Supplementary material

Supplementary Table S1. Search strategy

Ovid	Population
MEDLINE(R)	exp lupus erythematosus, systemic/ OR (lupus adj3 (erythematosus or
Epub Ahead of	erythematous or erythematosis) adj3 (disseminatus or systemic)).ti,ab,kf. OR
Print, In-Process	(libman sacks adj3 (disease or endocarditis or syndrom*)).ti,ab,kf. OR lupus.ti.
& Other Non-	OR (SLE and (lupus or autoimmun* or auto-immun*)).ti,ab,kf,hw.
Indexed	
Citations, Ovid	AND
MEDLINE(R)	
Daily and Ovid	Outcomes
MEDLINE(R):	cerebrovascular disorders/ or exp basal ganglia cerebrovascular disease/ or exp
1946 to 20 May	brain ischemia/ or exp carotid artery diseases/ or exp intracranial arterial
2020	diseases/ or exp intracranial hemorrhages/ or stroke/ or exp brain infarction/
	or exp vertebral artery dissection/ OR exp "Intracranial Embolism and
	Thrombosis"/ OR (stroke or cerebrovasc* or brain vasc* or cerebral vasc* or
	cva* or apoplex*).ti,ab,kf. OR ((brain* or cerebr* or cerebell* or
	vertebrobasilar or hemispher* or intracran* or intracerebral or infratentorial
	or supratentorial or MCA or anterior circulation or posterior circulation or basal
	ganglia) adj5 (isch?emi* or infarct* or thrombo* or emboli*)).ti,ab,kf. OR
	((brain* or cerebr* or cerebell* or intracerebral or intracran* or parenchymal
	or intraventricular or infratentorial or supratentorial or basal gangli*) adj5
	(haemorrhage* or hemorrhage* or haematoma* or hematoma* or
	bleed*)).ti,ab,kf. OR exp myocardial infarction/ OR heart failure/ OR
	myocardial ischemia/ OR ((myocardi* adj2 (infarct* or isch?emi*)) or (heart
	adj2 (attack? or failure? or infarct* or isch?emi*)) or (cardiovascular adj2
	(stroke? or thrombosis*))).ti,ab,kf. OR coronary artery disease/ OR (coronary
	adj (heart or arter*) adj disease).ti,ab,kf. OR exp Cardiovascular Diseases/ OR
	((cardio* or cardiac or heart or vascular) adj3 (abnormal* or accident? or
	burden? or complication? or comorbid* or co-morbid* or disease? or
	dysfunction* or event? or manifest* or risk? or hospitali#ation?)).ti,ab,kf. OR
	Angina Pectoris/ OR angina, unstable/ or angina pectoris, variant/ Or (angina?
	adj2 (unstable or preinfarction or pre-infarction or variant?)).ti,ab,kf. OR
	angina, stable/ OR (angina? adj2 (pectoris or stable or stabilised)).ti,ab,kf. OR angina?.ti,kf. (19985) OR *Heart Disease/ OR exp Heart Failure/ OR ((heart or
	cardiac or myocardi*) adj2 (failure or decompensation)).ti,ab,kf. OR
	((renocardiac or reno-cardiac or cardiorenal or cardio-renal) adj1
	syndrome?).ti,ab,kf. OR ((cardiac adj (asthma or edema? or oedema?)) or
	paroxysmal dyspnea?).ti,ab,kf. OR *brain ischemia/ OR ischemic attack,
	transient/ OR (transient isch?emi* attack? or (transient adj2 (brain stem or
	brainstem or cerebral) adj2 isch?emi*) or TIA).ti,ab,kf. Or exp Percutaneous
	Coronary Intervention/ OR exp Myocardial Revascularization/ OR Cerebral
	Revascularization/ OR revasculari#ation?.ti,ab,kf,hw. OR (angioplast* or
	atherectom* or bypass* or CABG).ti,ab,kf,hw. OR peripheral arterial disease/
	OR (acute and ((limb or vascular) adj2 isch?emi*)).ti,ab,kf,hw. OR (peripheral
	arter* and (event? or thrombo*)).ti,ab,kf. OR *Peripheral Vascular Diseases/
	OR Peripheral Vascular Diseases/co, et, pa OR *Arterial Occlusive Diseases/ OR
	Arterial Occlusive Diseases/co, et, pa [Complications, Etiology, Pathology] OR
	thromboembolism/ OR Venous Thromboembolism/ OR ((thromboemboli* or
	thrombo-emboli*) adj3 event*).ti,ab,kf. OR ((vein? or venous) adj2 (thrombo*

or thrombus or emboli*)).ti,ab,kf. OR (VTE or DVT).ti,ab,kf. OR exp Pulmonary Embolism/ OR ((pulmonary or lung?) adj2 (thrombo* or thrombus or emboli*)).ti,ab,kf.

AND

Outcome measures

risk factors/ OR risk assessment/ OR (incidence or prevalence or prediction or prognosis).ti,kf,hw. OR mortality/ or death/ or "cause of death"/ or survival rate/ OR (survival or morbidit* or comorbid* or co-morbid*).ti,kf,hw. OR (death? or mortality or fatal*).ti,kf. OR ((death? or mortalit* or fatal*) adj3 (cause? or causal or compar* or ratio or rate? or register* or registries or statistic* or lupus or SLE)).ti,ab,kf. OR Life Tables/

AND

Study design

	epidemiologic studies/ or case-control studies/ or cross-sectional studies/ or cohort studies/ or follow-up studies/ or longitudinal studies/ or prospective studies/ or retrospective studies/ OR (registry or registries).ti,ab,kf,hw. OR ((epidemiologic or prospective or retrospective or cross-sectional or case control* or cohort or longitudinal) adj3 (study or studies)).ti,ab,kf. OR (case control* or coss-sectional or cohort? or follow-up or followup or longitudinal or prospective or retrospective or observational or population).ti. OR ((cohort? adj2 (analys* or compar* or data or study or studies or lupus or SLE)) or (population adj2 (based or data* or study or studies or register? or registry or registries or survey? or surveillance))).ab. OR (compar* and (study or risk?) and population).ti,ab,kf. OR ((chart? adj (audit? or review?)) or ((autopsy or hospital*) adj2 (report? or record?)) or death certificate?).ti,ab,kf. OR hospital*ation?.ti,kf,hw OR (systematic or structured or evidence or studies).ti. and ((review or overview or look or examination or update* or summary).ti. or review.pt.) OR (0266-4623 or 1366-5278 or 1530-440X or 2046- 4053).is. OR meta-analysis.pt. or (meta-analys* or meta analys* or metaanalys* or meta synth* or meta-synth* or metasynth*).ti,ab,kf,hw. OR ((systematic or meta) adj2 (analys* or review)).ti,kf. or ((systematic* or quantitativ* or methodologic*) adj5 (review* or overview*)).ti,ab,kf,sh. or (quantitativ\$ adj5 synthesis\$).ti,ab,kf,sh. OR (integrative research review* or research integration).ti,ab,kf. or (review.ti,kf,pt. and (trials as topic or studies as topic).hw.) OR review.pt. and ((medline or medlars or embase or pubmed or scisearch or psychinfo or psychilt or psyclit or cinahl or electronic database* or poling or pooled or mantel haenszel or peto or dersimonian or der simonian or fixed effect or ((hand adj2 search*) or (manual* adj2 search*))).tw,hw. or (retraction of publication or retracted publication).pt.) and (cohort? or observational or studies).mp.
Ovid Embase:	Population
1974 to 2020	systemic lupus erythematosus/ or (lupus adj3 (erythematosus or erythematous
Week 20	or erythematosis) adj3 (disseminatus or systemic)).ti,ab,kw. OR
	(libman sacks adj3 (disease or endocarditis or syndrom*)).ti,ab,kw. OR (lupus

or SLE).ti.

AND

Outcomes

cerebrovascular disease/ or exp basal ganglion hemorrhage/ or exp brain hematoma/ or exp brain hemorrhage/ or exp brain infarction/ or exp brain ischemia/ or exp carotid artery disease/ or cerebral artery disease/ or exp cerebrovascular accident/ or exp intracranial aneurysm/ or exp occlusive cerebrovascular disease/ or vertebrobasilar insufficiency/ or stroke/ or stroke patient/ or stroke unit/ OR (stroke or poststroke or cerebrovasc* or brain vasc* or cerebral vasc* or cva* or apoplex*).ti,kw. OR ((brain* or cerebr* or cerebell* or vertebrobasilar or hemispher* or intracran* or intracerebral or infratentorial or supratentorial or MCA or anterior circulation or posterior circulation or basal ganglia) adj5 (isch?emi* or infarct* or thrombo* or emboli* or occlus*)).ti,kw. OR ((brain* or cerebr* or cerebell* or intracerebral or intracran* or parenchymal or intraventricular or infratentorial or supratentorial or basal gangli*) adj5 (haemorrhage* or hemorrhage* or haematoma* or hematoma* or bleed*)).ti,kw. OR ((lupus or SLE) adj3 (stroke? or brain* or cerebr* or cerebell* or intracerebral or intracran* or apoplex*)).ti,ab,kw. OR *systemic lupus erythematosus/ or (lupus or SLE).ti. OR exp *heart infarction/ OR exp *ischemic heart disease/ OR exp *coronary artery disease/ OR ((myocardi* adj2 (infarct* or isch?emi*)) or (heart adj2 (attack? or failure? or infarct* or isch?emi*)) or (cardiovascular adj2 (stroke? or thrombosis*))).ti,kw. OR ((lupus or SLE) adj3 (heart or cardiac or cardiovascular or coronary or myocardi*)).ti,ab,kw. OR

exp *cardiovascular disease/ OR ((cardio* or cardiac or heart or vascular) adj3 (abnormal* or accident? or burden? or complication? or comorbid* or comorbid* or disease? or dysfunction* or event? or manifest* or risk? or hospitali#ation?)).ti,kw. OR (lupus or SLE).af. and (cardiovascular mortality/ or heart death/) OR angina pectoris/ OR angina pectoris/ or exp unstable angina pectoris/ or exp variant angina pectoris/ OR (angina? adj2 (unstable or preinfarction or pre-infarction or variant? or revasculari*)).ti,ab,kw. OR angina?.ti,kw. OR *Heart Disease/ OR exp *heart failure/ OR ((heart or cardiac or myocardi*) adj2 (failure or decompensation)).ti,kw. OR ((renocardiac or reno-cardiac or cardiorenal or cardio-renal) adj1 syndrome?).ti,ab,kw. OR ((cardiac adj (asthma or edema? or oedema?)) or paroxysmal dyspnea?).ti,ab,kw. exp *brain ischemia/ transient ischemic attack/ OR (transient isch?emi* attack? or (transient adj2 (brain stem or brainstem or cerebral) adj2 isch?emi*) or TIA).ti,ab,kw. OR percutaneous coronary intervention/ or transluminal coronary angioplasty/ OR heart muscle revascularization/ or *coronary artery surgery/ OR cerebral revascularization/ or cerebrovascular surgery/ OR revasculari#ation?.ti,kw. OR (angioplast* or atherectom* or bypass* or CABG).ti,ab,kw,hw. OR peripheral occlusive artery disease/ or exp artery occlusion/ or exp claudication/ OR (acute and ((limb or vascular) adj2 isch?emi*)).ti,ab,kw,hw. OR (peripheral arter* and (event? or thrombo*)).ti,ab,kw. OR peripheral vascular disease/ OR *thromboembolism/ OR venous thromboembolism/ or deep vein thrombosis/ or lower extremity deep vein thrombosis/ or lung embolism/ or upper extremity deep vein thrombosis/ OR lung embolism/ OR (thrombo* or thrombus or emboli* or DVT or VTE).ti. OR ((lupus or SLE) adj5 (thrombo* or thrombus or emboli* or DVT or

VTE)).ti.

AND

Outcome measures

risk factors/ OR risk assessment/ OR (incidence or prevalence or prediction or prognosis).ti,kw,hw. OR mortality/ or all cause mortality/ or hospital mortality/ or exp mortality rate/ or premature mortality/ or standardized mortality ratio/OR *death/ or "cause of death"/ or survival rate/OR (death? or mortality or fatal*).ti,kw. OR ((death? or mortalit* or fatal*) adj3 (cause? or causal or compar* or ratio or rate? or register* or registries or statistic* or lupus or SLE)).ti,ab,kw. OR life table/OR *survival/ or *morbidity/ or *comorbidity/

AND

Study design

prospective study/ or retrospective study/ or longitudinal study/ or cohort analysis/ or cross-sectional study/ or case control study/ or population based case control study/OR (registry or registries).ti,ab,kw,hw. OR ((epidemiologic or prospective or retrospective or cross-sectional or case control* or cohort or longitudinal) adj3 study).ti,ab,kw. OR (case control* or cross-sectional or cohort? or follow-up or followup or longitudinal or prospective or retrospective or observational or population).ti. OR ((cohort? adj2 (analys* or compar* or data or study or studies or lupus or SLE)) or (population adj2 (based or data* or study or studies or register? or registry or registries or survey? or surveillance))).ab,kw. OR ((compar* adj3 (study or risk?)) and population).ti,ab,kw. OR ((chart? adj (audit? or review?)) or ((autopsy or hospital*) adj2 (report? or record?)) or death certificate?).ti,ab,kw. OR hospitali#ation?.ti,kw,hw. OR systematic review/ or meta analysis/ or network meta-analysis/ OR ((systematic or structured or evidence or trials or studies) and (review or overview or look or examination or update* or summary)).ti. OR (0266-4623 or 1469-493X or 1366-5278 or 1530-440X or 2046-4053).is. OR (systematic review? or evidence report* or technology assessment?).jw. OR (meta-analys* or meta analys* or metaanalys* or meta synth* or meta-synth* or metasynth*).ti,ab,kw,hw. OR ((systematic or meta) adj2 (analys* or review)).ti,kw. or ((systematic* or quantitativ* or methodologic*) adj5 (review* or overview*)).ti,ab,kw,sh. or (quantitativ* adj5 synthes*).ti,ab,kw,hw. OR exp "clinical trial (topic)"/ and review.ti,kw,pt. OR (integrative research review* or research integration).ti,ab,kw. or scoping review?.ti,kw. or (evidence adj3 review*).ti,ab,kw. OR review.pt. and (medline or medlars or embase or pubmed or scisearch or psychinfo or psycinfo or psychlit or psyclit or cinahl or electronic database* or bibliographic database* or computeri#ed database* or online database* or pooling or pooled or mantel haenszel or peto or dersimonian or der simonian or fixed effect or ((hand adj2 search*) or (manual* adj2 search*))).ti,ab,kw,hw. OR review.pt. and ((evidence based adj (medicine or practice)) or (outcome? adj (assessment or research)) or treatment outcome).hw.

Limitations No animals, case reports, editorials, or letters; no children

Supplementary Table S2. 12-point scale developed for studying observational studies in SLE and
CVD risk

Domains	12-point scale	Score
The source of the study sample*	Community-based	2
	Clinic-based	1
	Undefined	0
Cohort type**	Inception	2
	Non-inception	1
Definition of SLE	Use of the ACR classification criteria for SLE[1, 2]	2
	Other validated criteria	1
	Other pre-defined but non-validated criteria	0
SLE exposure	10 years or more	2
·	Less than 10, more than 5 years	1
	Less than 5 years or not defined	0
Assessment of individual outcome		
CVD death	Validated criteria	2
	Non-validated but clearly defined criteria	1
	(e.g., death certificates)	
	Not mentioned	0
МІ	Validated criteria	2
	Non-validated but clearly defined criteria	1
	(e.g., death certificates)	
	Not mentioned	0
Stroke	Validated criteria	2
	Non-validated but clearly defined criteria	1
	(e.g. death certificates)	
	Not mentioned	0

*Community-based is defined as a study using data from a geographically defined area (e.g., north Italy, national registry). Clinic-based is defined as a study using data from a number of single clinics or hospital registries.

**An inception cohort is defined as a study that follows patients prospectively from the time of first SLE diagnosis, therefore only includes incident cases of SLE and explicitly states that prevalent cases were excluded.

ACR, American College of Rheumatology; CVD, cardiovascular disease; MI, myocardial infarction; SLE, systemic lupus erythematosus.

Supplementary Table S3. Newcastle-Ottawa Scale[3]

Col	hort studies
Sel	ection
1.	Representativeness of the exposed cohort a. Truly representative of the average (describe) in the community ** b. Somewhat representative of the average in the community * c. Selected group of users e.g., nurses, volunteers d. No description of the derivation of the cohort
2.	 Selection of the non-exposed cohort a. Drawn from the same community as the exposed cohort * b. Drawn from a different source c. No description of the derivation of the non-exposed cohort
3.	Ascertainment of exposure a. Secure record (e.g., surgical records) ★ b. Structured interview ★ c. Written self-report d. No description
4.	 Demonstration that outcome of interest was not present at start of study a. Yes ★ b. No
Со	mparability
	 Comparability of cohorts on the basis of the design or analysis a. Study controls for (select the most important factor) ★ b. Study controls for any additional factor ★ (this criterion could be modified to indicate specific control for a second important factor)
	ltcome
1.	Assessment of outcome a. Independent blind assessment ★ b. Record linkage ★ c. Self-report d. No description
2.	Was follow up long enough for outcomes to occur a. Yes (select an adequate follow-up period for outcome of interest) * b. No
3.	 Adequacy of follow up of cohorts a. Complete follow up – all subjects accounted for ★ b. Subjects lost to follow up unlikely to introduce bias – small number lost – >% (select an adequate %) follow up, or description provided of those lost) ★ c. Follow-up rate <% (select an adequate %) and no description of those lost d. No statement
Day	printed with permission from Dr. GA Wells

Reprinted with permission from Dr. GA Wells.

Supplementary Table S4. Excluded studies

Study	Reference	Reason for exclusion
Abramovich 2018	Incidence and variables associated with short and long-term mortality in patients with systemic lupus erythematosus and sepsis admitted in intensive care units. <i>Lupus</i> 2018;27(12):1936–43.	Wrong comparison
Abu-Shakra 1995	Mortality studies in systemic lupus erythematosus. Results from a single center. I. Causes of death. <i>J Rheumatol</i> 1995:22(7):1259–64.	Wrong outcome
Adwan 2020	In-hospital mortality in patients with systemic lupus erythematosus: a study from Jordan 2002–2017. <i>Rheumatol Int</i> 2020;40(5):711–7.	Wrong comparison
Afeltra 2005	Thrombosis in systemic lupus erythematosus: congenital and acquired risk factors. <i>Arthritis Care Res (Hoboken)</i> 2005;53(3):452–9.	No usable data
Aguero 2015	Prevalence of lower extremity peripheral arterial disease in individuals with chronic immune mediated inflammatory disorders. <i>Atheroscler</i> 2015;242(1):1–7.	Wrong population
Ahlehoff 2017	Cutaneous lupus erythematosus and the risk of deep venous thrombosis and pulmonary embolism: a Danish nationwide cohort study. <i>Lupus</i> 2017;26(13):1435–9.	Did not report on stroke or MI
Ahn 2018	Prevalence, risk factors, and impact on mortality of neuropsychiatric lupus: a prospective, single-center study. <i>Lupus</i> 2018;27(8):1338–47.	Wrong comparison
Alessandri 2017	Anti-mutated citrullinated vimentin antibodies in antiphospholipid syndrome: diagnostic value and relationship with clinical features. <i>Immunol Res</i> 2017;65(2):524–31.	Too few patients of interest
Alian 2019	Charlson Comorbidity Index in patients with systemic lupus erythematosus in Egypt and its relation with disease characteristics. <i>Indian J Rheumatol</i> 2019;14(4):277.	Wrong comparison
Alliu 2020	Outcomes of percutaneous coronary intervention (PCI) among patients with connective tissue disease: propensity match analysis. <i>Int J Cardiol</i> 2020;304:29–34.	Wrong population
Ando 2019	Acute myocardial infarction outcomes in systemic lupus erythematosus (from the nationwide inpatient sample). <i>Am J</i> <i>Cardiol</i> 2019;123(2):227–32.	Wrong population
Annangi 2017	Prevalence of pulmonary embolism among systemic lupus erythematosus discharges: a decade of analysis of the National Hospital Discharge Survey. <i>J Clin Rheumatol</i> 2017;23(4):200–6.	No usable data
Anver 2019	Changing trends in mortality in systemic lupus erythematosus? An analysis of SLE inpatient mortality at University Hospital Coventry and Warwickshire NHS Trust from 2007 to 2016. <i>Rheumatol Int</i> 2019;39(12):2069–75.	No usable data
Arkema 2015	Cohort profile: systemic lupus erythematosus in Sweden: the Swedish Lupus Linkage (SLINK) cohort. <i>BMJ Open</i> 2015;5(8):14.	Wrong outcome
Arkema 2016	What to expect when expecting with systemic lupus erythematosus (SLE): a population-based study of maternal and fetal outcomes in SLE and pre-SLE. <i>Arthritis Care Res (Hoboken)</i> 2016;68(7):988–94.	Wrong comparison
Atta 2018	Clinical and laboratory aspects of dyslipidemia in Brazilian women with systemic lupus erythematosus. <i>Clin Rheumatol</i>	Wrong comparison

	2018-27/6-1520 46	
Andrea Zachtarta	2018;37(6):1539–46.	Diduct
Avina-Zubieta	The risk of pulmonary embolism and deep venous thrombosis in	Did not
2015	systemic lupus erythematosus: a general population-based study.	report on
Daharada 2017	Semin Arthritis Rheum 2015;45(2):195–201.	stroke or MI
Babazade 2017	Systemic lupus erythematosus is associated with increased adverse	Wrong
	postoperative renal outcomes and mortality: a historical cohort	comparison
	study using administrative health data. Anesth Analg	
Barber 2012	2017;124(4):1118–26. Evaluation of clinical outcomes and renal vascular pathology	Wrong
Darber 2012	among patients with lupus. <i>Clin J Am Soc Nephrol</i> 7(5):757–64.	outcome
Barbhaiya	Racial/ethnic variation in stroke rates and risks among patients	Wrong
2019	with systemic lupus erythematosus. Semin Arthritis Rheum	comparison
2015	2019;48(5):840–6.	companson
Barnado 2018	Phenome-wide association study identifies marked increased in	Did not
	burden of comorbidities in African Americans with systemic lupus	report on
	erythematosus. Arthritis Res Ther 2018;20(1):69.	stroke or MI
Bartels 2014	Mortality and cardiovascular burden of systemic lupus	No usable
	erythematosus in a US population-based cohort. J Rheumatol	data
	2014;41(4):680–7.	
Belfeki 2018	Thrombophilia in systemic lupus erythematosus: a case-control	Non-English
	study. J Med Vasc 2018;43(6):347–53.	0
Ben-Zvi 2016	The impact of inflammatory rheumatic diseases on the	Wrong
	presentation, severity, and outcome of acute coronary syndrome.	population
	Clin Rheumatol 2016;35(1):233–7.	
Bertolaccini	Antibodies directed to protein S in patients with systemic lupus	No usable
2003	erythematosus: prevalence and clinical significance. Thromb	data
	Haemost 2003;90(4):636–41.	
Bertolaccini	Antiprothrombin antibodies detected in two different assay	Wrong
2005	systems. Prevalence and clinical significance in systemic lupus	outcome
	erythematosus. Thromb Haemost 2005;93(2):289–97.	
Bertolaccini	Factor XII autoantibodies as a novel marker for thrombosis and	Wrong
2007	adverse obstetric history in patients with systemic lupus	outcome
	erythematosus. Ann Rheum Dis 2007;66(4):533–6.	
Bessant 2004	Risk of coronary heart disease and stroke in a large British cohort	Wrong
	of patients with systemic lupus erythematosus. <i>Rheumatology</i>	comparison
Bloou 2016	2004;43(7):924–9.	14/10000
Bleau 2016	Risk of venous thromboembolic events in pregnant patients with	Wrong
	autoimmune diseases: a population-based study. <i>Clin Appl Thromb</i>	comparison
Bohensky 2014	<i>Hemost</i> 2016;22(3):285–91. Statin initiation and treatment non-adherence following a first	Wrong
Donensky 2014	acute myocardial infarction in patients with inflammatory	outcome
	rheumatic disease versus the general population. Arthritis Res Ther	outcome
	2014;16(5):443.	
Brouwer 2004	[The contribution of inherited and acquired thrombophilic defects,	Wrong
21041101 2004	alone or combined with antiphospholipid antibodies, to venous	comparison
	and arterial thromboembolism in patients with systemic lupus	companson
	erythematosus]. [French]. <i>Rev Med Interne</i> 2004;26(2):163–4.	
Bruce 2003	Risk factors for coronary heart disease in women with systemic	Wrong
	lupus erythematosus: the Toronto Risk Factor study. Arthritis	outcome
	Rheum 2003;48(11):3159–67.	
Bultink 2014	Elevated risk of clinical fractures and associated risk factors in	Wrong
_		0

	patients with systemic lupus erythematosus versus matched	outcomo
	controls: a population-based study in the United Kingdom.	outcome
Burgos 2000	Osteoporos Int 2014;25(4):1275–83. Peripheral vascular damage in systemic lupus erythematosus: data	Wrong
Burgos 2009	from LUMINA, a large multi-ethnic U.S. cohort (LXIX). <i>Lupus</i>	Wrong
		comparison
Busch 2018	2009;18(14):1303–8. Hospitalizations among Danish SLE patients: a prospective study	Wrong
Busch 2018	on incidence, causes of admission and risk factors in a population-	Wrong
	based cohort. <i>Lupus</i> 2018;27(1):165–71.	comparison
Butkiewicz	Associations between the incidence of antiphosphatidylserine and	No usable
2014	antiphosphatidylethanolamine antibodies and clinical	data
2014	manifestations of systemic lupus erythematosus. Pol Arch Med	uala
	Wewn 2014;124(11):573–8.	
Cadaval 2009	[Assessment of the risk of coronary heart disease in women with	Wrong
	systemic lupus erythematosus]. [Portuguese, English]. <i>Rev Bras</i>	comparison
	Rheumatol 2009;49(6):658–69.	comparison
Cassano 2007	Accrual of organ damage over time in Argentine patients with	Wrong
CassallU 2007	systemic lupus erythematosus: a multi-centre study. <i>Clin</i>	comparison
	Rheumatol 2007;26(12):2017–22.	companson
Cervera 2009	Morbidity and mortality in the antiphospholipid syndrome during a	Wrong
	5-year period: a multicentre prospective study of 1000 patients.	comparison
	Ann Rheum Dis 2009;68(9):1428–32.	comparison
Cervera 2015	Morbidity and mortality in the antiphospholipid syndrome during a	Wrong
	10-year period: a multicentre prospective study of 1000 patients.	population
	Ann Rheum Dis 2015;74(6):1011–8.	population
Chang 2018	High health care utilization preceding diagnosis of systemic lupus	Wrong
	erythematosus in youth. Arthritis Care Res 2018;70(9):1303–11.	comparison
Chen 2011	Predictors of long-term mortality in patients with and without	Wrong
	systemic lupus erythematosus on maintenance dialysis: a	comparison
	comparative study. J Rheumatol 2011;38(11):2390–4.	•
Chen 2014	Onset age affects mortality and renal outcome of female systemic	No usable
	lupus erythematosus patients: a nationwide population-based	data
	study in Taiwan. Rheumatology (Oxford) 2014;53(1):180-5.	
Chen 2019	Incidence and survival impact of pulmonary arterial hypertension	Wrong
	among patients with systemic lupus erythematosus: a nationwide	comparison
	cohort study. Arthritis Res Ther 2019;21(1):82.	
Chen 2019	Heart failure risk in systemic lupus erythematosus compared to	Wrong
	diabetes mellitus and general medicaid patients. Semin Arthritis	outcome
	Rheum 2019;49(3):389–95.	
Chuang 2015	Risk of peripheral arterial occlusive disease in patients with	Wrong
	systemic lupus erythematosus: a nationwide population-based	outcome
	cohort study. <i>Medicine (Baltimore)</i> 2015;94(46): e2121.	
Chung 2014	Systemic lupus erythematosus increases the risks of deep vein	Did not
	thrombosis and pulmonary embolism: a nationwide cohort study. J	report on
	Thromb Haemost 2014;12(4):452–8.	stroke or MI
Ciccaci 2014	A multilocus genetic study in a cohort of Italian SLE patients	Wrong
	confirms the association with STAT4 gene and describes a new	outcome
	association with HCP5 gene. <i>PLoS One</i> 2014;9(11):e111991.	
Crosslin 2009	The impact of race and ethnicity on disease severity in systemic	Wrong
	lupus erythematosus. <i>Ethn Dis</i> 2009;19(3):301–7. [Summary for	comparison
	patients in <i>Ethn Dis</i> 2009;19(3):365].	companison

Crozier 1990	Cardiac involvement in systemic lupus erythematosus detected by	Wrong
	echocardiography. Am J Cardiol 1990;65(16):1145–8.	outcome
Dave 2014	Atherosclerotic cardiovascular disease in hospitalized patients with	Wrong
	systemic sclerosis: higher mortality than patients with lupus and	outcome
	rheumatoid arthritis. Arthritis Care Res (Hoboken) 2014;66(2):323-	
	7.	
Davey 2010	The role of endothelial dysfunction in the pathogenesis of	Wrong
	neuropsychiatric systemic lupus erythematosus. Lupus	comparison
	2010;19(7):797–802.	
Demina 2005	[Causes of death in patients with rheumatic diseases in Moscow].	Non-English
	[Russian]. Ter Arkh 2005;77(4):77–82.	
Dhital 2020	All-cause hospitalizations and mortality in systemic lupus	No usable
	erythematosus in the US: results from a national inpatient	data
	database. Rheumatol Int. 2020;40(3):393-7.	
Di 2019	Framingham, ACC/AHA or QRISK3: which is the best in systemic	Wrong
	lupus erythematosus cardiovascular risk estimation? [published	comparison
	online ahead of print, 2019 Oct 28]. <i>Clin Exp Rheumatol</i> 2019.	
Dubois 1974	Duration and death in systemic lupus erythematosus. An analysis	Wrong
	of 249 cases. JAMA 1974;227(12):1399–1402.	comparison
Duman 2019	Cerebral venous sinus thrombosis as a rare complication of	Wrong
	systemic lupus erythematosus: subgroup analysis of the VENOST	comparison
	study. J Stroke Cerebrovasc Dis 2019;28(12):104372.	
Elfving 2014	Mortality and causes of death among incident cases of systemic	No usable
	lupus erythematosus in Finland 2000-2008. Lupus	data
F	2014;23(13):1430–4.	14/1000
Esdaile 2001	Traditional Framingham risk factors fail to fully account for	Wrong
	accelerated atherosclerosis in systemic lupus erythematosus.	comparison
Falasinnu 2017	Arthritis Rheum 2001;44(10):2331–7. Impact of sex on systemic lupus erythematosus-related causes of	Wrong
Falasilliu 2017	premature mortality in the United States. Journal of Women's	outcome
	Health. 2017 Nov 1;26(11):1214-21.	outcome
Falasinnu 2018	Do death certificates underestimate the burden of rare diseases?	Wrong
	The example of systemic lupus erythematosus mortality, Sweden,	outcome
	2001-2013. <i>Public Health Rep</i> 2018;133(4):481–8.	outcome
Fasano 2018	The incidence of cardiovascular events in Italian patients with	Wrong
	systemic lupus erythematosus is lower than in North European and	outcome
	American cohorts: implication of disease-associated and	outcome
	traditional risk factors as emerged by a 16-year retrospective	
	GIRRCS study: GIRRCS= Gruppo Italiano di Ricerca in Reumatologia	
	Clinica e Sperimentale. <i>Medicine</i> 2018;97(15).	
Fesmire 2010	Effects of autoimmune antibodies anti-lipoprotein lipase, anti-low	Wrong
	density lipoprotein, and anti-oxidized low density lipoprotein on	outcome
	lipid metabolism and atherosclerosis in systemic lupus	
	erythematosus. Rev Bras Rheumatol 2010;50(5):545–51.	
Fischer 2004	Effect of rheumatoid arthritis or systemic lupus erythematosus on	Wrong
	the risk of first-time acute myocardial infarction. Am J Cardiol	outcome
	2004;93(2):198–200.	_
Fischer 2007	[Significance of antiphospholipid syndrome and antiphospholipid	Non-English
-	antibodies in patients with systemic lupus erythematosus in	
	estimation of risk of subclinical atherosclerosis development].	
	[Polish]. Pol Arch Med Wewn 2007;117:13–7.	

Caldhana 2000		
Goldberg 2009	Risk factors for development of coronary artery disease in women	Wrong
	with systemic lupus erythematosus. <i>J Rheumatol</i>	outcome
Goldstein 1996	2009;36(11):2454–61.	Neurophie
Goldstein 1996	MHC studies of the primary antiphospholipid antibody syndrome	No usable
	and of antiphospholipid antibodies in systemic lupus	data
Crease 2000	erythematosus. <i>J Rheumatol</i> 1996;23(7):1173–9.	14/2022
Greco 2009	Association between depression and coronary artery calcification	Wrong
	in women with systemic lupus erythematosus. <i>Rheumatology</i>	outcome
Cue en et e in	2009;48(5):576–81.	14/2020
Greenstein	Burden of comorbidities in South Africans with systemic lupus	Wrong
2019	erythematosus. <i>Clin Rheumatol</i> 2019;38(8):2077–82.	comparison
Han 2017	Comorbid conditions are associated with emergency department	Non-
	visits, hospitalizations, and medical charges of patients with	comparative
Hanly 2018	systemic lupus erythematosus. <i>J Clin Rheumatol</i> 2017;23(1):19–25. Cerebrovascular events in systemic lupus erythematosus: results	Wrong
Halliy 2010	from an international inception cohort study. Arthritis Care Res	
	2018;70(10):1478–87.	comparison
Haque 2018	Progression of subclinical and clinical cardiovascular disease in a	Wrong
naque 2010	UK SLE cohort: the role of classic and SLE-related factors. <i>Lupus Sci</i>	comparison
	Med 2018;5(1):e000267.	companson
Hawro 2015	Intractable headaches, ischemic stroke and seizures are linked to	Wrong
114410 2015	the presence of anti-beta2GPI antibodies in patients with systemic	outcome
	lupus erythematosus. <i>PLoS One</i> 2015:10(3):e0119911	outcome
Hesselvig 2017	Cutaneous lupus erythematosus and systemic lupus	No usable
	erythematosus are associated with clinically significant	data
	cardiovascular risk: A Danish nationwide cohort study. <i>Lupus</i>	aata
	2017;26(1):48–53.	
Ingvarsson	Good survival rates in systemic lupus erythematosus in southern	Wrong
2019	Sweden, while the mortality rate remains increased compared	outcome
	with the population. <i>Lupus</i> 2019;28(12):1488–94.	
Jacobsen 1999	Mortality and causes of death of 513 Danish patients with systemic	No usable
	lupus erythematosus. Scandinavian J Rheumatol 1999;28(2):75-80.	data
Jimenez 2008	Double heterozygosity polymorphisms for platelet glycoproteins	Wrong
	Ia/IIa and IIb/IIIa increases arterial thrombosis and arteriosclerosis	comparison
	in patients with the antiphospholipid syndrome or with systemic	
	lupus erythematosus. Ann Rheum Dis 2008;67(6):835–40.	
Johannesdottir	Autoimmune skin and connective tissue diseases and risk of	Did not
2012	venous thromboembolism: a population-based case-control study.	report on
	J Thromb Haemost 2012;10(5):815–21.	stroke or MI
Jonsson 1989	Outcome of neuropsychiatric systemic lupus erythematosus within	No usable
	a defined Swedish population: increased morbidity but low	data
	mortality. <i>Rheumatology</i> 1989;41(11):1308–12.	
June 2013	Peripheral vascular disease in systemic lupus patients. J Clin	Wrong
	Rheumatol 2013;19(7):367–72.	outcome
Karp 2012	Longitudinal evolution of risk of coronary heart disease in systemic	Wrong
	lupus erythematosus. J Rheumatol 2012;39(5):968–73.	outcome
Katz 2019	Systemic lupus erythematosus and increased prevalence of	Wrong
	atherosclerotic cardiovascular disease in hospitalized patients.	outcome
	Mayo Clin Proc 2019;94(8):1436–43.	
Kaul 2013	Association of systemic lupus erythematosus with angiographically	Wrong
	defined coronary artery disease: a retrospective cohort study.	comparison

	Arthritis Care Res (Hoboken) 2013;65(2):266–73.	
Ke 2019	Systemic lupus erythematosus is associated with poor outcome	Wrong
	after acute myocardial infarction. <i>Nutr, Metab Cardiovasc Dis</i>	comparison
	2019;29(12):1400–7.	
Kedves 2020	Large-scale mortality gap between SLE and control population is	Wrong
	associated with increased infection-related mortality in lupus.	comparison
	Rheumatology. 2020;keaa188.	
Khamashta	Association of antibodies against phospholipids with heart valve	Wrong
1990	disease in systemic lupus erythematosus. <i>Lancet</i>	outcome
	1990;335(8705):1541–4.	
Kim 2017	Elevated levels of soluble CD40 ligand are associated with	Non-
	antiphospholipid antibodies in patients with systemic lupus	comparative
	erythematosus. <i>Clin Exp Rheumatol</i> 2017;35(5):823–30.	
Kishore 2019	Systemic lupus erythematosus is associated with a high risk of	Wrong
	venous thromboembolism in hospitalized patients leading to poor	comparison
	outcomes and a higher cost: results from nationwide inpatient	
	sample database 2003-2011. ACR Open Rheumatol 2019;1(3):194–	
1.0:2015	200.	14/2022
Lai 2015	Outcomes of coronary artery bypass grafting in patients with	Wrong
	inflammatory rheumatic diseases: An 11-year nationwide cohort	comparison
Lai 2016	study. <i>J Thorac Cardiovasc Surg</i> 2015;149(3):859–66.e2.	Wrong
Lai 2016	Outcomes of percutaneous coronary intervention in patients with	Wrong
	rheumatoid arthritis and systemic lupus erythematosus: an 11-	comparison
1-: 2020	year nationwide cohort study. <i>Ann Rheum Dis</i> 2016;75(7):1350–6.	
Lai 2020	Outcomes of acute cardiovascular events in rheumatoid arthritis	Wrong
	and systemic lupus erythematosus: a population-based study.	comparison
Leonard 2018	Rheumatology 2020;59(6):1355–63. Novel gene variants associated with cardiovascular disease in	No usable
Leonaru 2016	systemic lupus erythematosus and rheumatoid arthritis. Ann	data
	Rheum Dis 2018;77(7):1063–9.	uala
Lerang 2014	Mortality and years of potential life loss in systemic lupus	Did not
Lerang 2014	erythematosus: a population-based cohort study. <i>Lupus</i>	report on
	2014;23(14):1546–52.	stroke or MI
Lim 2019	Racial disparities in mortality associated with systemic lupus	No usable
	erythematosus—Fulton and DeKalb Counties, Georgia, 2002–2016.	data
	<i>MMWR</i> 2019;68(18):419.	uuuu
Lim 2019	Pulmonary arterial hypertension in a multi-ethnic Asian	Wrong
	population: characteristics, survival and mortality predictors from	population
	a 14-year follow-up study. <i>Respirology</i> 2019;24(2):162–70.	202010101
Lin 2014	Adverse outcomes after major surgery in patients with systemic	Wrong
	lupus erythematosus: a nationwide population-based study. Ann	comparison
	Rheum Dis 2014;73(9):1646–51.	
Lood 2017	Decreased platelet size is associated with platelet activation and	Wrong
	anti-phospholipid syndrome in systemic lupus erythematosus.	outcome
	Rheumatology 2017;56(3):408–16.	-
López 2017	Serum levels of anti-PON1 and anti-HDL antibodies as potential	No usable
2017		data
2017	biomarkers of premature atherosclerosis in systemic lupus	data
-	biomarkers of premature atherosclerosis in systemic lupus erythematosus. <i>Thromb Haemost</i> 2017;117(11):2194–206.	
Lu 2015	biomarkers of premature atherosclerosis in systemic lupus	data Wrong outcome

Lundstrom	HLA-DRB1*04/*13 alleles are associated with vascular disease and	Wrong
2013	antiphospholipid antibodies in systemic lupus erythematosus. Ann Rheum Dis 2013;72(6):1018–25.	outcome
Magder 2012	Incidence of and risk factors for adverse cardiovascular events	Wrong
	among patients with systemic lupus erythematosus. <i>Am J</i> <i>Epidemiol</i> 2012;176(8):708–19.	comparison
Mageau 2019	The burden of chronic kidney disease in systemic lupus	Wrong
	erythematosus: a nationwide epidemiologic study. <i>Autoimmun Rev</i> 2019;18(7):733–7.	comparison
Maksimowicz-	Poor 1-year outcomes after percutaneous coronary interventions	Wrong
McKinnon	in systemic lupus erythematosus: report from the National Heart,	comparison
2008	Lung, and Blood Institute Dynamic Registry. Circulation	
Martinez 2020	2008;(3):201–8. Percutaneous coronary intervention outcomes in patients with	Mrong
Martinez 2020	rheumatoid arthritis, systemic lupus erythematosus and systemic	Wrong population
	sclerosis. <i>Rheumatology</i> 2020;kez639.	population
Mehta 2008	Platelet C4d is associated with acute ischemic stroke and stroke	Wrong
	severity. Stroke 2008;39(12):3236–41.	Wrong population
Mok 2010	Venous thromboembolism in southern Chinese patients with	Did not
WIOK 2010	systemic lupus erythematosus. <i>Clin Rheumatol</i> 2010;29(6):599–	report on
	604.	stroke or MI
Mok 2011	Epidemiology and survival of systemic lupus erythematosus in	Review
	Hong Kong Chinese. <i>Lupus</i> 2011;20(7):767–71.	article
Mok 2011	Life expectancy, standardized mortality ratios, and causes of death	No usable
	in six rheumatic diseases in Hong Kong, China. Arthritis Rheum	data
	2011;63(5):1182–9.	
Molad 2000	Renal outcome and vascular morbidity in systemic lupus	Wrong
	erythematosus (SLE): Lack of association with the angiotensin-	comparison
	converting enzyme gene polymorphism. Semin Arthritis Rheum	
	2000;30(2):132–7.	
Morgan 1993	Clinical analysis of 125 patients with the lupus anticoagulant. Aust	Wrong
-	N Z J Med 1993;23(2):151–6.	comparison
Moss 2002	Outcome of a cohort of 300 patients with systemic lupus	No usable
	erythematosus attending a dedicated clinic for over two decades.	data
	Ann Rheum Dis 2002;61(5):409–13.	
Mu 2018	Mortality and prognostic factors in Chinese patients with systemic	No usable
	lupus erythematosus. Lupus 2018;27(10):1742–52.	data
Munoz-	Prevalence and clinical significance of antiprothrombin antibodies	Wrong
Rodriguez 2000	in patients with systemic lupus erythematosus or with primary	comparison
	antiphospholipid syndrome. <i>Haematologica</i> 2000;85(6):632–7.	
Munoz-	Clinical significance of acquired activated protein C resistance in	Wrong
Rodriguez 2002	patients with systemic lupus erythematosus. Lupus	comparison
	2002;11(11):730–5.	
Nakamura	Autoantibody to CD40 ligand in systemic lupus erythematosus:	Wrong
2006	Association with thrombocytopenia but not thromboembolism.	comparison
	Rheumatology (Oxford) 2006;45(2):150–6.	
Nasonov 1992	The prediction of thrombosis development in patients with	Non-English
	systemic lupus erythematosus: the role of antibodies to	
	cardiolipin. [Russian]. Ter Arkh 1992;64(5):25–30.	
Nikpour 2009	Myocardial perfusion imaging in assessing risk of coronary events	Wrong
	in patients with systemic lupus erythematosus. J Rheumatol	comparison

	2009;36(2):288–94.	
Norby 2017	Outcome in biopsy-proven Lupus nephritis: Evaluation of biopsies	Wrong
/ -	from the Norwegian Kidney Biopsy Registry. <i>Lupus</i>	outcome
	2017;26(8):881–5.	
Oh 2003	Acquired activated protein C resistance, high tissue factor	Wrong
	expression, and hyper-homocysteinemia in systemic lupus	outcome
	erythematosus. Am J Hematol 2003;72(2):103–8.	
Ostanek 2006	Antiphospholipid syndrome and antiphospholipid antibodies as a	Non-English
	risk factors of ischaemic heart disease and myocardial infarction in	
	patients with systemic lupus erythematosus. [Polish]. Pol Arch Med	
	Wewn 2006;115(5):407–13.	
Panafidina	The significance of cardiovascular risk factors and C-reactive	Non-English
2006	protein to the development of atherosclerosis in women with	
	systemic lupus erythematosus. [Russian]. Klin Med (Mosk)	
	2006;84(10):49–54.	
Perez-Villa	Severe valvular regurgitation and antiphospholipid antibodies in	Wrong
2005	systemic lupus erythematosus: A prospective, long-term, followup	comparison
	study. Arthritis Care Res (Hoboken) 2005;53(3):460–7.	
Petri 1987	The frequency of lupus anticoagulant in systemic lupus	Wrong
	erythematosus. A study of sixty consecutive patients by activated	outcome
	partial thromboplastin time, Russell viper venom time, and	
	anticardiolipin antibody level. Ann Intern Med 1987;106(4):524–	
	31.	
Petri 2019	Development of a systemic lupus erythematosus cardiovascular	No usable
	risk equation. Lupus Sci Med 2019;6(1):e000346.	data
Plasín-	The H1 haplotype of the endothelial protein C receptor protects	No usable
Rodríguez 2018	against arterial thrombosis in patients with antiphospholipid	data
	syndrome. <i>Thromb Res</i> 2018;169:128–34.	
Ramagopalan	Risk of venous thromboembolism in people admitted to hospital	Did not
2011	with selected immune-mediated diseases: record-linkage study.	report on
D.:: 1 2020	<i>BMC Med</i> 2011;9:1.	stroke or MI
Reid 2020	High genetic risk score is associated with early disease onset,	No usable
	damage accrual and decreased survival in systemic lupus	data
Reshetniak	erythematosus. <i>Ann Rheum Dis</i> 2020;79(3):363–9. [Subclinical and clinical manifestations of atherosclerosis in	Non English
2008	antiphospholipid syndrome]. [Russian]. <i>Ter Arkh</i> 2008;80(10):60–7.	Non-English
Reshetniak	[Plasminogen activator inhibitor type 1 gene polymorphism and	Non-English
2013	thromboses in patients with antiphospholipid syndrome].	NOII-EIIglisii
2013	[Russian]. <i>Ter Arkh</i> 2013;85(1):76–84.	
Rodríguez-	Clinical and subclinical cardiovascular disease in female SLE	No usable
Carrio 2019	patients: interplay between body mass index and bone mineral	data
carrio 2015	density. <i>Nutr, Metab Cardiovasc Dis</i> 2019;29(2):135–43.	uutu
Roldan 1996	An echocardiographic study of valvular heart disease associated	Wrong
	with systemic lupus erythematosus. <i>New Engl J Med</i>	comparison
	1996;335(19):1424–30.	50
Roldan 2013	Libman-sacks endocarditis and embolic cerebrovascular disease.	Wrong
	JACC: Cardiovasc Imaging 2013;6(9):973–83.	comparison
Roldan 2015	Lamble excrescences: association with cerebrovascular disease and	Wrong
	pathogenesis. <i>Cerebrovasc Dis</i> 2015;40(1):18–27.	outcome
Rossides 2017	Mortality and functionality after stroke in patients with systemic	Wrong
	lupus erythematosus. J Rheumatol 2017;44(11):1590–6.	comparison
		30111011

Sairam 2002	Analysis of risk factors and comorbid disasses in the dayalanment	Mrong
Sairam 2003	Analysis of risk factors and comorbid diseases in the development	Wrong
	of thrombosis in patients with anticardiolipin antibodies. <i>Clin</i>	comparison
Sanmarco 1997	Rheumatol 2003;22(1):24–9.	
Sanmarco 1997	Prevalence and clinical significance of IgG isotype anti-beta 2-	Wrong
	glycoprotein I antibodies in antiphospholipid syndrome: a	outcome
	comparative study with anticardiolipin antibodies. <i>J Lab Clin Med</i>	
Caugh: 2005	1997;129(5):499–506.	New
Sarabi 2005	Incidence rates of arterial and venous thrombosis after diagnosis	Non-
	of systemic lupus erythematosus. <i>Arthritis Care Res (Hoboken)</i> 2005;53(4):609–12.	comparative
Sartori Vieira	Mortality profile related to the spectrum of systemic connective	Letter
2019	tissue diseases: a retrospective, population-based, case-control	
	study. Lupus 2019;28(12):1498–500.	
Scalzi 2010	Racial disparities in age at time of cardiovascular events and	No usable
	cardiovascular-related death in patients with systemic lupus	data
	erythematosus. Arthritis Rheum 2010;62(9):2767–75.	
Shafi 2007	Risk of vascular access thrombosis in patients with systemic lupus	Too few
	erythematosus on hemodialysis. J Vasc Access 2007;8(2):103–8.	patients of
		interest
Shah 2009	Poor outcomes after acute myocardial infarction in systemic lupus	Wrong
	erythematosus. J Rheumatol 2009;36(3):570–5.	comparison
Shang 2012	SLICC/ACR Damage Index independently associated with left	Wrong
	ventricular diastolic dysfunction in patients with systemic lupus	outcome
	erythematosus. Lupus 2012;21(10):1057-62.	
Shi 2017	Clinical characteristics and laboratory findings of 252 Chinese	Wrong
	patients with anti-phospholipid syndrome: comparison with Euro-	comparison
	Phospholipid cohort. Clin Rheumatol 2017;36(3):599–608.	
Singh 2016	Risk of cerebrovascular accidents and ischemic heart disease in	Wrong
	cutaneous lupus erythematosus: a population-based cohort study.	population
	Arthritis Care Res (Hoboken) 2016;68(11):1664–70.	
Smilowitz 2018	Systemic lupus erythematosus and the risk of perioperative major	Wrong
	adverse cardiovascular events. J Thromb Thrombolysis	comparison
	2018;45(1):13-7.	
Soh 2015	Association between pregnancy outcomes and death from	Wrong
	cardiovascular causes in parous women with systemic lupus	comparison
	erythematosus: a study using Swedish population registries.	
	Arthritis Rheumatol 2015;67(9):2376–82.	
Soh 2015	Brief report: association between pregnancy outcomes and death	Wrong
	from cardiovascular causes in parous women with systemic lupus	comparison
	erythematosus: a study using Swedish population registries.	
	Arthritis Rheumatol 2015;67(9):2376–82.	
Soltesz 2003	Evaluation of clinical and laboratory features of antiphospholipid	Wrong
	syndrome: a retrospective study of 637 patients. Lupus	comparison
	2003;12(4):302–7.	
Stahl-	Incidence studies of systemic lupus erythematosus in Southern	No usable
Hallengren	Sweden: increasing age, decreasing frequency of renal	data
2000	manifestations and good prognosis. <i>J Rheumatol</i> 2000;27(3):685–	
	91.	
Sun 2019	Association of lupus nephritis with coronary artery disease by	Wrong
	ISN/RPS classification: results from a large real-world lupus	comparison
	population. ACR Open Rheumatol 2019;1(4):244–50.	

<u>Current and and and and and and and and and and</u>	A STATA risk allele is accepted with independence and hypersonaler	14/2020
Svenungsson	A STAT4 risk allele is associated with ischaemic cerebrovascular	Wrong
2010	events and anti-phospholipid antibodies in systemic lupus	outcome
-	erythematosus. Ann Rheum Dis 2010;69(5):834–40.	
Svenungsson	Decreased levels of autoantibodies against apolipoprotein B-100	Wrong
2015	antigens are associated with cardiovascular disease in systemic	outcome
	lupus erythematosus. <i>Clin Exp Immunol</i> 2015;181(3):417–26.	
Tassies 2000	The 4G/5G polymorphism of the type 1 plasminogen activator	Wrong
	inhibitor gene and thrombosis in patients with antiphospholipid	outcome
	syndrome. <i>Arthritis Rheum</i> 2000;43(10):2349–58.	
Tektonidou	Brief report: trends in hospitalizations due to acute coronary	No usable
2016	syndromes and stroke in patients with systemic lupus	data
	erythematosus, 1996 to 2012. Arthritis Rheumatol	
	2016;68(11):2680–5.	
Thorburn 2003	Hospitalizations for coronary artery disease among patients with	Wrong
	systemic lupus erythematosus. Arthritis Rheum 2003;48(9):2519–	comparison
	23.	
Tkachenko	Profiling of non-criteria antiphospholipid antibodies in patients	Wrong
2020	with SLE: differentiation of thrombotic SLE patients and risk of	comparison
	recurrence of thrombosis. <i>Lupus</i> 2020;29(5):490–8.	
Toloza 2004	Systemic lupus erythematosus in a multiethnic US cohort	Non-
	(LUMINA). XXIII. Baseline predictors of vascular events. Arthritis	comparative
	Rheum 2004;50(12):3947–57.	
Tselios 2017	Evolution of risk factors for atherosclerotic cardiovascular events	No usable
	in systemic lupus erythematosus: a longterm prospective study. J	data
	Rheumatol 2017;44(12):1841–9.	
Tselios 2019	All-cause, cause-specific and age-specific standardised mortality	Wrong
	ratios of patients with systemic lupus erythematosus in Ontario,	outcome
	Canada over 43 years (1971–2013). Ann Rheum Dis	
	2019;78(6):802–6.	
Tuta 2006	[Clinical outcome of lupus nephritis in Constanta County].	Non-English
- 1 00/0	[Romanian]. <i>Rev Med Chir Soc Med Nat Iasi</i> 2006;110(2):299–304.	
Tyden 2013	Increased serum levels of S100A8/A9 and S100A12 are associated	Wrong
	with cardiovascular disease in patients with inactive systemic lupus	outcome
	erythematosus. <i>Rheumatology (Oxford)</i> 2013;52(11):2048–55.	
Tziomalos 2017	Arterial stiffness and peripheral arterial disease in patients with	Wrong
	systemic lupus erythematosus. <i>Rheumatol Int</i> 2017;37(2):293–8.	outcome
Ungprasert	Epidemiology of mixed connective tissue disease, 1985-2014: a	Wrong
2016	population-based study. Arthritis Care Res (Hoboken)	outcome
11	2016;68(12):1843–8.	14/2020
Unlu 2019	The impact of systemic lupus erythematosus on the clinical	Wrong
	phenotype of antiphospholipid antibody–positive patients: results	comparison
	from the AntiPhospholipid Syndrome Alliance for Clinical Trials and	
	InternatiOnal Clinical Database and Repository. <i>Arthritis Care Res</i>	
Urow:+- 2000	2019;71(1):134–41.	Non
Urowitz 2008	Accumulation of coronary artery disease risk factors over three	Non-
	years: data from an international inception cohort. Arthritis Care	comparative
	Res (Hoboken) 2008;59(2):176–80.	14/20 2 2
Van Doornum	Increased 30-day and 1-year mortality rates and lower coronary	Wrong
2015	revascularisation rates following acute myocardial infarction in	comparison
	patients with autoimmune rheumatic disease. Arthritis Res Ther	
	2015;17:38.	

Vaya 2008	Thrombotic events in systemic lupus erythematosus. Its	Wrong
	association with acquired and inherited thrombophilic defects. Clin	outcome
	Hemorheol Microcirc 2008;40(2):79–87.	
Vikerfors 2013	Clinical manifestations and anti-phospholipid antibodies in 712	Wrong
	patients with systemic lupus erythematosus: evaluation of two	outcome
	diagnostic assays. Rheumatology 2013;52(3):501–9.	
Voss 2013	Survival in systemic lupus erythematosus, 1995-2010. A	No usable
	prospective study in a Danish community. Lupus	data
	2013;22(11):1185–91.	
Ward 2000	Cardiovascular and cerebrovascular morbidity and mortality	Wrong
	among women with end-stage renal disease attributable to lupus	comparison
	nephritis. <i>Am J Kidney Dis</i> 2000;36(3):516–25.	
Ward 2004	Outcomes of hospitalizations for myocardial infarctions and	Wrong
	cerebrovascular accidents in patients with systemic lupus	comparison
	erythematosus. Arthritis Rheum 2004;50(10):3170–6.	
Watad 2017	Association between ischemic heart disease and systemic lupus	Wrong
	erythematosus-a large case-control study. Immunol Res	outcome
	2017;65(2):459–63.	
Watad 2017	The association between systemic lupus erythematosus and	Wrong
	valvular heart disease: an extensive data analysis. Eur J Clin Invest	outcome
	2017;47(5):366–71.	200000
Weng 2011	A retrospective study of pulmonary infarction in patients with	Non-
10016 2011	systemic lupus erythematosus from southern Taiwan. Lupus	comparative
	2011;20(8):876–85.	
Wu 2014	Major adverse cardiovascular events and mortality in systemic	Wrong
WU 2014	lupus erythematosus patients after successful delivery: a	comparison
	population-based study. Am J Med Sci 2014;347(1):42–9.	companison
Wu 2019	Causes of death in hospitalized patients with systemic lupus	No usable
WU 2015	erythematosus: a 10-year multicenter nationwide Chinese cohort.	data
	Clin Rheumatol 2019;38(1):107–15.	uutu
Yang 2012	Prevalence and correlation of conventional and lupus-specific risk	Wrong
	factors for cardiovascular disease in Chinese systemic lupus	outcome
	erythematosus patients. J Eur Acad Dermatol Venereol	outcome
	2012;26(1):95–101.	
Yap 2012	Survival analysis and causes of mortality in patients with lupus	Did not
1ah 2012		
	nephritis. Nephrol Dial Transplant 2012;27(8):3248–54.	report on
Varmaar 2001	Drognongy outcomes in women with systemic lysts	stroke or MI
Yasmeen 2001	Pregnancy outcomes in women with systemic lupus	Wrong
Varda	erythematosus. J Matern Fetal Med 2001;10(2):91–6.	comparison
Yazdanyar	Short-term perioperative all-cause mortality and cardiovascular	Wrong
2013	events in women with systemic lupus erythematosus. Arthritis	comparison
	<i>Care Res (Hoboken)</i> 2013;65(6):986–91.	
Yip 2009	Disease chronicity and activity predict subclinical left ventricular	Wrong
	systolic dysfunction in patients with systemic lupus erythematosus.	outcome
	Heart 2009;95(12):980–7.	
You 2020	Characteristics and risk factors of pulmonary embolism in patients	Wrong
	with systemic lupus erythematosus: a case control study.	comparison
	[published online ahead of print, 2020 Jan 20]. Clin Exp Rheumatol	
	2020.	
	2020.	
Yusuf 2014	Risk of venous thromboembolism among hospitalizations of adults	Did not

	2244 22(2) 225 42	
	2014;38(3):306–13.	stroke or MI
Yusuf 2015	Risk of venous thromboembolism occurrence among adults with	Did not
	selected autoimmune diseases: a study among a U.S. cohort of	report on
	commercial insurance enrollees. Thromb Res 2015;135(1):50-7.	stroke or MI
Zaccagni 2004	Soluble adhesion molecule levels, neuropsychiatric lupus and	Wrong
	lupus-related damage. Front Biosci 2004;9:1654–9.	comparison
Zhao 2017	Clinical characteristics and survival of pulmonary arterial	Non-
	hypertension associated with three major connective tissue	comparative
	diseases: a cohort study in China. Int J Cardiol 2017;236:432–7.	
Zirkzee 2014	Mortality in neuropsychiatric systemic lupus erythematosus	Wrong
	(NPSLE). Lupus 2014;23(1):31–8.	outcome
Zöller 2012	Risk of subsequent coronary heart disease in patients hospitalized	Wrong
	for immune-mediated diseases: a nationwide follow-up study from	outcome
	Sweden. <i>PloS One</i> 2012;7(3):e33442.	
Zöller 2012b	Risk of pulmonary embolism in patients with autoimmune	Did not
	disorders: a nationwide follow-up study from Sweden. Lancet	report on
	2012;379(9812):244–9.	stroke or MI
Zuily 2013	Superficial vein thrombosis, thrombin generation and activated	Wrong
	protein C resistance as predictors of thromboembolic events in	comparison
	lupus and antiphospholipid patients. A prospective cohort study.	
	Thromb Res 2013;132(1):e1–7.	

MI, myocardial infarction.

Supplementary Table S5. Quality assessment of the studies included in the meta-analysis that report the risk of stroke and MI in adult patients with SLE compared with the general population or healthy controls

Author/year		NOS		12-point scale							
	Selection (5 stars)	Comparability (2 stars)	Outcome (3 stars)	1. Source of the study sample	2. Cohort type (inception cohort)	3. Definition of SLE	4. Length of SLE exposure (years)	5. Assessment of outcome	6. Adjustment for confounders	Total 12-point SLE-specific scale	Overall estimate risk of bias
Arkema 2017[4]	4	2	3	Community	No	ICD-10	≥5 <10	Clearly defined	≥5 risk factors	8	Low risk
Avina-Zubieta 2017[5]	4	2	3	Community	No	ICD-9	≥5 <10	Clearly defined	<5 risk factors	8	Low risk
Barnado 2018[6]	5	1	3	Community	No	ICD-10	≥5 <10	Clearly defined	≥5 risk factors	11	Low risk
Bengtsson 2012[7]	3	2	3	Community	No	≥4 ACR	≥5 <10	Validated criteria	<5 risk factors	10	Low risk
Bernatsky 2006a[8]	3	2	3	Clinic based	Yes	≥4 ACR	≥10	Validated criteria	<5 risk factors	9	Low risk
Bernatsky 2006b[9]	3	2	3	Clinic based	No	≥4 ACR	≥10	Clearly defined	<5 risk factors	7	Moderate
Bjornadal 2004[10]	3	2	3	Clinic based	No	ICD-9	≥10	Clearly defined	<5 risk factors	7	Moderate
Chang 2013[11]	5	1	3	Community	No	ICD-9	≥5 <10	Validated criteria	<5 risk factors	9	Low risk
Chiu 2012[12]	4	2	3	Community	No	ICD-9	≥5 <10	Clearly defined	None	10	Low risk
Cook 2018[13]	1	0	2	Community	No	Self-report	NR	Self-report	≥5 risk factors	6	High risk
Dregan 2017[14]	2	2	2	Community	No	Clinician	<5	Not clear	≥5 risk factors	5	High risk
Faurschou 2011[15]	3	2	3	Clinic based	No	≥4 ACR	≥10	Validated criteria	<5 risk factors	8	Low risk
Hak 2009[16]	5	2	3	Community	Yes	≥4 ACR	≥10	Validated criteria	≥5 risk factors	12	Low risk
Hermansen 2017[17]	5	2	3	Community	Yes	ICD-10	≥10	Validated criteria	<5 risk factors	10	Low risk
Kim 2017[18]	3	2	3	Community	No	Unclear	≥10	Clearly defined	≥5 risk factors	8	Low risk
Krishnan 2005[19]	3	2	2	Community	No	ICD-9	NR	Clearly defined	<5 risk factors	5	High risk
Lim 2018[20]	4	2	3	Community	No	ICD-10	≥5 <10	Clearly defined	≥5 risk factors	11	Low risk

Lin 2014[21]	3	2	3	Community	Yes	ICD-9	≥5 <10	Validated criteria	<5 risk factors	10	Low risk
Liou 2014[22]	5	2	3	Community	No	≥4 ACR	≥10	Validated criteria	≥5 risk factors	11	Low risk
Manzi 1997[23]	4	2	3	Clinic based	No	≥4 ACR	≥10	Clearly defined	≥5 risk factors	9	Low risk
Mok 2009[24]	1	0	3	Clinic based	No	≥4 ACR	≥5 <10	Clearly defined	None	5	High risk
Ramagopalan 2013[25]	3	2	3	Community	No	ICD-9	≥10	Validated criteria	≥5 risk factors	9	Low risk
Rees 2016[26]	5	2	3	Community	Yes	Read codes	≥10	Clearly defined	≥5 risk factors	8	Low risk
Wang 2012[27]	5	2	3	Community	Yes	ICD-9	≥10	Validated criteria	<5 risk factors	11	Low risk
Ward 1999[28]	2	2	2	Community	No	ICD-9	<5	Clearly defined	≥5 risk factors	7	High risk
Zoller 2012[29]	5	2	3	Community	No	ICD-7-10	≥10	Clearly defined	≥5 risk factors	8	Low risk

ACR, American College of Rheumatology; ICD, International Classification of Diseases; MI, myocardial infarction; NOS, Newcastle-Ottawa Scale; NR, not reported; SLE, systemic lupus erythematosus.

Supplementary Table S6. Types of stroke and MI and definition reported by each study

Author/year	Ischaemic stroke	Intracerebral	Subarachnoid	Haemorrhagic	Unspecified	Composite stroke	MI
		haemorrhage	haemorrhage	stroke	stroke		
Arkema 2017[4]	ICD-8 432-434; ICD-	ICD-8 431; ICD-	ICD-8 430; ICD-9		ICD-8 436;	ICD-8 430-434,	
	9 433-434; ICD-10	9 431; ICD-10	430; ICD-10 I60		ICD-9 436;	436; ICD-9 430-	
	163	161			ICD-10 I64	431, 433-434,	
						436; ICD-10, I60-	
						161, 163-164	
Avina-Zubieta 2017[5]	ICD-9 433-434; ICD-						ICD-9 410; ICD-10 I21
	10 163-166						
Barnado 2018[6]							121, 122, 123, 124, 125, 151
Bengtsson 2012[7]						ICD-10 I61-6,	ICD-10 I21.0-9, I22.0-1, I22.8-9
						161.8-9, 162.9,	
						163.0-6, 163.8-9,	
						164	
Bernatsky 2006a[8]						ICD-9 430-459	
Bernatsky 2006b[9]	ICD-9 433-434		ICD-9 430			ICD-9 430-438	
Bjornadal 2004[10]						ICD-7 330-334;	
						ICD-8 430-438;	
						ICD-9 430-438	
Chang 2013[11]			ICD-9 430				
Chiu 2012[12]	Claims data						
Cook 2018[13]						Reported as	Reported as MI Biobank database based on
						stroke/ischaemic	ICD-10 codes
						stroke. Biobank	
						database based	
						on ICD-10 codes	
Dregan 2017[14]						ICD-10 and self-	
						report	
Faurschou 2011[15]							ICD-8 410; ICD-10 I21
Hak 2009[16]						National Survey of	WHO criteria plus in hospital measurements
						Stroke	or autopsy evidence for both
Hermansen 2017[17]						ICD-8 431-434;	ICD-8 410; ICD-10 I21
						ICD-10 I61, I62.9,	
						163-164	
Kim 2017[18]							NR
Krishnan 2005[19]	ICD-9 434	ICD-9 431	ICD-9 430		ICD-9 436	ICD-9 430-431,	
						434, 436	
Lim 2018[20]						Stroke ICD-10 I63,	ICD-10 (I21–22)

					164	
Lin 2014[21]						ICD-9 410
Liou 2014[22]	ICD-9 unspecified				ICD-9 430-438	
Manzi 1997[23]						Cardiovascular Health Study. Documentation required hospital data
Mok 2009[24]	Physician diagnosis	Physician			Physician	
		diagnosis			diagnosis	
Ramagopalan 2013[25]			ICD-9 430; ICD- 10 I60			
Rees 2016[26]					Read code available on request	
Wang 2012[27]	ICD-9 unspecified	ICD-9 unspecified	ICD-9 unspecified	ICD-9 unspecified	ICD-9 430-438	
Ward 1999[28]					ICD-9 431 or 434	ICD-9 410
Zoller 2012[29]	ICD-9 433-435, 437;	ICD-9 431-432;				
	ICD-10 I63, I65-I67	ICD-10 I61-I62				

ICD, International Classification of Diseases; MI, myocardial infarction; NR, not reported; WHO, World Health Organization.

Supplementary Table S7. Risk ratios and 95% confidence intervals for stroke and MI in adult patients with SLE compared with the general population or healthy controls: main meta-analysis and sensitivity analyses

Composite stroke								
Authors	Main analysis	Least adjusted	Published during or after 2014	Published before 2014	Only studies with low risk of bias	Fatal/non-fatal	Non-fatal	Excluding cross- sectional studies
Arkema 2017[4]	2.1 (1.7 to 2.6)	2.3 (1.8 to 2.8)	2.1 (1.7 to 2.6)	-	2.1 (1.7 to 2.6)	2.1 (1.7 to 2.6)	-	2.1 (1.7 to 2.6)
Bengtsson 2012[7]	-	-	-	1.6 (0.69 to 3.14)	-	-	-	-
Bernatsky 2006a[8]	1.1 (0.7 to 1.7)	1.1 (0.7 to 1.7)	-	1.1 (0.7 to 1.7)	1.1 (0.7 to 1.7)	-	-	1.1 (0.7 to 1.7)
Bernatsky 2006b[9]	-	-	-	-	-	-	-	-
Bjornadal 2004[10]	2.06 (1.74 to 2.43)	2.06 (1.74 to 2.43)	-	2.06 (1.74 to 2.43)	-	-	-	2.06 (1.74 to 2.43)
Cook 2018[13]	3.3 (1.8 to 5.9)	3.3 (1.8 to 5.9)	3.3 (1.8 to 5.9)	-	-	-	3.3 (1.8 to 5.9)	3.3 (1.8 to 5.9)
Dregan 2017[14]	5.34 (3.82 to 7.45)	5.34 (3.82 to 7.45)	5.34 (3.82 to 7.45)	-	-	5.34 (3.82 to 7.45)	-	-
Hak 2009[16]	2.29 (0.85 to 6.15)	2.51 (0.94 to 6.69)	-	2.29 (0.85 to 6.15)	2.29 (0.85 to 6.15)	2.29 (0.85 to 6.15)	-	2.29 (0.85 to 6.15)
Hermansen 2017[17]	2.4 (1.8 to 3.2)	2.4 (1.8 to 3.2)	2.4 (1.8 to 3.2)	-	2.4 (1.8 to 3.2)	2.4 (1.8 to 3.2)	-	2.4 (1.8 to 3.2)
Krishnan 2005[19]	1.67 (1.46 to 1.92)	1.54 (1.34 to 1.77)	-	1.67 (1.46 to 1.92)	-	-	1.67 (1.46 to 1.92)	-
Lim 2018[20]	3.31 (2.84 to 3.86)	3.61 (3.10 to 4.19)	3.31 (2.84 to 3.86)	-	3.31 (2.84 to 3.86)	-	3.31 (2.84 to 3.86)	3.31 (2.84 to 3.86)
Liou 2014[22]	-	-	1.88 (1.08 to 3.27)	-	-	-	-	-
Mok 2009[24]	2.02 (1.3 to 3.81)	2.02 (1.3 to 3.81)	-	2.02 (1.3 to 3.81)	-	2.02 (1.3 to 3.81)	-	2.02 (1.3 to 3.81)
Rees 2016[26]	1.47 (1.2 to 1.8)	1.81 (1.49 to 2.19)	1.47 (1.2 to 1.8)	-	1.47 (1.2 to 1.8)	-	1.47 (1.2 to 1.8)	1.47 (1.2 to 1.8)
Wang 2012[27]	2.9 (2.52 to 3.33)	3.17 (2.79 to 3.61)	-	2.9 (2.52 to 3.33)	2.9 (2.52 to 3.33)	-	2.9 (2.52 to 3.33)	2.9 (2.52 to 3.33)
Ward 1999[28]	-	-	-	-	-	-	-	-
Intracerebral haemo	orrhage							
Authors	Main analysis	Least adjusted	Published during or after 2014	Published before 2014	Only studies with low risk of bias	Fatal/non-fatal	Non-fatal	Excluding cross- sectional studies
Arkema 2017[4]	1.4 (0.7 to 2.9)	1.6 (0.8 to 3.2)	1.4 (0.7 to 2.9)	-	1.4 (0.7 to 2.9)	1.4 (0.7 to 2.9)	-	1.4 (0.7 to 2.9)
Krishnan 2005[19]	1.56 (1.11 to 2.21)	1.26 (0.89 to 1.77)	-	1.56 (1.11 to 2.21)	-	-	1.56 (1.11 to 2.21)	-
Mok 2009[24]	1.03 (0.26 to 4.11)	1.03 (0.26 to 4.11)	-	1.03 (0.26 to 4.11)	-	1.03 (0.26 to 4.11)	-	1.03 (0.26 to 4.11)
Wang 2012[27]	2.91 (2.12 to 3.99)	3.5 (2.64 to 4.64)	-	2.91 (2.12 to 3.99)	2.91 (2.12 to 3.99)	-	2.91 (2.12 to 3.99)	2.91 (2.12 to 3.99)
Zoller 2012[29]	-	-	-	2.65 (1.81 to 3.74)	-	-	2.65 (1.81 to 3.74)	-
Ischaemic stroke								
Authors	Main analysis	Least adjusted	Published during or after 2014	Published before 2014	Only studies with low risk of bias	Fatal/non-fatal	Non-fatal	Excluding cross- sectional studies

Arkema 2017[4]	2.2 (1.8 to 2.8)	2.4 (1.9 to 3.0)	2.2 (1.8 to 2.8)	-	2.2 (1.8 to 2.8)	2.2 (1.8 to 2.8)	-	2.2 (1.8 to 2.8)
Avina-Zubieta 2017[5]	2.14 (1.64 to 2.79)	2.81 (2.24 to 3.53)	2.14 (1.64 to 2.79)	-	2.14 (1.64 to 2.79)	2.14 (1.64 to 2.79)	-	2.14 (1.64 to 2.79)
Chiu 2012[12]	1.67 (1.45 to 1.91)	1.61 (1.39 to 1.87)	-	1.67 (1.45 to 1.91)	1.67 (1.45 to 1.91)	-	1.67 (1.45 to 1.91)	1.67 (1.45 to 1.91)
Krishnan 2005[19]	2.27 (1.89 to 2.73)	2.01 (1.67 to 2.42)	-	2.27 (1.89 to 2.73)	-	-	2.27 (1.89 to 2.73)	
Mok 2009[24]	3.72 (2.34 to 5.91)	3.72 (2.34 to 5.91)	-	3.72 (2.34 to 5.91)	-	3.72 (2.34 to 5.91)	-	3.72 (2.34 to 5.91)
Zoller 2012[29]	-	-	-	1.94 (1.68 to 2.24)	-	-	1.94 (1.68 to 2.24)	-
Wang 2012[27]	-	-	-	-	-	-	-	-
Liou 2014[22]	-	-	-	-	-	-	-	-
Bernatsky 2006b[9]	-	-	-	-	-	-	-	-
Subarachnoid haem	orrhage							
Authors	Main analysis	Least adjusted	Published during or after 2014	Published before 2014	Only studies with low risk of bias	Fatal/non-fatal	Non-fatal	Excluding cross- sectional studies
Arkema 2017[4]	1.4 (0.5 to 3.9)	1.6 (0.6 to 4.3)	1.4 (0.5 to 3.9)	-	1.4 (0.5 to 3.9)	1.4 (0.5 to 3.9)	-	1.4 (0.5 to 3.9)
Krishnan 2005[19]	0.53 (0.32 to 0.89)	0.57 (0.34 to 0.96)	-	0.53 (0.32 to 0.89)	-	-	0.53 (0.32 to 0.89)	-
Ramagopalan 2013[25]	3.76 (3.08 to 4.55)	3.76 (3.08 to 4.55)	3.76 (3.08 to 4.55)	-	3.76 (3.08 to 4.55)	3.76 (3.08 to 4.55)	-	3.76 (3.08 to 4.55)
Wang 2012[27]	4.8 (2.66 to 8.67)	5.16 (2.97 to 8.95)	-	4.8 (2.66 to 8.67)	4.8 (2.66 to 8.67)	-	4.8 (2.66 to 8.67)	4.8 (2.66 to 8.67)
Chang 2013[11]	-	-	-	-	-	-	-	-
Bernatsky 2006b[9]	-	-	-	-	-	-	-	-
Myocardial infarction	on							
Authors	Main analysis	Least adjusted	Published during or after 2014	Published before 2014	Only studies with low risk of bias	Fatal/non-fatal	Non-fatal	Including studies only reporting LN
Avina-Zubieta 2017[5]	2.61 (2.12 to 3.2)	3.04 (2.5 to 3.69)	2.61 (2.12 to 3.2)	-	2.61 (2.12 to 3.2)	2.61 (2.12 to 3.2)	-	2.61 (2.12 to 3.2)
Barnado 2018[6]	2.26 (1.53 to 3.35)	2.26 (1.53 to 3.35)	2.26 (1.53 to 3.35)	-	2.26 (1.53 to 3.35)	-	2.26 (1.53 to 3.35)	2.26 (1.53 to 3.35)
Bengtsson 2012[7]	-	-	-	2.13 (1.34 to 3.7)	-	-	-	-
Cook 2018[13]	2.9 (1.5 to 5.3)	2.9 (1.5 to 5.3)	2.9 (1.5 to 5.3)	-	-	-	2.9 (1.5 to 5.3)	2.9 (1.5 to 5.3)
Faurschou 2011[15]	-	-	-	-	-	-	-	7.9 (3.8 to 15)
Hak 2009[16]	1.81 (0.75 to 4.37)	1.81 (0.75 to 4.37)	-	1.81 (0.75 to 4.37)	1.81 (0.75 to 4.37)	1.81 (0.75 to 4.37)	-	1.81 (0.75 to 4.37)
Hermansen 2017[17]	3 (2 to 4.5)	3 (2 to 4.5)	3 (2 to 4.5)	-	3 (2 to 4.5)	3 (2 to 4.5)	-	3 (2 to 4.5)
Kim 2017[18]	4.1 (3.9 to 4.1)	4.1 (3.9 to 4.1)	4.1 (3.9 to 4.1)	-	4.1 (3.9 to 4.1)	-	4.1 (3.9 to 4.1)	4.1 (3.9 to 4.1)
Lim 2018[20]	2.74 (2.28 to 3.37)	3.13 (2.63 to 3.73)	3.13 (2.63 to 3.73)	-	3.13 (2.63 to 3.73)	-	3.13 (2.63 to 3.73)	3.13 (2.63 to 3.73)

Lin 2014[21]	5.11 (2.63 to 9.92)	5.11 (2.63 to 9.92)	5.11 (2.63 to 9.92)	-	5.11 (2.63 to 9.92)	5.11 (2.63 to 9.92)	-	5.11 (2.63 to 9.92)
--------------	---------------------	---------------------	---------------------	---	---------------------	---------------------	---	---------------------

In the case of population overlap, an additional analysis was performed for each overlap study replacing the study used in the main analysis with the overlapping study.

LN, lupus nephritis; MI, myocardial infarction; SLE, systemic lupus erythematosus.

Supplementary Figure S1. Forest plot of risk ratios and stroke/MI in adult patients with SLE compared with the general population or healthy controls by age at SLE diagnosis, for stroke and subtypes

Study	Outcome	Age	RR (95% CI)		
<30 years	Cuttomic	Age			
Mok 2009	Ischaemic or haemorrhagic stroke	<30 years	22.8 (6.67 to 91.7)		_
Wang 2012	All stroke	18-24 years	36.6 (15.2 to 83.4)		_
Mok 2009	Ischaemic stroke	<30 years	53.9 (7.47 to 389)		_
Mok 2009	Haemorrhagic stroke	<30 years	14.5 (2.03 to 103)		──■ →
30–40 years					
Mok 2009	Ischaemic or haemorrhagic stroke	30–40 years	21 (7.84 to 56.5)		
Rees 2016	Stoke	<40 years	15 (4.13 to 54.49)		
Wang 2012	All stroke	25–34 years	13 (7.89 to 21.5)		_ _
Mok 2009	Ischaemic stroke	30-40 years	29.1 (9.28 to 91)		
Mok 2009	Haemorrhagic stroke	30-40 years	11.5 (1.61 to 82.2)		
40–50 years					
Mok 2009	Ischaemic or haemorrhagic stroke	40–50 years	7.44 (3.33 to 16.6)		
Wang 2012	All stroke	35-44 years	4.34 (2.99 to 6.29)		
Zoller 2012a	Haemorrhagic stroke	<50 years	7.5 (2.7 to 16.43)		
Zoller 2012a	Ischaemic stroke				_
Mok 2009	Ischaemic stroke	40–50 years	10.9 (4.88 to 24.4)		
50–60 years					
Arkema 2017	Ischaemic stroke	50 to <60 years	1.7 (0.9 to 3.1)	-	
Avina-Zubieta 2017		45–59 years	3.29 (1.89 to 5.73)		_
Mok 2009	Ischaemic or haemorrhagic stroke		1.88 (0.6 to 5.84)		
Wang 2012	All stroke	45–54 years	1.68 (1.18 to 2.39)		- -
Zoller 2012a	Haemorrhagic stroke	50–59 years	3.1 (0.81 to 8.02)		
Mok 2009	Ischaemic stroke	50–60 years	3.37 (1.26 to 9.02)		-
60–70 years					
Arkema 2017	Ischaemic stroke	60+ years	2.1 (1.6 to 2.8)		
Avina-Zubieta 2017	Ischaemic stroke	60–74 years	1.49 (0.95 to 2.35)		
Mok 2009	Ischaemic or haemorrhagic stroke		2.8 (1.03 to 7.6)		
Wang 2012	All stroke	55–64 years	1.68 (1.18 to 2.4)		
Ward 1999	Cardiovascular accident	45-64 years	0.96 (0.7 to 1.3)		
Zoller 2012a	Haemorrhagic stroke	60–69 years	1.05 (0.1 to 3.85)		
Arkema 2017	Intracerebral harmorrhage	60+ years	1 (0.4 to 2.4)		
Mok 2009	Ischaemic stroke	60–70 years	2.52 (0.8 to 7.94)	_	
70+ years	lachaomia atraka	75	1 76 (1 10 to 0 70)		
Avina-Zubieta 2017 Mok 2009	Ischaemic stroke Ischaemic or haemorrhagic stroke	75+ years	1.76 (1.12 to 2.78) 0.53 (0.07 to 3.81)		
Rees 2016	Stroke	70+ years 70+ years	1.51 (1.16 to 1.95)	-	
	All stroke	70+ years 65+ years	•	_	
Wang 2012 Ward 1999	Cardiovascular accident	65+ years	0.8 (0.56 to 1.12) 0.76 (0.59 to 0.98)		T
Zoller 2012a	Haemorrhagic stroke	70+ years	1.57 (0.75 to 2.91)		
Mok 2009	Ischaemic stroke	70+ years 70+ years	0.63 (0.09 to 4.56)		
MOR 2003		ior years	0.00 (0.09 (0 4.00)		
			C).10 1	.0 10.0 100.0

Descriptive analysis. No meta-analysis or synthesis was conducted.

CI, confidence interval; MI, myocardial infarction; RR, risk ratio; SLE, systemic lupus erythematosus.

References

- 1. Hochberg MC. Updating the American College of Rheumatology revised criteria for the classification of systemic lupus erythematosus. *Arthritis Rheum* 1997;40:1725. doi: 10.1002/1529-0131(199709)40:9<1725::AID-ART29>3.0.CO;2-Y.
- 2. Tan EM, Cohen AS, Fries JF, et al. The 1982 revised criteria for the classification of systemic lupus erythematosus. *Arthritis Rheum* 1982;25:1271–77. doi: 10.1002/art.1780251101.
- Wells G, Shea B, O'Connell D, et al. The Newcastle-Ottawa Scale (NOS) for assessing the quality of nonrandomised studies in meta-analyses. 2014. Available from: <u>http://www.ohri.ca/programs/clinical_epidemiology/oxford.asp</u>. Accessed October 21, 2019.
- Arkema VE, Svenungsson E, Von Euler M, et al. Stroke in systemic lupus erythematosus: a Swedish population-based cohort study. *Ann Rheum Dis* 2017;76:1544–49. doi: 10.1136/annrheumdis-2016-210973.
- 5. Avina-Zubieta JA, To F, Vostretsova K, et al. Risk of myocardial infarction and stroke in newly diagnosed systemic lupus erythematosus: a general population-based study. *Arthritis Care Res* 2017;69:849–56. doi: 10.1002/acr.23018.
- 6. Barnado A, Carroll RJ, Casey C, et al. Phenome-wide association study identifies marked increased in burden of comorbidities in African Americans with systemic lupus erythematosus. *Arthritis Res Ther* 2018;20:69. doi: 10.1186/s13075-018-1561-8.
- Bengtsson C, Ohman ML, Nived O, et al. Cardiovascular event in systemic lupus erythematosus in northern Sweden: incidence and predictors in a 7-year follow-up study. *Lupus* 2012;21:452–59. doi: 10.1177/0961203311425524.
- 8. Bernatsky S, Boivin JF, Joseph L, et al. Mortality in systemic lupus erythematosus. *Arthritis Rheum* 2006;54:2550–57. doi: 10.1002/art.21955.
- 9. Bernatsky S, Clarke A, Gladman DD, et al. Mortality related to cerebrovascular disease in systemic lupus erythematosus. *Lupus* 2006;15:835–39.
- 10. Björnådal L, Yin L, Granath F, et al. Cardiovascular disease a hazard despite improved prognosis in patients with systemic lupus erythematosus: results from a Swedish population based study 1964-95. *J Rheumatol* 2004;31:713–19. doi: 0315162X-31-713.
- 11. Chang Y-SS, Liu C-JJ, Chen W-SS, et al. Increased risk of subarachnoid hemorrhage in patients with systemic lupus erythematosus: a nationwide population-based study. *Arthritis Care Res* 2013;65:601–06. doi: 10.1002/acr.21846.
- 12. Chiu CC, Huang CC, Chan WL, et al. Increased risk of ischemic stroke in patients with systemic lupus erythematosus: a nationwide population-based study. *Intern Med* 2012;51:17–21.
- 13. Cook MJ, Bellou E, Bowes J, et al. The prevalence of co-morbidities and their impact on physical activity in people with inflammatory rheumatic diseases compared with the general population: results from the UK Biobank. *Rheumatology (Oxford)* 2018;57:2172–82. doi: 10.1093/rheumatology/key224.
- 14. Dregan A, Chowienczyk P, Molokhia M. Cardiovascular and type 2 diabetes morbidity and allcause mortality among diverse chronic inflammatory disorders. *Heart* 2017;103:1867–73. doi: 10.1136/heartjnl-2017-311214.
- 15. Faurschou M, Mellemkjaer L, Starklint H, et al. High risk of ischemic heart disease in patients with lupus nephritis. *J Rheumatol* 2011;38:2400–05. doi: 10.3899/jrheum.110329.
- Hak AE, Karlson EW, Feskanich D, et al. Systemic lupus erythematosus and the risk of cardiovascular disease: results from the nurses' health study. *Arthritis Rheum* 2009;61:1396–402. doi: 10.1002/art.24537.
- 17. Hermansen ML, Lindhardsen J, Torp-Pedersen C, et al. The risk of cardiovascular morbidity and cardiovascular mortality in systemic lupus erythematosus and lupus nephritis: a Danish

nationwide population-based cohort study. *Rheumatology* 2017;56:709–15. doi: 10.1093/rheumatology/kew475.

- 18. Kim CH, Al-Kindi SG, Jandali B, et al. Incidence and risk of heart failure in systemic lupus erythematosus. *Heart* 2017;103:227–33. doi: 10.1136/heartjnl-2016-309561.
- 19. Krishnan E. Stroke subtypes among young patients with systemic lupus erythematosus. *Am J Med* 2005;118:1415. doi: 10.1016/j.amjmed.2005.026.
- 20. Lim SY, Bae EH, Han KD, et al. Systemic lupus erythematosus is a risk factor for cardiovascular disease: a nationwide, population-based study in Korea. *Lupus* 2018;27:2050–56. doi: 10.1177/0961203318804883.
- 21. Lin CY, Shih CC, Yeh CC, et al. Increased risk of acute myocardial infarction and mortality in patients with systemic lupus erythematosus: two nationwide retrospective cohort studies. *Int J Cardiol* 2014;176:847–51. doi: /10.1016/j.ijcard.2014.08.006.
- 22. Liou TH, Huang SW, Lin JW, et al. Risk of stroke in patients with rheumatism: a nationwide longitudinal population-based study. *Sci Rep* 2014;4:5110. doi: 10.1038/srep05110.
- 23. Manzi S, Meilahn EN, Rairie JE, et al. Age-specific incidence rates of myocardial infarction and angina in women with systemic lupus erythematosus: comparison with the Framingham Study. *Am J Epidemiol* 1997;145:408–15.
- 24. Mok CC, Ho LY, To CH. Annual incidence and standardized incidence ratio of cerebrovascular accidents in patients with systemic lupus erythematosus. *Scand J Rheumatol* 2009;38:362–68. doi: 10.1080/03009740902776927.
- 25. Ramagopalan VS, Pakpoor J, Seminog O, et al. Risk of subarachnoid haemorrhage in people admitted to hospital with selected immune-mediated diseases: record-linkage studies. *BMC Neurol* 2013;13:176. doi: 10.1186/1471-2377-13-176.
- 26. Rees F, Doherty M, Grainge M, et al. Burden of comorbidity in systemic lupus erythematosus in the UK, 1999-2012. *Arthritis Care Res* 2016;68:819–27. doi: 10.1002/acr.22751.
- 27. Wang IK, Muo CH, Chang YC, et al. Risks, subtypes, and hospitalization costs of stroke among patients with systemic lupus erythematosus: a retrospective cohort study in Taiwan. *J Rheumatol* 2012;39:1611–18. doi: 10.3899/jrheum.111510.
- 28. Ward MM. Premature morbidity from cardiovascular and cerebrovascular diseases in women with systemic lupus erythematosus. *Arthritis Rheum* 1999;42:338–46. doi: 10.1002/1529-0131(199902)42:2<338::AID-ANR17>3.0.CO;2-U.
- 29. Zoller B, Li X, Sundquist J, et al. Risk of subsequent ischemic and hemorrhagic stroke in patients hospitalized for immune-mediated diseases: a nationwide follow-up study from Sweden. *BMC Neurol* 2012;12:41. doi: 10.1186/1471-2377-12-41.