT03100253

Table S2. Biologic drugs for treating SLE, SSc and RA that are currently under clinical trials

Agent	Action, Cells	Disease	Phase of trial	Clinical trials code
B-cell activity inhibitors				
<i>Blisibimod</i>	Anti-BAFF fusion protein consisting of four BAFF binding domains	SLE	III	NCT01395745 NCT02514967
<i>Obexelimab (XmAb5871)</i>	BSAb, targeting CD19 and Fc γ RIIb on B cells	SLE	II	NCT02725515
<i>Epratuzumab</i>	Anti-CD22 human moAb	SLE	III	NCT01262365 NCT01408576

				NCT00383214 NCT00111306
<i>Obinutuzumab</i>	Human anti CD20 moAb	SLE	III	NCT04963296
<i>Mosunetuzumab</i>	BsAb, binds CD20 and CD3 to engage T-cells	SLE	I	NCT05155345
<i>PRV-3279</i>	Fv specific BsAb, simultaneously ligate the inhibitory molecule CD32B (FcγRIIb) with the B-cell receptor component CD79B on B cells	SLE	II	NCT05087628
<i>Telitacicept</i>	Fully human TACI-Fc fusion protein that targets B lymphocyte stimulator (BLyS) and a proliferating-inducing ligand (APRIL)	SLE	IV	NCT05899907
<i>Ocrelizumab</i>	Anti CD20 moAb	SLE	III	NCT00539838 NCT00626197
<i>Abetimus</i>	It is made of four double-stranded oligodeoxyribonucleotides that are attached to a carrier platform and are designed to block specific B-cell anti double stranded DNA antibodies	SLE	III	NCT00089804 NCT00035308
<i>Rozibafusp Alfa (AMG-570)</i>	BsAb, targeting inducible costimulator ligand (ICOSL) and B cell activating factor (BAFF)	SLE	II	NCT04058028
		RA	I	NCT03156023
<i>Ianalumab</i>	MoAb, anti-BAFFR	SLE	III	NCT05639114 NCT05624749
		RA	I	NCT03574545
<i>Atacicept</i>	Recombinant fusion protein, anti-BLyS and APRIL	SLE	III	NCT00624338
		RA	II	NCT00430495 NCT00595413
<i>Tabalumab</i>	Anti-BAFF moAb	SLE	III	NCT02041091

		RA	III	NCT01676701 NCT01202760
<i>Rituximab</i>	Chimeric anti-CD20 moAb. Eliminates B cells, and a subset of T cells	SLE	IV	NCT05828147
		SSc	III	NCT01748084 NCT04274257
<i>Brentuximab vedotin</i>	Anti-CD30 moAb	SLE	II	NCT02533570
		SSc	III	NCT03222492 NCT03198689
<i>Inebilizumab</i>	anti-CD19 monoclonal antibody	SSc	III	NCT05198557
<i>Belimumab</i>	Anti-BAFF, IgG1 moAb that specifically binds to the soluble form of human B-cell-stimulating protein	SSc	II/III	NCT05878717 NCT0384406 NCT01670565
T-cell or T-cell co-stimulator blockers				
<i>Itolizumab</i>	Anti-CD6 moAb	SLE	I	NCT04128579
<i>Iscalimab</i>	Anti-CD40 moAb	SLE	II	NCT03656562
<i>Milatumzumab</i>	Anti-CD74 moAb	SLE	I	NCT01845740
<i>Lulizumab pegol</i>	Anti-CD28 moAb	SLE	II	NCT02265744
<i>Acazicolcept</i>	Anti-CD28 and ICOS BsAb	SLE	II	NCT04835441
<i>LY3361237</i>	Agonist moAb to the checkpoint inhibitory receptor BTLA (B and T lymphocyte attenuator)	SLE	II	NCT03933943
<i>Dapirolizumab pegol</i>	Anti-CD40L moAb	SLE	III	NCT04294667
<i>Dazodalibep (VIB4920)</i>	anti-CD40 ligand-Tn3 fusion protein	SLE	II	NCT02804763
		RA	II	NCT04163991 NCT02780388 NCT04129164

BI655064	Anti-CD40 moAb	SLE	II	NCT03385564 NCT02770170
		RA	I	NCT01751776
Abatacept	Protein, CTLA4-Ig construct. CD80/86 costimulation inhibitor, selectively modulates a key co-stimulatory signal for the full activation of T lymphocytes expressing CD28. Inhibits activated Tcells, and antigen-presenting cells.	SLE	III	NCT00430677
		SSc	II	NCT00442611 NCT02161406
Romilkimab	BsAb against IL-4/IL-13 cytokines, which are derived from Th2 cells, mast cells eosinophils, innate lymphoid cells. Inhibits M2 macrophages, and myofibroblasts.	SSc	II	NCT02921971
Theralizumab (TAB08)	anti-CD28 moAb	RA	II	NCT01990157
Targeting cytokine or cytokine receptor				
Sifalimumab	Anti-IFN- α moAb	SLE	II	NCT00979654 NCT01283139
Rontalizumab	Anti-IFN- α moAb	SLE	II	NCT00962832
JNJ-55920839	Anti-IFN- α moAb	SLE	I	NCT02609789
IFN-α kinoid	A therapeutic vaccine composed of IFN α 2b coupled to a carrier protein	SLE	II	NCT02665364
BT-063	Anti-IL-10 moAb	SLE	II	NCT02554019
Vunakizumab (SHR1314)	Anti-IL17A moAb	SLE	II	NCT04924296
Avizakimab (BOS161721)	Anti-IL21 moAb	SLE	II	NCT03371251
Infliximab	Chimeric human-mouse anti-(TNF α) moAb	SLE	II/III	NCT00368264

<i>Etanercept</i>	It's a fusion protein produced by recombinant DNA, TNF inhibitor	SLE	II	NCT02656082
<i>Ustekinumab</i>	MoAb that binds to the p-40 subunit of both IL-12 and IL-23 so that they subsequently cannot bind to their receptors to activate T-cells	SLE	III	NCT03517722
		RA	II	NCT01645280
<i>Secukinumab</i>	anti-IL17A moAb	SLE	III	NCT05232864 NCT04181762
		RA	III	NCT01770379 NCT01377012 NCT01350804
<i>ALX-0061</i>	anti-IL-6R BsAb	SLE	II	NCT02437890
		RA	II	NCT02518620 NCT01284569
<i>Guselkumab</i>	anti-IL23 moAb	SLE	II	NCT04376827
		SSc	II	NCT01645280
<i>Tocilizumab</i>	moAb against IL-6R. Inhibits B cell differentiation, M2 macrophages, TH17 polarization, and myofibroblast activation	SLE	I	NCT05835986
		SSc	III	NCT01532869 NCT02453256 NCT05963048
<i>Brodalumab (KHK4827)</i>	anti-IL17A moAb	SSc	III	NCT03957681
<i>Bermekimab</i>	anti-IL-1a mAb	SSc	II	NCT04045743
<i>Nemolizumab</i>	anti-IL-31 mAb	SSc	II	NCT05214794
<i>Fresolimumab</i>	anti-TGFβ mAb	SSc	I	NCT01284322

<i>Sirukumab</i>	anti-IL-6 moAb	RA	III	NCT01856309 NCT02019472
<i>Clazakizumab</i>	anti-IL-6 moAb	RA	II	NCT02015520
<i>Olokizumab</i>	anti-IL-6 moAb	RA	III	NCT02760433 NCT02760368 NCT03120949
<i>Anifrolumab</i>	MoAb, Anti-IFN- α receptor	SSc	III	NCT05925803
		RA	II	NCT01438489
Drugs targeting the plasmacytoid dendritic cells (pDCs)				
<i>Litifilimab</i>	Anti-BDCA2 (Blood dendritic cell antigen 2) moAb	SLE	II	NCT02847598
<i>Daxdilimab</i>	Anti-Immunoglobulin-Like Transcript (ILT)-7 moAb, which specifically induces the depletion of pDCs	SLE	II	NCT04925934

Table S3. Common biologic agents used to treat SLE, SSc, and RA

Agent	Action, Cells	Lupus	Sclerosis	RA
<i>Rituximab</i>	Chimeric anti-CD20 moAb. Eliminates B cells, and a subset of T cells	Phase IV	Phase III	Phase IV Approved by FDA in 2006
<i>Abatacept</i>	Protein, CTLA4-Ig construct. CD80/86 costimulation inhibitor, selectively modulates a key co-stimulatory signal for the full activation of T lymphocytes expressing CD28. Inhibits activated T cells, and antigen-presenting cells.	Phase III	Phase II	Phase IV Approved by FDA in 2005

<i>Anifrolumab</i>	MoAb, Anti-IFN- α receptor	Phase III Approved by FDA in 2021	Phase III	Phase II
<i>Tocilizumab</i>	moAb against IL-6R. Inhibits B cell differentiation, M2 macrophages, TH17 polarization, and myofibroblast activation	Phase I	Phase III	Phase IV Approved by FDA in 2010