

A Trans-Atlantic Analysis of Real-world Evidences showing preventive Effects of Statins on Cancer Incidence

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Supplementary Methods

Study population

The total study population includes patients from the Charité and UVA hospitals. Therefore, in an initial data-processing stage, all diagnoses and discharge medications were extracted, merged, unified and transformed into a centralized database. The medications for the Charité datasets were available in the doctor letters at discharge. The medications were automatically extracted based on a complete list of German drug products, including an ingredient mapping. At the UVA hospital, an EHR was established in 2010, in which the discharge medication could be extracted. Cancer was defined by ICD10 diagnoses codes from 'C00' to 'D48' with an exclusion of ICD10 codes 'D10' to 'D37'. Newborns and patients older than 100 years were excluded from the study.

This process yielded in 277,980 distinct patients, of whom 53,113 patients were diagnosed with cancer (Age and gender distribution is visualized in Supplementary Figure 2).

All patient records were anonymized prior to study cohort selection in accordance with the Health Insurance Portability and Accountability Act (HIPAA) [1]. The Ethical Review Committee of the Charité – Universitätsmedizin Berlin granted formal approval for this study (EA1/171/15), as well as the Institutional Review Board (IRB) of the University of Virginia (UVA IRB-HSR Study Tracking # 22282). A Data Use Agreement (DUA) between UVA and Charite, as regards the Statins project, was finally executed.

Trial design / Cohort design

The RWE for treatment of cancer with statins in a large scale, multicentric, randomized observational study was scrutinized. The Charité patient data, as well as the UVA patients, were separately analyzed, and also considered in a transatlantic cohort. A detailed overview of the dataset is shown in Supplementary Table 1. Patients with a cancer diagnosis were randomly matched 1:1 with non-cancer patients according to age and gender.

Given statistically significant large case counts, subtypes of cancer were analyzed. High frequency cancer entities, like breast, colorectal, liver, pancreas or prostate were separately considered. In this context (based on sufficient data), Atorvastatin, Fluvastatin, Lovastatin, Pravastatin, Rosuvastatin and Simvastatin were inspected individually.

The primary end-point of this study describes the status of patients at discharge considering the transatlantic cohort. Due to incomplete data, patient progression is not available for a significant number of cases. The second endpoint describes the 1:1 matched study including cancer subtypes by considering the different statins.

Confounder / Co-medication

Different confounding factors were considered during the planning phase. Because of the variability in prescription or diagnosis habits by doctors, the data could be biased. Here, the study design was set-up as a transatlantic cohort, including two completely independent datasets, to minimize such effects. In this regard, other confounding factors, such as family history, will be minimized.

Further, to mitigate the risk of describing false-positive cancer preventive effect of statins, co-medication for every patient was extensively studied. Therefore, the most commonly prescribed drugs along with statins were considered. Each drug was then evaluated independently of each other (Figure 3 - Co-Medication). Specifically, we also analyzed whether co-diagnoses (based on ICD10 codes) like hypercholesterolemia (E78 & Z83), ischemic heart (I20-I25), hypertension (I10), stroke (I63), diabetes (E10-E14), obesity (E65-E68) or chronic kidney diseases (N18) influence the conclusion reached in our study.

Statistical analysis

Our primary goal was to establish whether or not statins have a cancer preventive effect. RWE studies generally cannot prove a causal inference, given effect modifiers and confounding factors. A stratified analysis was performed in this study, considering an overall cancer preventive effect for the collection of all prescribed statins as well as analyzing different cancer subtypes and individual statins separately; moreover, it is also for this reason that the study was conducted across two distinct cohorts, one in Germany and the other in the U.S.

As common measures of the effect size in randomized controlled trials, Odds ratio (OR) and risk ratio (RR) were calculated for the respective case and control groups. To account for the study design, a 1:1 matched analysis was performed, taking into consideration the contingency-table for different statins and cancer subtypes. In order to overcome artifacts from the matching process, this procedure was performed 25 times and OR and RR were considered as mean and median values.

Both measurements are reported with the 95%-confidence intervals and p-values. All p-values are two-sided for which 0.025 were considered to indicate statistical significance. We statistically analyzed the mRNA expression data with a one-way analysis of variance (ANOVA) procedure, including Dunnett's multiple comparison test and a confidence interval of 95%. The functional assays were analyzed using two-way ANOVA, including Turkey's additivity test and a confidence interval of 95% (* = $p < 0.05$, ** = $p < 0.01$, *** = $p < 0.001$, **** = $p < 0.0001$).

Cell lines and growth conditions

Human cancer cell lines HCT116 (CRC), HT-29 (CRC), SW48 (CRC), BxPC3 (pancreatic cancer) were purchased from the American Type Culture Collection and MKN45 (gastric cancer) was kindly provided by Experimentelle Pharmakologie & Onkologie Berlin-Buch GmbH (EPO GmbH, Berlin, Germany). Cells were grown in DMEM or RPMI1640 medium (Thermo Fisher Scientific, Waltham, Massachusetts) supplemented with 10% fetal bovine serum (Bio & Sell, Feucht, Germany) in a humidified incubator at 37°C with 5% CO₂. Cell lines were regularly tested for mycoplasma using the MycoAlert Mycoplasma

detection kit (Lonza, Basel, Switzerland). We verified the authentication of all cell lines by short tandem repeat (STR) genotyping at the Leibniz-Institute DSMZ (Braunschweig, Germany): STR genotypes concurred with published genotypes.

Drug treatment was performed with statins purchased from Selleckchem (Houston, Texas) (atorvastatin) or Thermo Fisher Scientific (Waltham, Massachusetts) (fluvastatin, simvastatin, lovastatin). The 20 mM stock solutions were prepared freshly for every application in dimethylsulfoxide (DMSO). Control cells were treated with equal amounts of solvent to rule out adverse effects caused by DMSO.

Transfection of human CRC cells

Cell line generation for HTS was described previously [2]. For constitutive or inducible MACC1 overexpression, cells were transduced with lentiviruses produced in HEK293T cells using the plasmids RC224774L2 (MACC1-GFP, Origene, Rockville, Maryland) or pCW-GW-rtTA/MACC1-P2A-nLuc (kindly provided by Dr. Nikolas Gunkel, DKFZ, Heidelberg) and pMD2.G, psPAX2 for packaging following standard protocols. pMD2.G and psPAX2 were kind gifts from Didier Trono (Addgene plasmid # 12259, # 12260). Cells were selected either by FACS or blasticidin (Invivogen, San Diego, California) treatment. Knock-out of MACC1 in HCT116 cells (HCT116/MACC1^{-/-}) was performed according to the protocol by Ran et al. with the vector pSpCas9(BB)-2A-GFP (Addgene, Watertown, Massachusetts) and the single guide RNAs: sgMACC1 fwd – GTT TGA AGA GTA CCC GGG TTT GG and sgMACC1 rev – ACA TGC CTT GCT CCG TAT GCA GG (Biotex, Berlin, Germany) [3].

High throughput drug screening (HTS)

We seeded HCT116-MACC1p-Luc CRC cells stably expressing the MACC1-promoter driven luciferase gene in 384-well plates (Perkin Elmer, Waltham, Massachusetts) at 4,000 cells/well. The plates already contained the 4241 compounds in two different concentrations: 5 µg/ml and 0.5 µg/ml of Prestwick library compounds, and 1 µM and 0.1 µM of NIH and Microsource library compounds, respectively.

Cells were incubated with compounds for 24 h at 37°C in a humidified incubator with 5% CO₂. To assess the luminescent signal, plates were treated with 25 µl BriteLite plus luminescent reagent for 5 min (Perkin Elmer, Waltham, Massachusetts) and measured in an Envision Reader with ultrasensitive luminescence detector (Perkin Elmer, Waltham, Massachusetts).

In vivo validation of statins

Animal experiments were conducted according to the United Kingdom Coordinating Committee of Cancer Research (UKCCCR) guidelines and granted by the State Office of Health and Social Affairs (Landesamt für Gesundheit und Soziales, LaGeSo, Berlin, Germany).

We transplanted 3x10⁵ of HCT116/CMVp-Luc cells into the spleens of 6-week-old female SCID-beige mice (Charles River, Wilmington, Massachusetts). Mice were randomly assigned to 3 groups of 10 animals for oral application with daily doses of either solvent (10% Kolliphor in 0.9% NaCl) or a human equivalent dose of approximately 1 mg/kg body weight (13 mg/kg body weight) of fluvastatin or atorvastatin (in 0.9% NaCl). This dose is widely used in blood lipid reduction therapy [4,5]. We continuously monitored tumor growth and metastasis formation via intraperitoneal application of 150 mg/kg D-luciferin (Biosynth, Staad, Switzerland) and the bioluminescence imaging system NightOWL LB 981 (Berthold Technologies, Bad Wildbad, Germany). For imaging and quantification, WinLight (Berthold Technologies) and ImageJ (version 1.51j8, National Institutes of Health, USA) were used. The experiment was terminated due to ethical reasons. Afterwards, spleens and livers were removed and shock frozen in liquid nitrogen for preparation of cryosections and further analysis. Extraction of RNA and genomic DNA with subsequent qRT-PCR and qPCR analysis was performed as described previously.

For immunohistochemistry (IHC) cryosections were fixed with 4% para-formaldehyde (PFA, Carl Roth GmbH & Co. KG, Karlsruhe, Germany) and quenched with 0.1 M glycine (Carl Roth GmbH & Co. KG, Karlsruhe, Germany). Blocking of endogenous peroxidases with 3% H₂O₂ (Carl Roth GmbH & Co. KG, Karlsruhe, Germany) was followed by permeabilization with 0.2% TritonX100 and blocking with 5%

bovine serum albumin (BSA). Primary antibody (1:200 in 2.5% BSA, anti-CK19; OriGene Technologies Inc., Rockville, Maryland) incubation was carried out for 2 h followed by 5 washing steps and 1 h secondary antibody incubation (1:500 in 2.5% BSA, anti-rabbit-HRP; Promega, Madison, Wisconsin) at room temperature. Final staining with DAB solution (DAKO DAB Liquid + chromogen substrate; Agilent Technologies, Santa Clara, California) and nuclear staining with hemalum (Carl Roth GmbH & Co. KG, Karlsruhe, Germany) was followed by washing under running tap water. Slices were mounted on microscope slides with Glycergel (Agilent Technologies, Santa Clara, California) and imaged with a Keyence BZ-X800 microscope (Keyence, Osaka, Japan).

RNA extraction and qRT-PCR

Drug treatment was performed in 12-well dishes with 1.25×10^5 cells/well. To isolate total RNA, we used the Universal RNA Purification Kit (Roboklon, Berlin, Germany) according to manufacturer's instructions. RNA was quantified (Nanodrop, Peqlab, Erlangen, Germany) and 50 ng RNA were applied to reverse transcription with random hexamers in a reaction mix (5 mM MgCl₂, 1x RT-buffer, 4 mM pooled dNTPs, 1 U/μl RNase inhibitor and 2.5 U/μL Moloney Murine Leukemia Virus reverse transcriptase; Thermo Fisher Scientific) at 23°C for 15 min, 42°C for 45 min, 99°C for 5 min with subsequent cooling at 4°C for 5 min. The LightCycler 480 (Roche Diagnostics, Mannheim, Germany) with GoTaq dye (Promega, Fitchburg, Wisconsin) chemistry was employed for cDNA amplification in quantitative polymerase chain reaction (qPCR) under following conditions: 95°C for 2 min followed by 45 cycles of 95°C for 7 s, 60°C for 10 s and 72°C for 5 s. Primers for MACC1 and G6PD were described previously [20]. Data was analyzed with the LightCycler 480 Software release 1.5.0SP3 (RocheDiagnostics). Duplicate qRT-PCR reaction values were averaged and each mean value of the expressed gene was normalized to the respective mean amount of the G6PD cDNA.

Protein extraction and Western blotting

Similar to RNA extraction, 1.25×10^5 cells were seeded in 12-well plates. After drug treatment, cells were scraped off in ice cold RIPA buffer (50 mM Tris, 150 mM NaCl and 1% Nonidet P-40; pH 7.5 supplemented with complete protease inhibitor tablets; Roche Diagnostics) and lysed for 30 min on ice. We used the Bicinchoninic Acid Protein Assays Reagent (Thermo Fisher Scientific) to determine protein concentration according to manufacturer's instructions. Equal amounts of protein were separated by SDS-PAGE and transferred to PVDF membranes (Bio-Rad Laboratories Inc., Hercules, California). Membranes were blocked for 1 h at room temperature with 5% BSA in TBST buffer (10mM Tris-HCl, 0.1% Tween20 and 150 mM NaCl; pH 7.5). Incubation of membranes with rabbit anti-MACC1 antibody (Sigma-Aldrich, dilution 1:10,000) or mouse anti- β -actin (Sigma-Aldrich, dilution 1:25,000) and vinculin (Sigma-Aldrich, 1:2,000) at 4°C overnight was followed by incubation with HRP-conjugated anti-rabbit IgG (Promega, dilution 1:20,000) or anti-mouse IgG (Thermo Fisher Scientific, dilution 1:20,000) for 1 h at room temperature. Antibody-protein complexes were visualized with WesternBright ECL HRP substrate (Advansta, Menlo Park, California) and subsequent exposure to CL-Xposure Films (Thermo Fisher Scientific). Immunoblotting for β -actin and vinculin served as protein loading control.

Proliferation assay

Cell viability and proliferation was assessed using the IncuCyte® ZOOM System (Essen BioScience, Ann Arbor, Michigan). 5×10^3 cells were seeded in 96-well plates and allowed to accommodate for 24 h before drug treatment. After drug treatment, plates were placed into the IncuCyte for 72 h at 37°C in a humidified incubator with 5% CO₂. Four pictures of each well were taken every 2 h which allowed a detailed analysis of the area covered by proliferating cells over time. For data analysis and export the IncuCyte ZOOM software 2016B was used. Each cell proliferation experiment was performed in triplicate. Results are expressed as confluence percent area under the curve of cell confluence compared normalized to solvent-treated controls.

Clonogenic assay

We used the clonogenic assay to assess the reproductive viability and ability to form colonies. 400 cells/well of HCT116/GFP and HCT116/MACC1-GFP cells were seeded in 6-well dishes and allowed to attach for 24 h before drug treatment. After drug treatment, plates were placed into a humidified incubator at 37°C with 5% CO₂ for 7 days. Medium was removed and the colonies were fixed and stained with a solution of PBS containing 1% formaldehyde and 0.1% crystal violet. The colony covered area was determined with the colony area plug in of ImageJ (version 1.51j8, National Institutes of Health, USA). Results are expressed as area percent compared to solvent-treated.

References for Supplementary Methods

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5. Stone NJ, Robinson JG, Lichtenstein AH, Bairey Merz CN, Blum CB, Eckel RH, et al. 2013 ACC/AHA guideline on the treatment of blood cholesterol to reduce atherosclerotic cardiovascular risk in adults: a report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines. *J Am Coll Cardiol.* 2014; 63: 2889-934.

Supplementary Table 1: Statistical characteristics of the sampled population and study design.
[caption on next page]

Characteristics	Full Trans-Atlantic Cohort			1:1 Matched Cohort		
	Statin use?		p-value	Statin use?		p-value
	✓	✗		✓	✗	
All Subjects (#)	42,682	235,298		17,941	88,279	
Age (yrs)						
Median	69	54		68	59	
Interquartile range	60–76	34–69		62–76	51–72	
Gender (# [%])			< 0.001			< 0.001
♀	16,903 [40]	114,640 [49]		11,718 [65]	46,669 [52]	
♂	25,779 [60]	120,658 [51]		6,223 [34]	41,610 [47]	
Cancer patients by Age (median yrs)						
♀	70	59		70	59	
♂	70	61		70	60	
Cancer patients by Gender (# [%])			< 0.001			< 0.001
♀	2,154 [33]	21,791* [46]		2,154 [33]	21,790 [46]	
♂	4,349* [67]	24,896 [54]		4,348 [67]	24,896 [54]	
Censored Confounders (# subjects)						
Hypercholesterolemia			< 0.001			
♀	987	13,366				
♂	2,098	16,744				
Hypertension			< 0.001			
♀	623	9,240				
♂	1,391	11,370				
Ischemic heart			< 0.001			
♀	1,369	13,269				
♂	2,151	15,974				
Stroke			< 0.001			
♀	1,630	13,669				
♂	3,514	17,390				
Obesity			< 0.001			
♀	1,589	13,231				
♂	3,429	16,918				
Diabetes			< 0.001			
♀	1,178	11,914				
♂	2,398	14,342				
Chronic kidney disease			< 0.001			
♀	1,451	12,758				
♂	2,989	15,782				

This Table summarizes characteristic features and properties of the population sampled in this work, for both of the trans-Atlantic cohorts (Charite, UVA) as well as the 1:1 matched cohort. Both cohorts are split into statin and non-statin patients. If determined, a p-value for the underlying condition is provided. The censored confounders are included as well as interquartile ranges for the age distribution for statin and non-statin users. Note that the columns in the bottom half of the table (i.e., below the Censored Confounders section) contain values that apply to both cohorts, "full trans-Atlantic" and "1:1 matched": this is intrinsic to the layout of the table (the "Statin use?" query applies in both cohorts), and thus the data are not replicated in the bottom-half columns. A similar pattern occurs in parts of the top-half of the table too, though the data there are explicitly provided because the pairs of numerical values are not perfectly identical (e.g., the "Cancer patients by Gender" row has a discrepancy of two patients between the two cohorts, for the parameters marked by '*' in the first pair of columns).

Supplementary Table 2

In this table, the co-diagnoses of cancer were analyzed, taking into consideration statin intake as well. In tab 1 "Counts", co-diagnoses are calculated. The resulting diagnoses are presented via a short description as well as the ICD10 code. "Counts" consists of the number of patients, also diagnosed together with cancer in descending order. The second tab "Odds-Ratios", shows the ORs. The first results row displays the OR for cancer given (statins, atorvastatin, simvastatin) without exclusion of any co-diagnoses. Results are displayed for the transatlantic as well as 1:1 matched cohort.

Tab 1:

Co-diagnoses		
ICD10-Code	Counts	Description
Z11	35.397	Encounter for screening for infectious and parasitic diseases
I10	33.633	Essential primary hypertension
E87	16.140	Other disorders of fluid electrolyte and acid-base balance
Z45	15.226	Encounter for adjustment and management of implanted device
Z92	13.022	Personal history of medical treatment
E11	12.614	Type 2 diabetes mellitus
N18	10.697	Chronic kidney disease CKD
Z29	9.638	Encounter for other prophylactic measures
Z95	9.472	Presence of cardiac and vascular implants and grafts
I25	8.454	Chronic ischemic heart disease
E78	8.361	Disorders of lipoprotein metabolism and other lipidemias
E03	7.921	Other hypothyroidism
Z90	7.354	Acquired absence of organs not elsewhere classified
Z43	7.193	Encounter for attention to artificial openings

Tab 2

Odds-Ratios

TRANSATLANTIC COHORT							
Exclusion	Drug	#Drug / Diagnosis	OR	Conf-low	Conf-up	P-value	
cancer	atorvastatin	1235	0,408	0,380	0,430	1,35005195E-155	
cancer	Diabetes	atorvastatin	517	0,490	0,450	5,31000000E-48	
cancer	Hypercholesterolemia	atorvastatin	407	0,699	0,630	1,41000000E-10	
cancer	Ischemic Heart	atorvastatin	452	0,490	0,440	1,16000000E-38	
cancer	Stroke	atorvastatin	642	0,583	0,540	5,18000000E-39	
cancer	Hypertension	atorvastatin	268	0,615	0,540	9,36000000E-13	
cancer	CKD	atorvastatin	601	0,467	0,430	1,85000000E-61	
cancer	Obesity	atorvastatin	664	0,469	0,430	8,24000000E-61	
1:1 MATCHED COHORT							
Exclusion	Drug	#Drug / Diagnosis	OR	Conf-low	Conf-up	P-value	
cancer	atorvastatin	1235	0,299	0,280	0,32	5,347E-250	
cancer	Diabetes	atorvastatin	517	0,335	0,302	8,180E-88	
cancer	Hypercholesterolemia	atorvastatin	407	0,465	0,413	2,230E-33	
cancer	Ischemic Heart	atorvastatin	452	0,340	0,305	2,910E-75	
cancer	Stroke	atorvastatin	642	0,406	0,370	3,000E-71	
cancer	Hypertension	atorvastatin	268	0,387	0,335	2,530E-35	
cancer	CKD	atorvastatin	601	0,328	0,299	4,180E-105	
cancer	Obesity	atorvastatin	664	0,335	0,306	1,480E-108	
ALL STATINS							
TRANSATLANTIC COHORT							
Exclusion	Drug	#Drug / Diagnosis	#Drug / Diagnosis	OR	Conf-low	Conf-up	P-value
	statins	53113	6426	0,72	0,70	0,74	2,1139E-108
Diabetes	statins	29834	3576	0,81	0,77	0,84	5,0600E-21
Hypercholesterolemia	statins	33198	3085	1,01	0,96	1,05	8,3842E-01
Ischemic Heart	statins	32766	3520	0,88	0,85	0,92	2,2900E-09
Stroke	statins	36205	5144	0,88	0,85	0,91	5,8400E-13
Hypertension	statins	22627	2014	0,94	0,89	0,99	2,2559E-02
CKD	statins	32982	4440	0,81	0,78	0,84	1,6800E-27
Obesity	statins	35169	5018	0,80	0,77	0,83	6,5700E-29
1:1 MATCHED COHORT							
Exclusion	Drug	#Diagnoses	#Drug / Diagnosis	OR	Conf-low	Conf-up	P-value
	statins	53113	6425	0,496	0,480	0,51	0,000E+00
Diabetes	statins	29834	3576	0,521	0,498	0,55	1,180E-154
Hypercholesterolemia	statins	33198	3085	0,643	0,613	0,68	1,210E-64
Ischemic Heart	statins	32766	3520	0,580	0,555	0,61	7,500E-111
Stroke	statins	36205	5144	0,584	0,562	0,61	1,300E-144
Hypertension	statins	22627	2014	0,546	0,515	0,58	3,990E-81
CKD	statins	32982	4440	0,530	0,508	0,55	3,150E-172
Obesity	statins	35169	5018	0,536	0,516	0,56	2,270E-195
SIMVASTATIN							
TRANSATLANTIC COHORT							
Exclusion	Drug	#Drug / Diagnosis	OR	Conf-low	Conf-up	P-value	
	simvastatin	4609	0,904	0,870	0,940	4,85394737E-07	
Diabetes	simvastatin	2814	0,897	0,860	0,940	2,30000000E-06	
Hypercholesterolemia	simvastatin	2452	1,081	1,030	1,130	1,02896600E-03	
Ischemic Heart	simvastatin	2845	1,007	0,960	1,050	7,72965426E-01	
Stroke	simvastatin	4179	0,946	0,910	0,980	3,37182700E-03	
Hypertension	simvastatin	1600	1,031	0,970	1,090	3,09244571E-01	
CKD	simvastatin	3575	0,915	0,880	0,950	6,98000000E-06	
Obesity	simvastatin	4038	0,898	0,870	0,930	6,40000000E-10	
1:1 MATCHED COHORT							
Exclusion	Drug	#Drug / Diagnosis	OR	Conf-low	Conf-up	P-value	
	simvastatin	4608	0,634	0,610	0,66	1,115E-100	
Diabetes	simvastatin	2814	0,590	0,561	0,62	6,360E-82	
Hypercholesterolemia	simvastatin	2452	0,699	0,662	0,74	6,840E-35	
Ischemic Heart	simvastatin	2845	0,666	0,633	0,70	5,710E-51	
Stroke	simvastatin	4179	0,640	0,613	0,67	9,330E-85	
Hypertension	simvastatin	1600	0,601	0,563	0,64	3,350E-47	
CKD	simvastatin	3575	0,615	0,588	0,64	1,590E-86	
Obesity	simvastatin	4038	0,615	0,589	0,64	5,480E-96	

Supplementary Table 2
Odds-Ratios

Supplementary Table 3

This table consists of the raw-results from the RWE study. Odds-Ratios and Relative-Risks are calculated for different statins and different cancer entities. Also both cohorts Charité and UVA were considered together in the transatlantic cohort as well as separately.

		Odds-Ratio Transatlatic-Cohort						
		No Exclusion						
Diagnose	Drug	NbDrug	NbDiag	NbDiag/Diag	Odds Ratio	Conf-Low	Conf-Up	P-Value
bladder	atorvastatin_charite_Cohort	6420	2100	75	0,936	0,740	1,180	0,59048164
brain	atorvastatin_charite_Cohort	6439	1296	20	0,393	0,250	0,610	0,00004770
breast	atorvastatin_charite_Cohort	6436	1373	19	0,351	0,220	0,550	0,00000957
cancer	atorvastatin_charite_Cohort	6302	36203	711	0,470	0,430	0,510	0,00000000
colorectal	atorvastatin_charite_Cohort	6424	3151	50	0,404	0,300	0,530	0,00000000
esophagus	atorvastatin_charite_Cohort	6431	678	7	0,262	0,120	0,550	0,00059876
kidney	atorvastatin_charite_Cohort	6431	1530	45	0,763	0,570	1,030	0,07273126
liver	atorvastatin_charite_Cohort	6432	1651	17	0,261	0,160	0,420	0,00000008
lung	atorvastatin_charite_Cohort	6421	2630	69	0,678	0,530	0,860	0,00170116
ovaries	atorvastatin_charite_Cohort	6438	828	6	0,183	0,080	0,410	0,00005408
pancreas	atorvastatin_charite_Cohort	6435	1464	28	0,489	0,340	0,710	0,00015610
prostate	atorvastatin_charite_Cohort	6428	2409	59	0,631	0,490	0,820	0,00048792
seq77	atorvastatin_charite_Cohort	6417	5507	73	0,336	0,270	0,420	0,00000000
seq78	atorvastatin_charite_Cohort	6411	7200	73	0,254	0,200	0,320	0,00000000
seq79	atorvastatin_charite_Cohort	6426	6024	81	0,340	0,270	0,420	0,00000000
stomach	atorvastatin_charite_Cohort	6438	1057	12	0,288	0,160	0,510	0,00003077
bladder	atorvastatin_transatlantic_Cohort	13837	2489	88	0,668	0,540	0,830	0,00025616
brain	atorvastatin_transatlantic_Cohort	13861	1815	31	0,315	0,220	0,450	0,00000000
breast	atorvastatin_transatlantic_Cohort	13854	2081	29	0,256	0,180	0,370	0,00000000
cancer	atorvastatin_transatlantic_Cohort	13460	52882	1235	0,408	0,380	0,430	0,00000000
colorectal	atorvastatin_transatlantic_Cohort	13841	3827	71	0,343	0,270	0,430	0,00000000
esophagus	atorvastatin_transatlantic_Cohort	13855	870	20	0,428	0,270	0,670	0,00027528
kidney	atorvastatin_transatlantic_Cohort	13853	1708	50	0,548	0,410	0,730	0,00005121
liver	atorvastatin_transatlantic_Cohort	13860	1822	22	0,222	0,150	0,340	0,00000000
lung	atorvastatin_transatlantic_Cohort	13822	3443	110	0,601	0,500	0,730	0,00000021
ovaries	atorvastatin_transatlantic_Cohort	13861	1219	11	0,165	0,090	0,300	0,00000001
pancreas	atorvastatin_transatlantic_Cohort	13855	1766	40	0,421	0,310	0,580	0,00000010
prostate	atorvastatin_transatlantic_Cohort	13837	3178	98	0,579	0,470	0,710	0,00000032
seq77	atorvastatin_transatlantic_Cohort	13815	6925	106	0,282	0,230	0,340	0,00000000
seq78	atorvastatin_transatlantic_Cohort	13795	9688	130	0,245	0,210	0,290	0,00000000
seq79	atorvastatin_transatlantic_Cohort	13816	8363	148	0,326	0,280	0,380	0,00000000
stomach	atorvastatin_transatlantic_Cohort	13863	1176	13	0,203	0,120	0,350	0,00000001
bladder	atorvastatin_UVA_Cohort	7417	389	13	0,421	0,240	0,730	0,00235283
brain	atorvastatin_UVA_Cohort	7422	519	11	0,263	0,140	0,480	0,0002599
breast	atorvastatin_UVA_Cohort	7418	708	10	0,174	0,090	0,320	0,00000011
cancer	atorvastatin_UVA_Cohort	7158	16679	524	0,379	0,350	0,420	0,00000000
colorectal	atorvastatin_UVA_Cohort	7417	676	21	0,390	0,250	0,600	0,00002990
esophagus	atorvastatin_UVA_Cohort	7424	192	13	0,885	0,500	1,550	0,00021530
kidney	atorvastatin_UVA_Cohort	7422	178	5	0,352	0,140	0,860	0,02395751
liver	atorvastatin_UVA_Cohort	7428	171	5	0,367	0,150	0,890	0,02710168
lung	atorvastatin_UVA_Cohort	7401	813	41	0,649	0,470	0,890	0,00795611
ovaries	atorvastatin_UVA_Cohort	7423	391	5	0,157	0,070	0,380	0,00002180
pancreas	atorvastatin_UVA_Cohort	7420	302	12	0,504	0,280	0,900	0,02126562
prostate	atorvastatin_UVA_Cohort	7409	769	39	0,651	0,470	0,900	0,00958381
seq77	atorvastatin_UVA_Cohort	7398	1418	33	0,290	0,210	0,410	0,00000000
seq78	atorvastatin_UVA_Cohort	7384	2488	57	0,285	0,220	0,370	0,00000000
seq79	atorvastatin_UVA_Cohort	7390	2339	67	0,359	0,280	0,460	0,00000000
stomach	atorvastatin_UVA_Cohort	7425	119	1	0,103	0,010	0,740	0,03811341
bladder	fluvastatin_charite_Cohort	860	2119	12	1,105	0,620	1,960	0,74682672
brain	fluvastatin_charite_Cohort	864	1295	0	0,000	0,000	nan	nan
breast	fluvastatin_charite_Cohort	861	1370	2	0,281	0,070	1,130	0,07323226
cancer	fluvastatin_charite_Cohort	810	36199	118	0,639	0,530	0,780	0,00000718
colorectal	fluvastatin_charite_Cohort	863	3162	8	0,486	0,240	0,980	0,04403310
esophagus	fluvastatin_charite_Cohort	862	683	5	1,425	0,590	3,440	0,43930927
kidney	fluvastatin_charite_Cohort	857	1535	14	1,800	1,060	3,060	0,02949958
liver	fluvastatin_charite_Cohort	863	1656	4	0,465	0,170	1,240	0,13103053
lung	fluvastatin_charite_Cohort	861	2633	8	0,586	0,290	1,180	0,13568962
ovaries	fluvastatin_charite_Cohort	864	825	1	0,233	0,030	1,650	0,15467825
pancreas	fluvastatin_charite_Cohort	864	1462	2	0,262	0,070	1,050	0,05213356
prostate	fluvastatin_charite_Cohort	860	2420	12	0,965	0,550	1,710	0,90975237
seq77	fluvastatin_charite_Cohort	854	5530	10	0,350	0,190	0,650	0,00086204
seq78	fluvastatin_charite_Cohort	856	7220	18	0,480	0,300	0,770	0,00232437
seq79	fluvastatin_charite_Cohort	855	6028	18	0,581	0,360	0,930	0,02470929
stomach	fluvastatin_charite_Cohort	862	1050	1	0,183	0,030	1,300	0,07696706
bladder	fluvastatin_transatlantic_Cohort	870	2517	12	1,446	0,820	2,560	0,20570011
brain	fluvastatin_transatlantic_Cohort	874	1813	0	0,000	0,000	nan	nan
breast	fluvastatin_transatlantic_Cohort	871	2087	2	0,286	0,070	1,150	0,07922611
cancer	fluvastatin_transatlantic_Cohort	820	53012	119	0,697	0,570	0,850	0,00042853
colorectal	fluvastatin_transatlantic_Cohort	873	3844	9	0,701	0,360	1,350	0,29606307
esophagus	fluvastatin_transatlantic_Cohort	872	878	5	1,719	0,710	4,150	0,23125908
kidney	fluvastatin_transatlantic_Cohort	867	1719	14	2,498	1,470	4,250	0,00076389
liver	fluvastatin_transatlantic_Cohort	873	1825	4	0,656	0,250	1,750	0,40292487

Supplementary Table 3
Odds-Ratio Transatlantic Cohort

lung	fluvastatin_transatlantic_Cohort	871	3469	8	0,692	0,340	1,390	0,30978342
ovaries	fluvastatin_transatlantic_Cohort	874	1219	1	0,244	0,030	1,740	0,17414313
pancreas	fluvastatin_transatlantic_Cohort	874	1769	2	0,337	0,080	1,350	0,13148467
prostate	fluvastatin_transatlantic_Cohort	870	3208	12	1,131	0,640	2,000	0,68504579
seq77	fluvastatin_transatlantic_Cohort	864	6974	10	0,432	0,230	0,810	0,00895920
seq78	fluvastatin_transatlantic_Cohort	866	9744	18	0,555	0,350	0,890	0,01333511
seq79	fluvastatin_transatlantic_Cohort	865	8395	18	0,649	0,410	1,030	0,06540593
stomach	fluvastatin_transatlantic_Cohort	872	1173	1	0,255	0,040	1,810	0,16060165
bladder	fluvastatin_UVA_Cohort	10	398	0	0,000	0,000	nan	nan
brain	fluvastatin_UVA_Cohort	10	518	0	0,000	0,000	nan	nan
breast	fluvastatin_UVA_Cohort	10	717	0	0,000	0,000	nan	nan
cancer	fluvastatin_UVA_Cohort	10	16813	1	0,542	0,070	4,280	0,57105569
colorectal	fluvastatin_UVA_Cohort	10	682	1	15,394	1,950	121,670	0,00950846
esophagus	fluvastatin_UVA_Cohort	10	195	0	0,000	0,000	nan	nan
kidney	fluvastatin_UVA_Cohort	10	184	0	0,000	0,000	nan	nan
liver	fluvastatin_UVA_Cohort	10	169	0	0,000	0,000	nan	nan
lung	fluvastatin_UVA_Cohort	10	836	0	0,000	0,000	nan	nan
ovaries	fluvastatin_UVA_Cohort	10	394	0	0,000	0,000	nan	nan
pancreas	fluvastatin_UVA_Cohort	10	307	0	0,000	0,000	nan	nan
prostate	fluvastatin_UVA_Cohort	10	788	0	0,000	0,000	nan	nan
seq77	fluvastatin_UVA_Cohort	10	1444	0	0,000	0,000	nan	nan
seq78	fluvastatin_UVA_Cohort	10	2524	0	0,000	0,000	nan	nan
seq79	fluvastatin_UVA_Cohort	10	2367	0	0,000	0,000	nan	nan
stomach	fluvastatin_UVA_Cohort	10	123	0	0,000	0,000	nan	nan
bladder	pravastatin_charite_Cohort	947	2121	18	1,514	0,950	2,420	0,08172758
brain	pravastatin_charite_Cohort	950	1294	2	0,269	0,070	1,080	0,05955814
breast	pravastatin_charite_Cohort	951	1373	5	0,636	0,260	1,540	0,32342309
cancer	pravastatin_charite_Cohort	914	36213	178	0,907	0,770	1,070	0,24746687
colorectal	pravastatin_charite_Cohort	948	3157	6	0,331	0,150	0,740	0,00663901
esophagus	pravastatin_charite_Cohort	950	684	3	0,769	0,250	2,400	0,66191383
kidney	pravastatin_charite_Cohort	946	1535	11	1,271	0,700	2,310	0,43937018
liver	pravastatin_charite_Cohort	949	1652	1	0,105	0,010	0,750	0,04039165
lung	pravastatin_charite_Cohort	948	2631	12	0,802	0,450	1,420	0,46055382
ovaries	pravastatin_charite_Cohort	950	826	6	1,280	0,570	2,870	0,56083404
pancreas	pravastatin_charite_Cohort	949	1462	5	0,599	0,250	1,440	0,25402749
prostate	pravastatin_charite_Cohort	949	2423	23	1,697	1,120	2,570	0,1250808
seq77	pravastatin_charite_Cohort	938	5526	24	0,777	0,520	1,170	0,22461704
seq78	pravastatin_charite_Cohort	943	7215	22	0,534	0,350	0,820	0,00391808
seq79	pravastatin_charite_Cohort	938	6023	18	0,528	0,330	0,840	0,00738607
stomach	pravastatin_charite_Cohort	952	1053	4	0,664	0,250	1,780	0,42126692
bladder	pravastatin_transatlantic_Cohort	2590	2516	18	0,723	0,450	1,150	0,17631538
brain	pravastatin_transatlantic_Cohort	2596	1812	6	0,331	0,150	0,740	0,00663901
breast	pravastatin_transatlantic_Cohort	2597	2092	9	0,431	0,220	0,830	0,01290727
cancer	pravastatin_transatlantic_Cohort	2485	52982	331	0,631	0,560	0,710	0,00000000
colorectal	pravastatin_transatlantic_Cohort	2591	3836	12	0,313	0,180	0,550	0,00005350
esophagus	pravastatin_transatlantic_Cohort	2596	879	4	0,457	0,170	1,220	0,11931669
kidney	pravastatin_transatlantic_Cohort	2590	1717	13	0,765	0,440	1,320	0,34459225
liver	pravastatin_transatlantic_Cohort	2595	1821	2	0,110	0,030	0,440	0,00132245
lung	pravastatin_transatlantic_Cohort	2584	3459	25	0,732	0,490	1,090	0,12618731
ovaries	pravastatin_transatlantic_Cohort	2595	1219	9	0,744	0,390	1,440	0,38140528
pancreas	pravastatin_transatlantic_Cohort	2593	1768	8	0,454	0,230	0,910	0,02421165
prostate	pravastatin_transatlantic_Cohort	2590	3206	41	1,308	0,960	1,780	0,08794177
seq77	pravastatin_transatlantic_Cohort	2579	6965	27	0,390	0,270	0,570	0,00000113
seq78	pravastatin_transatlantic_Cohort	2570	9722	40	0,414	0,300	0,570	0,00000012
seq79	pravastatin_transatlantic_Cohort	2572	8379	35	0,421	0,300	0,590	0,00000079
stomach	pravastatin_transatlantic_Cohort	2597	1175	4	0,341	0,130	0,910	0,02995527
bladder	pravastatin_UVA_Cohort	1643	395	0	0,000	0,000	nan	nan
brain	pravastatin_UVA_Cohort	1646	518	4	0,443	0,170	1,190	0,10075232
breast	pravastatin_UVA_Cohort	1646	719	4	0,318	0,120	0,850	0,02162087
cancer	pravastatin_UVA_Cohort	1571	16769	153	0,527	0,450	0,620	0,00000000
colorectal	pravastatin_UVA_Cohort	1643	679	6	0,509	0,230	1,140	0,09792977
esophagus	pravastatin_UVA_Cohort	1646	195	1	0,294	0,040	2,100	0,22779074
kidney	pravastatin_UVA_Cohort	1644	182	2	0,634	0,160	2,560	0,53012127
liver	pravastatin_UVA_Cohort	1646	169	1	0,339	0,050	2,420	0,27783347
lung	pravastatin_UVA_Cohort	1636	828	13	0,918	0,530	1,590	0,77285988
ovaries	pravastatin_UVA_Cohort	1645	393	3	0,438	0,140	1,370	0,15651161
pancreas	pravastatin_UVA_Cohort	1644	306	3	0,565	0,180	1,760	0,33135588
prostate	pravastatin_UVA_Cohort	1641	783	18	1,352	0,850	2,160	0,20652570
seq77	pravastatin_UVA_Cohort	1641	1439	3	0,040	0,040	0,370	0,00019509
seq78	pravastatin_UVA_Cohort	1627	2507	18	0,415	0,260	0,660	0,00023613
seq79	pravastatin_UVA_Cohort	1634	2356	17	0,416	0,260	0,670	0,00030595
stomach	pravastatin_UVA_Cohort	1645	122	0	0,000	0,000	nan	nan
bladder	simvastatin_charite_Cohort	22402	2105	431	1,733	1,560	1,930	0,00000000
brain	simvastatin_charite_Cohort	22371	1300	104	0,579	0,470	0,710	0,00000032
breast	simvastatin_charite_Cohort	22372	1366	102	0,537	0,440	0,660	0,00000000

Supplementary Table 3
Odds-Ratio Transatlantic Cohort

cancer	simvastatin_charite_Cohort	22271	36506	4257	0,883	0,850	0,920	0,00000000
colorectal	simvastatin_charite_Cohort	22364	3139	319	0,755	0,670	0,850	0,00000485
esophagus	simvastatin_charite_Cohort	22370	704	98	1,080	0,870	1,340	0,49473117
kidney	simvastatin_charite_Cohort	22368	1523	243	1,272	1,110	1,460	0,00061550
liver	simvastatin_charite_Cohort	22358	1646	140	0,620	0,520	0,740	0,00000018
lung	simvastatin_charite_Cohort	22366	2646	415	1,249	1,120	1,390	0,00006325
ovaries	simvastatin_charite_Cohort	22375	833	53	0,452	0,340	0,600	0,00000007
pancreas	simvastatin_charite_Cohort	22372	1473	155	0,784	0,660	0,930	0,00544550
prostate	simvastatin_charite_Cohort	22393	2401	389	1,297	1,160	1,450	0,00000641
seq77	simvastatin_charite_Cohort	22319	5374	510	0,703	0,640	0,770	0,00000000
seq78	simvastatin_charite_Cohort	22300	7141	576	0,583	0,530	0,640	0,00000000
seq79	simvastatin_charite_Cohort	22348	5961	540	0,665	0,610	0,730	0,00000000
stomach	simvastatin_charite_Cohort	22363	1059	116	0,821	0,680	1,000	0,04463320
bladder	simvastatin_transatlantic_Cohort	25863	2502	437	1,999	1,800	2,220	0,00000000
brain	simvastatin_transatlantic_Cohort	25830	1818	113	0,620	0,510	0,750	0,00000167
breast	simvastatin_transatlantic_Cohort	25827	2077	116	0,553	0,460	0,670	0,00000000
cancer	simvastatin_transatlantic_Cohort	25608	53267	4609	0,904	0,870	0,940	0,00000049
colorectal	simvastatin_transatlantic_Cohort	25818	3816	329	0,886	0,790	0,990	0,03521205
esophagus	simvastatin_transatlantic_Cohort	25830	897	104	1,230	1,000	1,510	0,04856346
kidney	simvastatin_transatlantic_Cohort	25828	1705	243	1,564	1,360	1,790	0,00000000
liver	simvastatin_transatlantic_Cohort	25819	1814	141	0,790	0,660	0,940	0,00897216
lung	simvastatin_transatlantic_Cohort	25816	3472	442	1,377	1,250	1,520	0,00000000
ovaries	simvastatin_transatlantic_Cohort	25834	1224	59	0,473	0,360	0,620	0,00000011
pancreas	simvastatin_transatlantic_Cohort	25831	1778	161	0,933	0,790	1,100	0,41921404
prostate	simvastatin_transatlantic_Cohort	25848	3183	408	1,385	1,250	1,540	0,00000000
seq77	simvastatin_transatlantic_Cohort	25766	6807	534	0,803	0,730	0,880	0,00000550
seq78	simvastatin_transatlantic_Cohort	25735	9644	616	0,640	0,590	0,690	0,00000000
seq79	simvastatin_transatlantic_Cohort	25791	8314	579	0,703	0,650	0,770	0,00000000
stomach	simvastatin_transatlantic_Cohort	25824	1182	119	1,050	0,870	1,270	0,62571728
bladder	simvastatin_UVA_Cohort	3461	397	6	0,410	0,180	0,920	0,03189318
brain	simvastatin_UVA_Cohort	3459	518	9	0,473	0,240	0,910	0,02743977
breast	simvastatin_UVA_Cohort	3455	711	14	0,538	0,320	0,910	0,01992112
cancer	simvastatin_UVA_Cohort	3337	16761	352	0,575	0,510	0,640	0,00000000
colorectal	simvastatin_UVA_Cohort	3454	677	10	0,401	0,210	0,750	0,00493380
esophagus	simvastatin_UVA_Cohort	3460	193	6	0,859	0,380	1,940	0,72790244
kidney	simvastatin_UVA_Cohort	3460	182	0	0,000	0,000	nan	nan
liver	simvastatin_UVA_Cohort	3461	168	1	0,160	0,020	1,140	0,07522293
lung	simvastatin_UVA_Cohort	3450	826	27	0,910	0,620	1,340	0,64425689
ovaries	simvastatin_UVA_Cohort	3459	391	6	0,417	0,190	0,930	0,03059616
pancreas	simvastatin_UVA_Cohort	3459	305	6	0,537	0,240	1,210	0,13206059
prostate	simvastatin_UVA_Cohort	3455	782	19	0,667	0,420	1,050	0,08284148
seq77	simvastatin_UVA_Cohort	3447	1433	24	0,457	0,310	0,690	0,00014003
seq78	simvastatin_UVA_Cohort	3435	2503	40	0,436	0,320	0,600	0,00000035
seq79	simvastatin_UVA_Cohort	3443	2353	39	0,452	0,330	0,620	0,00000115
stomach	simvastatin_UVA_Cohort	3461	123	3	0,669	0,210	2,100	0,50384829
bladder	statins_charite_Cohort	30087	2094	548	1,699	1,540	1,870	0,00000000
brain	statins_charite_Cohort	30054	1302	127	0,512	0,430	0,620	0,00000000
breast	statins_charite_Cohort	30048	1363	128	0,491	0,410	0,590	0,00000000
cancer	statins_charite_Cohort	29926	36497	5295	0,790	0,760	0,820	0,00000000
cancersubentities	statins_charite_Cohort	30019	21890	3344	0,858	0,830	0,890	0,00000000
colorectal	statins_charite_Cohort	30046	3133	386	0,667	0,600	0,740	0,00000000
esophagus	statins_charite_Cohort	30045	699	115	0,937	0,770	1,140	0,52629374
kidney	statins_charite_Cohort	30047	1513	318	1,270	1,120	1,440	0,00021278
liver	statins_charite_Cohort	30037	1645	162	0,518	0,440	0,610	0,00000000
lung	statins_charite_Cohort	30038	2651	507	1,129	1,020	1,250	0,01919563
ovaries	statins_charite_Cohort	30054	837	66	0,406	0,320	0,520	0,00000000
pancreas	statins_charite_Cohort	30053	1476	191	0,706	0,610	0,820	0,00000524
prostate	statins_charite_Cohort	30077	2395	492	1,235	1,120	1,360	0,00002466
seq77	statins_charite_Cohort	29994	5354	628	0,633	0,580	0,690	0,00000000
seq78	statins_charite_Cohort	29966	7125	699	0,513	0,470	0,560	0,00000000
seq79	statins_charite_Cohort	30040	5955	671	0,602	0,550	0,650	0,00000000
stomach	statins_charite_Cohort	30041	1058	132	0,677	0,560	0,810	0,00004065
bladder	statins_transatlantic_Cohort	43286	2480	570	1,593	1,450	1,750	0,00000000
brain	statins_transatlantic_Cohort	43261	1823	154	0,488	0,410	0,580	0,00000000
breast	statins_transatlantic_Cohort	43242	2067	158	0,438	0,370	0,520	0,00000000
cancer	statins_transatlantic_Cohort	42682	53113	6426	0,716	0,700	0,740	0,00000000
colorectal	statins_transatlantic_Cohort	43238	3803	425	0,667	0,600	0,740	0,00000000
esophagus	statins_transatlantic_Cohort	43249	886	140	0,997	0,830	1,190	0,97639961
kidney	statins_transatlantic_Cohort	43251	1688	325	1,269	1,120	1,430	0,00014824
liver	statins_transatlantic_Cohort	43248	1814	169	0,544	0,460	0,640	0,00000000
lung	statins_transatlantic_Cohort	43207	3449	596	1,114	1,020	1,220	0,01797753
ovaries	statins_transatlantic_Cohort	43257	1225	82	0,380	0,300	0,480	0,00000000
pancreas	statins_transatlantic_Cohort	43252	1774	213	0,724	0,630	0,840	0,00001351
prostate	statins_transatlantic_Cohort	43258	3151	573	1,184	1,080	1,300	0,00038379
seq77	statins_transatlantic_Cohort	43154	6753	693	0,608	0,560	0,660	0,00000000

Supplementary Table 3
Odds-Ratio Transatlantic Cohort

seq78	statins_transatlantic_Cohort	43085	9574	827	0,499	0,460	0,540	0,00000000
seq79	statins_transatlantic_Cohort	43178	8269	808	0,574	0,530	0,620	0,00000000
stomach	statins_transatlantic_Cohort	43249	1176	136	0,694	0,580	0,830	0,00007447
bladder	statins_UVA_Cohort	13199	386	22	0,396	0,260	0,610	0,00002502
brain	statins_UVA_Cohort	13207	521	27	0,358	0,240	0,530	0,00000056
breast	statins_UVA_Cohort	13194	704	30	0,291	0,200	0,420	0,00000000
cancer	statins_UVA_Cohort	12756	16616	1131	0,465	0,440	0,500	0,00000000
cancersubentities	statins_UVA_Cohort	12987	7514	501	0,465	0,420	0,510	0,00000000
colorectal	statins_UVA_Cohort	13192	670	39	0,405	0,290	0,560	0,00000012
esophagus	statins_UVA_Cohort	13204	187	25	1,013	0,660	1,540	0,95663733
kidney	statins_UVA_Cohort	13204	175	7	0,273	0,130	0,580	0,00070476
liver	statins_UVA_Cohort	13211	169	7	0,283	0,130	0,600	0,00126272
lung	statins_UVA_Cohort	13169	798	89	0,827	0,660	1,030	0,09405443
ovaries	statins_UVA_Cohort	13203	388	16	0,282	0,170	0,460	0,00000091
pancreas	statins_UVA_Cohort	13199	298	22	0,523	0,340	0,810	0,00347392
prostate	statins_UVA_Cohort	13181	756	81	0,789	0,630	0,990	0,03951355
seq77	statins_UVA_Cohort	13160	1399	65	0,320	0,250	0,410	0,00000000
seq78	statins_UVA_Cohort	13119	2449	128	0,361	0,300	0,430	0,00000000
seq79	statins_UVA_Cohort	13138	2314	137	0,413	0,350	0,490	0,00000000
stomach	statins_UVA_Cohort	13208	118	4	0,230	0,080	0,620	0,00494112

Relative Risk Transatlatic-Cohort						
No Exclusion						
Diagnose	Drug	RelativRisk	Conf-Upper	Conf-Lower	P-Value	NNT-ALL
bladder	atorvastatin_charite_Cohort	1,214	1,483	0,994	0,05703431	-370,59
brain	atorvastatin_charite_Cohort	0,413	0,636	0,269	0,00006509	216,13
breast	atorvastatin_charite_Cohort	0,371	0,576	0,239	0,00001249	189,85
cancer	atorvastatin_charite_Cohort	0,584	0,623	0,548	0,00000000	11,27
colorectal	atorvastatin_charite_Cohort	0,510	0,654	0,398	0,00000017	106,70
esophagus	atorvastatin_charite_Cohort	0,373	0,695	0,200	0,00196494	382,57
kidney	atorvastatin_charite_Cohort	0,858	1,134	0,650	0,28435302	766,64
liver	atorvastatin_charite_Cohort	0,353	0,533	0,234	0,00000103	153,13
lung	atorvastatin_charite_Cohort	0,791	0,986	0,635	0,03643041	301,53
ovaries	atorvastatin_charite_Cohort	0,184	0,410	0,082	0,00004414	241,38
pancreas	atorvastatin_charite_Cohort	0,559	0,792	0,394	0,00113989	254,64
prostate	atorvastatin_charite_Cohort	0,765	0,966	0,606	0,02412469	291,94
seq77	atorvastatin_charite_Cohort	0,469	0,570	0,386	0,00000000	56,66
seq78	atorvastatin_charite_Cohort	0,348	0,424	0,285	0,00000000	35,15
seq79	atorvastatin_charite_Cohort	0,417	0,508	0,342	0,00000000	47,15
stomach	atorvastatin_charite_Cohort	0,336	0,570	0,199	0,00005661	233,27
bladder	atorvastatin_transatlantic_Cohort	0,860	1,036	0,714	0,11211379	740,91
brain	atorvastatin_transatlantic_Cohort	0,356	0,497	0,255	0,00000000	218,77
breast	atorvastatin_transatlantic_Cohort	0,291	0,411	0,207	0,00000000	172,77
cancer	atorvastatin_transatlantic_Cohort	0,512	0,539	0,488	0,00000000	10,30
colorectal	atorvastatin_transatlantic_Cohort	0,411	0,509	0,331	0,00000000	113,77
esophagus	atorvastatin_transatlantic_Cohort	0,552	0,815	0,374	0,00283963	656,42
kidney	atorvastatin_transatlantic_Cohort	0,620	0,807	0,477	0,00039409	397,41
liver	atorvastatin_transatlantic_Cohort	0,283	0,410	0,195	0,00000000	195,24
lung	atorvastatin_transatlantic_Cohort	0,702	0,835	0,589	0,00008113	251,18
ovaries	atorvastatin_transatlantic_Cohort	0,210	0,356	0,124	0,00000001	263,58
pancreas	atorvastatin_transatlantic_Cohort	0,471	0,633	0,350	0,00000093	274,26
prostate	atorvastatin_transatlantic_Cohort	0,666	0,802	0,553	0,00002221	243,18
seq77	atorvastatin_transatlantic_Cohort	0,374	0,442	0,317	0,00000000	59,35
seq78	atorvastatin_transatlantic_Cohort	0,315	0,367	0,270	0,00000000	38,74
seq79	atorvastatin_transatlantic_Cohort	0,395	0,458	0,341	0,00000000	51,07
stomach	atorvastatin_transatlantic_Cohort	0,234	0,389	0,140	0,00000005	281,88
bladder	atorvastatin_UVA_Cohort	0,504	0,831	0,306	0,00719458	471,88
brain	atorvastatin_UVA_Cohort	0,332	0,564	0,195	0,00005501	263,73
breast	atorvastatin_UVA_Cohort	0,224	0,388	0,130	0,00000014	165,29
cancer	atorvastatin_UVA_Cohort	0,473	0,510	0,438	0,00000000	10,89
colorectal	atorvastatin_UVA_Cohort	0,403	0,616	0,264	0,00003139	228,21
esophagus	atorvastatin_UVA_Cohort	1,065	1,774	0,639	0,82064966	-7606,49
kidney	atorvastatin_UVA_Cohort	0,408	0,921	0,181	0,03051310	854,36
liver	atorvastatin_UVA_Cohort	0,367	0,893	0,151	0,02682121	861,19
lung	atorvastatin_UVA_Cohort	0,748	0,997	0,561	0,04740669	449,94
ovaries	atorvastatin_UVA_Cohort	0,249	0,502	0,124	0,00011013	308,51
pancreas	atorvastatin_UVA_Cohort	0,530	0,923	0,304	0,02483085	644,38
prostate	atorvastatin_UVA_Cohort	0,682	0,930	0,500	0,01553347	379,75
seq77	atorvastatin_UVA_Cohort	0,339	0,465	0,246	0,00000000	97,52
seq78	atorvastatin_UVA_Cohort	0,333	0,424	0,262	0,00000000	55,45
seq79	atorvastatin_UVA_Cohort	0,426	0,532	0,341	0,00000000	68,96
stomach	atorvastatin_UVA_Cohort	0,100	0,714	0,014	0,02152952	823,82
bladder	fluvastatin_charite_Cohort	1,278	2,153	0,759	0,36236861	-283,95
brain	fluvastatin_charite_Cohort	0,000	nan	0,000	nan	128,39
breast	fluvastatin_charite_Cohort	0,421	1,303	0,136	0,13359885	209,36
cancer	fluvastatin_charite_Cohort	0,768	0,892	0,661	0,00059087	20,41
colorectal	fluvastatin_charite_Cohort	0,611	1,132	0,329	0,11804412	135,82
esophagus	fluvastatin_charite_Cohort	1,698	3,781	0,762	0,19643705	-350,44
kidney	fluvastatin_charite_Cohort	2,253	3,569	1,423	0,00056940	-86,80
liver	fluvastatin_charite_Cohort	0,583	1,399	0,243	0,22906666	241,57
lung	fluvastatin_charite_Cohort	0,661	1,267	0,344	0,21501041	186,81
ovaries	fluvastatin_charite_Cohort	0,233	1,657	0,033	0,14516057	263,10
pancreas	fluvastatin_charite_Cohort	0,263	1,052	0,066	0,05823774	154,52
prostate	fluvastatin_charite_Cohort	1,194	1,975	0,722	0,49972728	-356,47
seq77	fluvastatin_charite_Cohort	0,597	0,957	0,373	0,03159736	75,69
seq78	fluvastatin_charite_Cohort	0,619	0,927	0,413	0,01989581	61,27
seq79	fluvastatin_charite_Cohort	0,832	1,216	0,569	0,34797885	166,13
stomach	fluvastatin_charite_Cohort	0,365	1,461	0,091	0,15520242	249,09
bladder	fluvastatin_transatlantic_Cohort	1,668	2,809	0,991	0,05384184	-156,01
brain	fluvastatin_transatlantic_Cohort	0,000	nan	0,000	nan	144,04
breast	fluvastatin_transatlantic_Cohort	0,429	1,329	0,139	0,14202220	219,14

Supplementary Table 3
Relative Risk Transatlatic-Cohort

cancer	fluvastatin_transatlantic_Cohort	0,822	0,955	0,708	0,01022126	28,73
colorectal	fluvastatin_transatlantic_Cohort	0,857	1,543	0,476	0,61949024	477,96
esophagus	fluvastatin_transatlantic_Cohort	2,048	4,559	0,920	0,07876138	-284,63
kidney	fluvastatin_transatlantic_Cohort	3,121	4,943	1,971	0,00000172	-71,85
liver	fluvastatin_transatlantic_Cohort	0,821	1,969	0,342	0,67176963	799,50
lung	fluvastatin_transatlantic_Cohort	0,778	1,492	0,406	0,45845290	340,63
ovaries	fluvastatin_transatlantic_Cohort	0,245	1,740	0,035	0,15870180	283,97
pancreas	fluvastatin_transatlantic_Cohort	0,338	1,349	0,085	0,12405946	223,08
prostate	fluvastatin_transatlantic_Cohort	1,397	2,309	0,845	0,19361733	-206,09
seq77	fluvastatin_transatlantic_Cohort	0,733	1,175	0,458	0,19762215	141,96
seq78	fluvastatin_transatlantic_Cohort	0,711	1,065	0,475	0,09748340	94,03
seq79	fluvastatin_transatlantic_Cohort	0,926	1,353	0,634	0,70411316	425,08
stomach	fluvastatin_transatlantic_Cohort	0,508	2,030	0,127	0,34351616	451,50
bladder	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	238,43
brain	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	183,16
breast	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	132,46
cancer	fluvastatin_UVA_Cohort	0,588	3,775	0,092	0,58714551	14,27
colorectal	fluvastatin_UVA_Cohort	13,954	89,721	2,170	0,00553781	-10,77
esophagus	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	486,35
kidney	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	515,34
liver	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	561,09
lung	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	113,78
ovaries	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	240,78
pancreas	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	309,08
prostate	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	120,54
seq77	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	66,13
seq78	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	37,96
seq79	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	40,44
stomach	fluvastatin_UVA_Cohort	0,000	nan	0,000	nan	770,74
bladder	pravastatin_charite_Cohort	1,820	2,757	1,202	0,00473027	-96,32
brain	pravastatin_charite_Cohort	0,405	1,256	0,131	0,11696336	215,91
breast	pravastatin_charite_Cohort	0,637	1,530	0,265	0,31793719	334,39
cancer	pravastatin_charite_Cohort	1,003	1,132	0,889	0,96481003	-1609,80
colorectal	pravastatin_charite_Cohort	0,499	0,957	0,260	0,03621028	105,48
esophagus	pravastatin_charite_Cohort	1,023	2,727	0,384	0,96709036	-10659,15
kidney	pravastatin_charite_Cohort	1,484	2,552	0,863	0,15402595	-224,75
liver	pravastatin_charite_Cohort	0,212	0,846	0,053	0,02793005	127,62
lung	pravastatin_charite_Cohort	1,000	1,655	0,605	1,00000000	-261485,81
ovaries	pravastatin_charite_Cohort	1,487	3,120	0,708	0,29839227	-415,50
pancreas	pravastatin_charite_Cohort	0,837	1,754	0,399	0,65047280	698,67
prostate	pravastatin_charite_Cohort	1,740	2,588	1,170	0,00626592	-93,58
seq77	pravastatin_charite_Cohort	1,078	1,501	0,774	0,66969261	-392,78
seq78	pravastatin_charite_Cohort	0,752	1,064	0,531	0,10778773	94,09
seq79	pravastatin_charite_Cohort	0,871	1,239	0,612	0,45136894	216,04
stomach	pravastatin_charite_Cohort	0,665	1,772	0,250	0,42193321	472,88
bladder	pravastatin_transatlantic_Cohort	0,880	1,336	0,579	0,56038745	863,95
brain	pravastatin_transatlantic_Cohort	0,388	0,814	0,185	0,01220153	235,07
breast	pravastatin_transatlantic_Cohort	0,432	0,831	0,225	0,01175244	219,94
cancer	pravastatin_transatlantic_Cohort	0,738	0,810	0,672	0,00000000	19,46
colorectal	pravastatin_transatlantic_Cohort	0,419	0,684	0,257	0,00052834	117,27
esophagus	pravastatin_transatlantic_Cohort	0,571	1,374	0,237	0,21307558	691,86
kidney	pravastatin_transatlantic_Cohort	0,878	1,457	0,529	0,62727756	1247,07
liver	pravastatin_transatlantic_Cohort	0,165	0,512	0,053	0,00189729	171,16
lung	pravastatin_transatlantic_Cohort	0,844	1,214	0,587	0,36631804	484,91
ovaries	pravastatin_transatlantic_Cohort	0,908	1,642	0,502	0,76239373	2333,24
pancreas	pravastatin_transatlantic_Cohort	0,624	1,128	0,346	0,11769346	392,79
prostate	pravastatin_transatlantic_Cohort	1,357	1,829	1,007	0,04458478	-230,04
seq77	pravastatin_transatlantic_Cohort	0,550	0,755	0,401	0,00023361	84,04
seq78	pravastatin_transatlantic_Cohort	0,561	0,730	0,430	0,00002267	61,62
seq79	pravastatin_transatlantic_Cohort	0,614	0,806	0,467	0,00049152	81,38
stomach	pravastatin_transatlantic_Cohort	0,342	0,911	0,128	0,03183007	336,85
bladder	pravastatin_UVA_Cohort	0,000	nan	0,000	nan	236,13
brain	pravastatin_UVA_Cohort	0,444	1,187	0,166	0,10550477	328,90
breast	pravastatin_UVA_Cohort	0,320	0,853	0,120	0,02258455	193,37
cancer	pravastatin_UVA_Cohort	0,614	0,708	0,533	0,00000000	15,21
colorectal	pravastatin_UVA_Cohort	0,592	1,245	0,282	0,16713509	341,28
esophagus	pravastatin_UVA_Cohort	0,294	2,097	0,041	0,22464588	685,88
kidney	pravastatin_UVA_Cohort	0,627	2,524	0,156	0,52148463	1383,09
liver	pravastatin_UVA_Cohort	0,340	2,424	0,048	0,28456782	846,44
lung	pravastatin_UVA_Cohort	0,971	1,643	0,574	0,91974648	3959,03

Supplementary Table 3
Relative Risk Transatlantic-Cohort

ovaries	pravastatin_UVA_Cohort	0,584	1,562	0,218	0,28807270	577,54
pancreas	pravastatin_UVA_Cohort	0,749	2,005	0,280	0,57670855	1226,12
prostate	pravastatin_UVA_Cohort	1,410	2,217	0,897	0,13683920	-297,87
seq77	pravastatin_UVA_Cohort	0,159	0,425	0,060	0,00025366	78,07
seq78	pravastatin_UVA_Cohort	0,528	0,794	0,351	0,00221617	80,22
seq79	pravastatin_UVA_Cohort	0,514	0,787	0,335	0,00230768	82,95
stomach	pravastatin_UVA_Cohort	0,000	nan	0,000	nan	763,31
bladder	simvastatin_charite_Cohort	1,867	2,063	1,690	0,00000000	-99,74
brain	simvastatin_charite_Cohort	0,610	0,742	0,502	0,00000103	318,82
breast	simvastatin_charite_Cohort	0,586	0,710	0,483	0,00000009	282,78
cancer	simvastatin_charite_Cohort	0,945	0,972	0,920	0,00006386	86,05
colorectal	simvastatin_charite_Cohort	0,847	0,944	0,760	0,00273241	341,15
esophagus	simvastatin_charite_Cohort	1,152	1,413	0,938	0,17673512	-1589,86
kidney	simvastatin_charite_Cohort	1,371	1,563	1,203	0,00000313	-309,11
liver	simvastatin_charite_Cohort	0,714	0,839	0,609	0,00004441	341,90
lung	simvastatin_charite_Cohort	1,371	1,513	1,243	0,00000000	-176,63
ovaries	simvastatin_charite_Cohort	0,494	0,644	0,378	0,00000033	376,29
pancreas	simvastatin_charite_Cohort	0,854	1,001	0,729	0,05065770	766,44
prostate	simvastatin_charite_Cohort	1,391	1,543	1,255	0,00000000	-186,72
seq77	simvastatin_charite_Cohort	0,879	0,952	0,812	0,00153184	249,95
seq78	simvastatin_charite_Cohort	0,704	0,760	0,652	0,00000000	76,19
seq79	simvastatin_charite_Cohort	0,785	0,851	0,725	0,00000001	127,11
stomach	simvastatin_charite_Cohort	0,891	1,071	0,741	0,22131364	1428,70
bladder	simvastatin_transatlantic_Cohort	2,163	2,383	1,962	0,00000000	-98,22
brain	simvastatin_transatlantic_Cohort	0,656	0,788	0,545	0,00000943	409,97
breast	simvastatin_transatlantic_Cohort	0,610	0,729	0,511	0,00000008	314,16
cancer	simvastatin_transatlantic_Cohort	0,964	0,989	0,939	0,00563314	141,33
colorectal	simvastatin_transatlantic_Cohort	0,994	1,104	0,894	0,91817358	10726,16
esophagus	simvastatin_transatlantic_Cohort	1,308	1,592	1,075	0,00736776	-971,41
kidney	simvastatin_transatlantic_Cohort	1,686	1,919	1,481	0,00000000	-237,41
liver	simvastatin_transatlantic_Cohort	0,909	1,065	0,775	0,24182110	1558,71
lung	simvastatin_transatlantic_Cohort	1,509	1,658	1,374	0,00000000	-154,04
ovaries	simvastatin_transatlantic_Cohort	0,519	0,667	0,405	0,00000040	429,90
pancreas	simvastatin_transatlantic_Cohort	1,018	1,188	0,873	0,83178373	-8103,83
prostate	simvastatin_transatlantic_Cohort	1,493	1,648	1,353	0,00000000	-174,64
seq77	simvastatin_transatlantic_Cohort	1,001	1,082	0,926	0,98185959	-42610,79
seq78	simvastatin_transatlantic_Cohort	0,769	0,828	0,715	0,00000000	115,64
seq79	simvastatin_transatlantic_Cohort	0,829	0,895	0,768	0,00000216	181,96
stomach	simvastatin_transatlantic_Cohort	1,136	1,361	0,948	0,16764596	-1648,44
bladder	simvastatin_UVA_Cohort	0,409	0,915	0,183	0,02919174	399,30
brain	simvastatin_UVA_Cohort	0,523	0,977	0,280	0,04169286	379,88
breast	simvastatin_UVA_Cohort	0,647	1,046	0,401	0,07466324	373,91
cancer	simvastatin_UVA_Cohort	0,661	0,726	0,602	0,00000000	17,25
colorectal	simvastatin_UVA_Cohort	0,477	0,843	0,270	0,01078588	263,11
esophagus	simvastatin_UVA_Cohort	0,850	1,915	0,377	0,70823554	3270,60
kidney	simvastatin_UVA_Cohort	0,000	nan	0,000	nan	503,72
liver	simvastatin_UVA_Cohort	0,159	1,138	0,022	0,06734275	656,30
lung	simvastatin_UVA_Cohort	0,961	1,390	0,665	0,84344230	2953,97
ovaries	simvastatin_UVA_Cohort	0,483	1,020	0,229	0,05578103	462,48
pancreas	simvastatin_UVA_Cohort	0,623	1,317	0,295	0,21687114	816,98
prostate	simvastatin_UVA_Cohort	0,766	1,169	0,502	0,21825597	515,81
seq77	simvastatin_UVA_Cohort	0,530	0,769	0,365	0,00088099	139,39
seq78	simvastatin_UVA_Cohort	0,521	0,692	0,393	0,0000806	78,65
seq79	simvastatin_UVA_Cohort	0,521	0,698	0,388	0,00001667	83,65
stomach	simvastatin_UVA_Cohort	0,664	2,085	0,211	0,49327588	2276,65
bladder	statins_charite_Cohort	1,831	2,007	1,671	0,00000000	-106,95
brain	statins_charite_Cohort	0,542	0,648	0,454	0,00000000	264,09
breast	statins_charite_Cohort	0,531	0,632	0,446	0,00000000	243,35
cancer	statins_charite_Cohort	0,865	0,887	0,844	0,00000000	34,35
cancersubentities	statins_charite_Cohort	0,935	0,967	0,905	0,00008053	119,31
colorectal	statins_charite_Cohort	0,753	0,832	0,682	0,00000004	207,62
esophagus	statins_charite_Cohort	1,019	1,232	0,843	0,85628348	-12446,38
kidney	statins_charite_Cohort	1,343	1,512	1,194	0,00000140	-337,59
liver	statins_charite_Cohort	0,601	0,698	0,517	0,00000000	237,39
lung	statins_charite_Cohort	1,235	1,352	1,127	0,00000711	-276,08
ovaries	statins_charite_Cohort	0,441	0,562	0,347	0,00000000	329,88
pancreas	statins_charite_Cohort	0,764	0,883	0,660	0,00031403	462,77
prostate	statins_charite_Cohort	1,329	1,460	1,210	0,00000001	-223,03
seq77	statins_charite_Cohort	0,787	0,847	0,731	0,00000000	138,90
seq78	statins_charite_Cohort	0,622	0,667	0,580	0,00000000	58,12

Supplementary Table 3
Relative Risk Transatlantic-Cohort

seq79	statins_charite_Cohort	0,708	0,762	0,658	0,00000000	91,47
stomach	statins_charite_Cohort	0,743	0,884	0,625	0,00082432	588,99
bladder	statins_transatlantic_Cohort	1,723	1,882	1,578	0,00000000	-159,82
brain	statins_transatlantic_Cohort	0,517	0,607	0,441	0,00000000	282,62
breast	statins_transatlantic_Cohort	0,480	0,560	0,412	0,00000000	227,40
cancer	statins_transatlantic_Cohort	0,797	0,815	0,779	0,00000000	24,53
colorectal	statins_transatlantic_Cohort	0,748	0,822	0,681	0,00000000	264,99
esophagus	statins_transatlantic_Cohort	1,085	1,287	0,914	0,35586601	-3505,14
kidney	statins_transatlantic_Cohort	1,346	1,511	1,199	0,00000071	-470,86
liver	statins_transatlantic_Cohort	0,627	0,726	0,541	0,00000000	367,74
lung	statins_transatlantic_Cohort	1,206	1,310	1,109	0,00001308	-377,70
ovaries	statins_transatlantic_Cohort	0,426	0,527	0,345	0,00000000	345,91
pancreas	statins_transatlantic_Cohort	0,781	0,895	0,681	0,00042104	655,76
prostate	statins_transatlantic_Cohort	1,274	1,388	1,169	0,00000006	-316,16
seq77	statins_transatlantic_Cohort	0,751	0,805	0,700	0,00000000	148,13
seq78	statins_transatlantic_Cohort	0,601	0,641	0,564	0,00000000	64,71
seq79	statins_transatlantic_Cohort	0,672	0,718	0,629	0,00000000	92,31
stomach	statins_transatlantic_Cohort	0,761	0,901	0,643	0,00155665	901,38
bladder	statins_UVA_Cohort	0,433	0,649	0,289	0,00005825	403,96
brain	statins_UVA_Cohort	0,381	0,554	0,262	0,00000066	280,26
breast	statins_UVA_Cohort	0,342	0,478	0,245	0,00000000	190,74
cancer	statins_UVA_Cohort	0,554	0,585	0,526	0,00000000	12,72
cancersubentities	statins_UVA_Cohort	0,528	0,574	0,486	0,00000000	25,96
colorectal	statins_UVA_Cohort	0,434	0,590	0,319	0,00000017	235,43
esophagus	statins_UVA_Cohort	1,113	1,649	0,751	0,60591437	-4490,73
kidney	statins_UVA_Cohort	0,336	0,655	0,172	0,00143681	741,44
liver	statins_UVA_Cohort	0,280	0,597	0,132	0,00098864	734,53
lung	statins_UVA_Cohort	0,863	1,063	0,700	0,16756369	837,78
ovaries	statins_UVA_Cohort	0,361	0,560	0,233	0,00000682	355,42
pancreas	statins_UVA_Cohort	0,566	0,851	0,376	0,00633191	689,46
prostate	statins_UVA_Cohort	0,856	1,063	0,689	0,16044605	854,23
seq77	statins_UVA_Cohort	0,367	0,462	0,292	0,00000000	99,68
seq78	statins_UVA_Cohort	0,426	0,501	0,363	0,00000000	63,37
seq79	statins_UVA_Cohort	0,477	0,559	0,407	0,00000000	74,40
stomach	statins_UVA_Cohort	0,273	0,669	0,112	0,00445089	994,24

		1:1 matched Cohort						
		MEAN Odds-Ratio						
Diagnose	Drug	NbDrug	NbDiag	NbDiag/Drug	Odds Ratio	Conf-Low	Conf-Up	P-Value
bladder	atorvastatin_charite_Cohort	199	2100	75	0,574	0,428	0,77	0,000
brain	atorvastatin_charite_Cohort	66	1296	20	0,424	0,249	0,72	0,002
breast	atorvastatin_charite_Cohort	70	1373	19	0,374	0,219	0,64	0,000
cancer	atorvastatin_charite_Cohort	2727	36203	711	0,341	0,312	0,37	0,000
colorectal	atorvastatin_charite_Cohort	216	3151	50	0,286	0,208	0,39	0,000
esophagus	atorvastatin_charite_Cohort	50	678	7	0,170	0,075	0,38	0,000
kidney	atorvastatin_charite_Cohort	141	1530	45	0,512	0,354	0,74	0,000
liver	atorvastatin_charite_Cohort	110	1651	17	0,179	0,106	0,30	0,000
lung	atorvastatin_charite_Cohort	211	2630	69	0,457	0,341	0,61	0,000
ovaries	atorvastatin_charite_Cohort	35	828	6	0,203	0,083	0,49	0,000
pancreas	atorvastatin_charite_Cohort	115	1464	28	0,371	0,238	0,58	0,000
prostate	atorvastatin_charite_Cohort	239	2409	59	0,328	0,242	0,44	0,000
seq77	atorvastatin_charite_Cohort	317	5507	73	0,261	0,201	0,34	0,000
seq78	atorvastatin_charite_Cohort	430	7200	73	0,203	0,158	0,26	0,000
seq79	atorvastatin_charite_Cohort	376	6024	81	0,257	0,200	0,33	0,000
stomach	atorvastatin_charite_Cohort	66	1057	12	0,212	0,113	0,40	0,000
bladder	atorvastatin_transatlantic_Cohort	288	2489	88	0,397	0,308	0,51	0,000
brain	atorvastatin_transatlantic_Cohort	133	1815	31	0,332	0,220	0,50	0,000
breast	atorvastatin_transatlantic_Cohort	139	2081	29	0,257	0,170	0,39	0,000
cancer	atorvastatin_transatlantic_Cohort	5125	52882	1235	0,299	0,280	0,32	0,000
colorectal	atorvastatin_transatlantic_Cohort	344	3827	71	0,235	0,181	0,31	0,000
esophagus	atorvastatin_transatlantic_Cohort	86	870	20	0,281	0,169	0,47	0,000
kidney	atorvastatin_transatlantic_Cohort	178	1708	50	0,380	0,272	0,53	0,000
liver	atorvastatin_transatlantic_Cohort	157	1822	22	0,150	0,095	0,24	0,000
lung	atorvastatin_transatlantic_Cohort	359	3443	110	0,406	0,323	0,51	0,000
ovaries	atorvastatin_transatlantic_Cohort	82	1219	11	0,170	0,089	0,33	0,000
pancreas	atorvastatin_transatlantic_Cohort	168	1766	40	0,311	0,216	0,45	0,000
prostate	atorvastatin_transatlantic_Cohort	390	3178	98	0,310	0,245	0,39	0,000
seq77	atorvastatin_transatlantic_Cohort	557	6925	106	0,218	0,176	0,27	0,000
seq78	atorvastatin_transatlantic_Cohort	747	9688	130	0,188	0,156	0,23	0,000
seq79	atorvastatin_transatlantic_Cohort	719	8363	148	0,244	0,203	0,29	0,000
stomach	atorvastatin_transatlantic_Cohort	77	1176	13	0,153	0,085	0,28	0,000
bladder	atorvastatin_UVA_Cohort	67	389	13	0,250	0,133	0,47	0,000
brain	atorvastatin_UVA_Cohort	51	519	11	0,263	0,133	0,52	0,000
breast	atorvastatin_UVA_Cohort	63	708	10	0,187	0,094	0,37	0,000
cancer	atorvastatin_UVA_Cohort	2328	16679	524	0,281	0,254	0,31	0,000
colorectal	atorvastatin_UVA_Cohort	100	676	21	0,264	0,160	0,44	0,000
esophagus	atorvastatin_UVA_Cohort	34	192	13	0,585	0,282	1,22	0,151
kidney	atorvastatin_UVA_Cohort	20	178	5	0,274	0,098	0,77	0,014
liver	atorvastatin_UVA_Cohort	26	171	5	0,247	0,089	0,68	0,007
lung	atorvastatin_UVA_Cohort	135	813	41	0,409	0,279	0,60	0,000
ovaries	atorvastatin_UVA_Cohort	34	391	5	0,175	0,066	0,46	0,000
pancreas	atorvastatin_UVA_Cohort	47	302	12	0,345	0,174	0,68	0,002
prostate	atorvastatin_UVA_Cohort	136	769	39	0,340	0,232	0,50	0,000
seq77	atorvastatin_UVA_Cohort	166	1418	33	0,227	0,154	0,33	0,000
seq78	atorvastatin_UVA_Cohort	266	2488	57	0,210	0,157	0,28	0,000
seq79	atorvastatin_UVA_Cohort	296	2339	67	0,256	0,194	0,34	0,000
stomach	atorvastatin_UVA_Cohort	10	119	1	0,080	0,010	0,63	0,017
bladder	fluvastatin_charite_Cohort	31	2119	12	0,810	0,373	1,76	0,607
brain	fluvastatin_charite_Cohort	4	1295	0	0,000	0,000	nan	nan
breast	fluvastatin_charite_Cohort	8	1370	2	0,366	0,070	1,95	0,239
cancer	fluvastatin_charite_Cohort	354	36199	118	0,483	0,387	0,60	0,000
colorectal	fluvastatin_charite_Cohort	28	3162	8	0,383	0,168	0,87	0,022
esophagus	fluvastatin_charite_Cohort	7	683	5	1,143	0,299	4,53	0,858
kidney	fluvastatin_charite_Cohort	19	1535	14	1,392	0,609	3,21	0,443
liver	fluvastatin_charite_Cohort	16	1656	4	0,416	0,123	1,43	0,162
lung	fluvastatin_charite_Cohort	27	2633	8	0,446	0,192	1,04	0,060
ovaries	fluvastatin_charite_Cohort	4	825	1	inf	0,030	inf	nan
pancreas	fluvastatin_charite_Cohort	12	1462	2	0,233	0,048	1,14	0,071
prostate	fluvastatin_charite_Cohort	31	2420	12	0,606	0,292	1,26	0,180
seq77	fluvastatin_charite_Cohort	48	5530	10	0,280	0,138	0,57	0,000
seq78	fluvastatin_charite_Cohort	73	7220	18	0,370	0,215	0,64	0,000
seq79	fluvastatin_charite_Cohort	57	6028	18	0,446	0,255	0,78	0,005
stomach	fluvastatin_charite_Cohort	11	1050	1	0,183	0,021	1,61	0,125
bladder	fluvastatin_transatlantic_Cohort	28	2517	12	0,855	0,392	1,87	0,707
brain	fluvastatin_transatlantic_Cohort	9	1813	0	0,000	0,000	nan	nan
breast	fluvastatin_transatlantic_Cohort	8	2087	2	0,329	0,065	1,67	0,180
cancer	fluvastatin_transatlantic_Cohort	361	53012	119	0,514	0,412	0,64	0,000
colorectal	fluvastatin_transatlantic_Cohort	25	3844	9	0,492	0,220	1,10	0,084
esophagus	fluvastatin_transatlantic_Cohort	11	878	5	inf	0,303	inf	nan
kidney	fluvastatin_transatlantic_Cohort	23	1719	14	1,934	0,749	5,25	0,185
liver	fluvastatin_transatlantic_Cohort	17	1825	4	0,445	0,136	1,46	0,183
lung	fluvastatin_transatlantic_Cohort	24	3469	8	0,513	0,216	1,22	0,132
ovaries	fluvastatin_transatlantic_Cohort	6	1219	1	inf	0,032	inf	nan

Supplementary Table 3

1:1 matched Cohort

Mean Odds-Ratio

pancreas	fluvastatin_transatlantic_Cohort	11	1769	2	0,287	0,058	1,42	0,126
prostate	fluvastatin_transatlantic_Cohort	31	3208	12	0,557	0,273	1,14	0,107
seq77	fluvastatin_transatlantic_Cohort	36	6974	10	0,321	0,157	0,66	0,002
seq78	fluvastatin_transatlantic_Cohort	54	9744	18	0,443	0,254	0,77	0,004
seq79	fluvastatin_transatlantic_Cohort	53	8395	18	0,475	0,270	0,84	0,010
stomach	fluvastatin_transatlantic_Cohort	12	1173	1	0,206	0,023	1,89	0,161
bladder	fluvastatin_UVA_Cohort	0	398	0	nan	nan	nan	nan
brain	fluvastatin_UVA_Cohort	0	518	0	0,000	0,000	nan	nan
breast	fluvastatin_UVA_Cohort	0	717	0	0,000	0,000	nan	nan
cancer	fluvastatin_UVA_Cohort	4	16813	1	0,476	0,041	5,92	0,570
colorectal	fluvastatin_UVA_Cohort	1	682	1	inf	0,062	inf	nan
esophagus	fluvastatin_UVA_Cohort	0	195	0	nan	nan	nan	nan
kidney	fluvastatin_UVA_Cohort	0	184	0	0,000	0,000	nan	nan
liver	fluvastatin_UVA_Cohort	0	169	0	0,000	0,000	nan	nan
lung	fluvastatin_UVA_Cohort	0	836	0	0,000	0,000	nan	nan
ovaries	fluvastatin_UVA_Cohort	0	394	0	0,000	0,000	nan	nan
pancreas	fluvastatin_UVA_Cohort	0	307	0	0,000	0,000	nan	nan
prostate	fluvastatin_UVA_Cohort	0	788	0	0,000	0,000	nan	nan
seq77	fluvastatin_UVA_Cohort	0	1444	0	0,000	0,000	nan	nan
seq78	fluvastatin_UVA_Cohort	0	2524	0	0,000	0,000	nan	nan
seq79	fluvastatin_UVA_Cohort	0	2367	0	0,000	0,000	nan	nan
stomach	fluvastatin_UVA_Cohort	0	123	0	0,000	0,000	nan	nan
bladder	pravastatin_charite_Cohort	37	2121	18	1,000	0,516	1,94	1,000
brain	pravastatin_charite_Cohort	7	1294	2	0,335	0,065	1,73	0,193
breast	pravastatin_charite_Cohort	9	1373	5	0,699	0,217	2,26	0,561
cancer	pravastatin_charite_Cohort	461	36213	178	0,662	0,547	0,80	0,000
colorectal	pravastatin_charite_Cohort	34	3157	6	0,264	0,106	0,66	0,004
esophagus	pravastatin_charite_Cohort	8	684	3	0,652	0,138	3,43	0,614
kidney	pravastatin_charite_Cohort	18	1535	11	1,051	0,440	2,53	0,918
liver	pravastatin_charite_Cohort	15	1652	1	0,069	0,009	0,53	0,010
lung	pravastatin_charite_Cohort	43	2631	12	0,577	0,281	1,19	0,134
ovaries	pravastatin_charite_Cohort	12	826	6	1,282	0,367	4,58	0,713
pancreas	pravastatin_charite_Cohort	13	1462	5	0,481	0,162	1,43	0,189
prostate	pravastatin_charite_Cohort	53	2423	23	0,934	0,526	1,66	0,827
seq77	pravastatin_charite_Cohort	68	5526	24	0,594	0,357	0,99	0,045
seq78	pravastatin_charite_Cohort	80	7215	22	0,432	0,261	0,71	0,001
seq79	pravastatin_charite_Cohort	57	6023	18	0,417	0,240	0,73	0,002
stomach	pravastatin_charite_Cohort	12	1053	4	0,589	0,165	2,12	0,424
bladder	pravastatin_transatlantic_Cohort	52	2516	18	0,459	0,261	0,81	0,007
brain	pravastatin_transatlantic_Cohort	29	1812	6	0,363	0,140	0,94	0,037
breast	pravastatin_transatlantic_Cohort	25	2092	9	0,438	0,198	0,97	0,041
cancer	pravastatin_transatlantic_Cohort	1032	52982	331	0,475	0,416	0,54	0,000
colorectal	pravastatin_transatlantic_Cohort	60	3836	12	0,231	0,123	0,44	0,000
esophagus	pravastatin_transatlantic_Cohort	19	879	4	0,382	0,117	1,25	0,111
kidney	pravastatin_transatlantic_Cohort	37	1717	13	0,585	0,292	1,17	0,130
liver	pravastatin_transatlantic_Cohort	23	1821	2	0,080	0,019	0,34	0,001
lung	pravastatin_transatlantic_Cohort	70	3459	25	0,544	0,333	0,89	0,015
ovaries	pravastatin_transatlantic_Cohort	18	1219	9	0,796	0,326	1,95	0,630
pancreas	pravastatin_transatlantic_Cohort	32	1768	8	0,330	0,147	0,74	0,007
prostate	pravastatin_transatlantic_Cohort	88	3206	41	0,827	0,545	1,26	0,379
seq77	pravastatin_transatlantic_Cohort	113	6965	27	0,322	0,208	0,50	0,000
seq78	pravastatin_transatlantic_Cohort	132	9722	40	0,325	0,227	0,47	0,000
seq79	pravastatin_transatlantic_Cohort	144	8379	35	0,316	0,215	0,46	0,000
stomach	pravastatin_transatlantic_Cohort	12	1175	4	0,294	0,095	0,91	0,033
bladder	pravastatin_UVA_Cohort	14	395	0	0,000	0,000	nan	nan
brain	pravastatin_UVA_Cohort	12	518	4	0,435	0,131	1,46	0,176
breast	pravastatin_UVA_Cohort	18	719	4	0,359	0,112	1,15	0,085
cancer	pravastatin_UVA_Cohort	550	16769	153	0,390	0,323	0,47	0,000
colorectal	pravastatin_UVA_Cohort	16	679	6	0,415	0,158	1,10	0,075
esophagus	pravastatin_UVA_Cohort	7	195	1	0,242	0,025	2,47	0,228
kidney	pravastatin_UVA_Cohort	7	182	2	0,528	0,087	3,40	0,505
liver	pravastatin_UVA_Cohort	5	169	1	0,279	0,029	2,73	0,274
lung	pravastatin_UVA_Cohort	34	828	13	0,653	0,320	1,33	0,244
ovaries	pravastatin_UVA_Cohort	14	393	3	0,423	0,103	1,78	0,239
pancreas	pravastatin_UVA_Cohort	15	306	3	0,412	0,104	1,66	0,211
prostate	pravastatin_UVA_Cohort	40	783	18	0,862	0,452	1,64	0,665
seq77	pravastatin_UVA_Cohort	34	1439	3	0,087	0,027	0,29	0,000
seq78	pravastatin_UVA_Cohort	81	2507	18	0,321	0,188	0,55	0,000
seq79	pravastatin_UVA_Cohort	71	2356	17	0,309	0,178	0,53	0,000
stomach	pravastatin_UVA_Cohort	4	122	0	0,000	0,000	nan	nan
bladder	simvastatin_charite_Cohort	899	2105	431	0,930	0,802	1,08	0,341
brain	simvastatin_charite_Cohort	248	1300	104	0,665	0,511	0,87	0,002
breast	simvastatin_charite_Cohort	282	1366	102	0,515	0,399	0,66	0,000
cancer	simvastatin_charite_Cohort	10668	36506	4257	0,610	0,585	0,64	0,000
colorectal	simvastatin_charite_Cohort	862	3139	319	0,509	0,439	0,59	0,000
esophagus	simvastatin_charite_Cohort	210	704	98	0,734	0,551	0,98	0,035
kidney	simvastatin_charite_Cohort	541	1523	243	0,828	0,686	1,00	0,049

Supplementary Table 3
1:1 matched Cohort
Mean Odds-Ratio

liver	simvastatin_charite_Cohort	466	1646	140	0,409	0,331	0,51	0,000
lung	simvastatin_charite_Cohort	897	2646	415	0,807	0,700	0,93	0,003
ovaries	simvastatin_charite_Cohort	165	833	53	0,461	0,327	0,65	0,000
pancreas	simvastatin_charite_Cohort	396	1473	155	0,563	0,454	0,70	0,000
prostate	simvastatin_charite_Cohort	936	2401	389	0,627	0,543	0,72	0,000
seq77	simvastatin_charite_Cohort	1385	5374	510	0,541	0,481	0,61	0,000
seq78	simvastatin_charite_Cohort	1782	7141	576	0,444	0,400	0,49	0,000
seq79	simvastatin_charite_Cohort	1586	5961	540	0,486	0,435	0,54	0,000
stomach	simvastatin_charite_Cohort	293	1059	116	0,611	0,475	0,79	0,000
bladder	simvastatin_transatlantic_Cohort	866	2502	437	1,063	0,917	1,23	0,424
brain	simvastatin_transatlantic_Cohort	274	1818	113	0,665	0,519	0,85	0,001
breast	simvastatin_transatlantic_Cohort	331	2077	116	0,540	0,427	0,69	0,000
cancer	simvastatin_transatlantic_Cohort	11555	53267	4608	0,634	0,610	0,66	0,000
colorectal	simvastatin_transatlantic_Cohort	849	3816	329	0,597	0,516	0,69	0,000
esophagus	simvastatin_transatlantic_Cohort	231	897	104	0,782	0,593	1,03	0,082
kidney	simvastatin_transatlantic_Cohort	475	1705	243	1,020	0,841	1,24	0,851
liver	simvastatin_transatlantic_Cohort	396	1814	141	0,501	0,404	0,62	0,000
lung	simvastatin_transatlantic_Cohort	929	3472	442	0,896	0,780	1,03	0,119
ovaries	simvastatin_transatlantic_Cohort	168	1224	59	0,462	0,335	0,64	0,000
pancreas	simvastatin_transatlantic_Cohort	382	1778	161	0,646	0,523	0,80	0,000
prostate	simvastatin_transatlantic_Cohort	1010	3183	407	0,662	0,577	0,76	0,000
seq77	simvastatin_transatlantic_Cohort	1402	6807	534	0,601	0,536	0,67	0,000
seq78	simvastatin_transatlantic_Cohort	1843	9644	616	0,476	0,430	0,53	0,000
seq79	simvastatin_transatlantic_Cohort	1644	8314	579	0,502	0,451	0,56	0,000
stomach	simvastatin_transatlantic_Cohort	261	1182	119	0,743	0,576	0,96	0,022
bladder	simvastatin_UVA_Cohort	32	397	6	0,234	0,095	0,58	0,002
brain	simvastatin_UVA_Cohort	29	518	9	0,525	0,231	1,19	0,124
breast	simvastatin_UVA_Cohort	39	711	14	0,521	0,269	1,01	0,052
cancer	simvastatin_UVA_Cohort	1152	16761	351	0,431	0,379	0,49	0,000
colorectal	simvastatin_UVA_Cohort	39	677	10	0,269	0,132	0,55	0,000
esophagus	simvastatin_UVA_Cohort	16	193	6	0,587	0,208	1,66	0,320
kidney	simvastatin_UVA_Cohort	11	182	0	0,000	0,000	nan	nan
liver	simvastatin_UVA_Cohort	6	168	1	0,142	0,017	1,20	0,072
lung	simvastatin_UVA_Cohort	70	826	27	0,597	0,366	0,97	0,039
ovaries	simvastatin_UVA_Cohort	27	391	6	0,383	0,146	1,01	0,051
pancreas	simvastatin_UVA_Cohort	15	305	6	0,391	0,148	1,03	0,058
prostate	simvastatin_UVA_Cohort	60	782	18	0,377	0,217	0,66	0,001
seq77	simvastatin_UVA_Cohort	82	1433	24	0,363	0,226	0,58	0,000
seq78	simvastatin_UVA_Cohort	159	2503	40	0,331	0,230	0,48	0,000
seq79	simvastatin_UVA_Cohort	147	2353	39	0,325	0,225	0,47	0,000
stomach	simvastatin_UVA_Cohort	9	123	3	0,584	0,133	2,59	0,487
bladder	statins_charite_Cohort	1115	2094	548	0,891	0,778	1,02	0,096
brain	statins_charite_Cohort	303	1302	127	0,583	0,460	0,74	0,000
breast	statins_charite_Cohort	372	1363	128	0,465	0,371	0,59	0,000
cancer	statins_charite_Cohort	14079	36497	5295	0,535	0,515	0,56	0,000
cancersubentities	statins_charite_Cohort	8604	21890	3344	0,560	0,533	0,59	0,000
colorectal	statins_charite_Cohort	1145	3133	386	0,442	0,387	0,51	0,000
esophagus	statins_charite_Cohort	279	699	115	0,578	0,445	0,75	0,000
kidney	statins_charite_Cohort	705	1513	318	0,816	0,688	0,97	0,019
liver	statins_charite_Cohort	557	1645	162	0,343	0,281	0,42	0,000
lung	statins_charite_Cohort	1164	2651	507	0,720	0,631	0,82	0,000
ovaries	statins_charite_Cohort	206	837	66	0,428	0,314	0,58	0,000
pancreas	statins_charite_Cohort	535	1476	191	0,490	0,403	0,60	0,000
prostate	statins_charite_Cohort	1253	2395	492	0,564	0,495	0,64	0,000
seq77	statins_charite_Cohort	1791	5354	628	0,480	0,432	0,53	0,000
seq78	statins_charite_Cohort	2252	7125	699	0,389	0,353	0,43	0,000
seq79	statins_charite_Cohort	1985	5955	671	0,433	0,392	0,48	0,000
stomach	statins_charite_Cohort	341	1058	132	0,496	0,393	0,63	0,000
bladder	statins_transatlantic_Cohort	1231	2480	570	0,842	0,739	0,96	0,009
brain	statins_transatlantic_Cohort	434	1823	154	0,525	0,426	0,65	0,000
breast	statins_transatlantic_Cohort	506	2067	158	0,431	0,352	0,53	0,000
cancer	statins_transatlantic_Cohort	18016	53113	6425	0,496	0,480	0,51	0,000
colorectal	statins_transatlantic_Cohort	1285	3803	425	0,439	0,386	0,50	0,000
esophagus	statins_transatlantic_Cohort	333	886	140	0,630	0,496	0,80	0,000
kidney	statins_transatlantic_Cohort	734	1688	325	0,808	0,685	0,95	0,012
liver	statins_transatlantic_Cohort	634	1814	169	0,351	0,289	0,43	0,000
lung	statins_transatlantic_Cohort	1358	3449	596	0,688	0,611	0,77	0,000
ovaries	statins_transatlantic_Cohort	295	1225	82	0,365	0,279	0,48	0,000
pancreas	statins_transatlantic_Cohort	597	1774	213	0,490	0,408	0,59	0,000
prostate	statins_transatlantic_Cohort	1467	3151	572	0,564	0,501	0,64	0,000
seq77	statins_transatlantic_Cohort	2040	6753	693	0,452	0,410	0,50	0,000
seq78	statins_transatlantic_Cohort	2782	9574	827	0,370	0,339	0,40	0,000
seq79	statins_transatlantic_Cohort	2553	8269	808	0,412	0,376	0,45	0,000
stomach	statins_transatlantic_Cohort	380	1176	136	0,497	0,396	0,63	0,000
bladder	statins_UVA_Cohort	96	386	22	0,215	0,131	0,35	0,000
brain	statins_UVA_Cohort	91	521	27	0,366	0,230	0,58	0,000
breast	statins_UVA_Cohort	138	704	30	0,298	0,194	0,46	0,000

Supplementary Table 3
1:1 matched Cohort
Mean Odds-Ratio

cancer	statins_UVA_Cohort	4086	16616	1130	0,338	0,314	0,36	0,000
cancersubentities	statins_UVA_Cohort	1901	7514	500	0,319	0,286	0,36	0,000
colorectal	statins_UVA_Cohort	172	670	39	0,277	0,190	0,40	0,000
esophagus	statins_UVA_Cohort	52	187	25	0,608	0,349	1,06	0,078
kidney	statins_UVA_Cohort	43	175	7	0,196	0,083	0,46	0,000
liver	statins_UVA_Cohort	36	169	7	0,183	0,078	0,43	0,000
lung	statins_UVA_Cohort	244	798	89	0,506	0,382	0,67	0,000
ovaries	statins_UVA_Cohort	87	388	16	0,269	0,151	0,48	0,000
pancreas	statins_UVA_Cohort	86	298	22	0,330	0,196	0,56	0,000
prostate	statins_UVA_Cohort	245	756	80	0,411	0,308	0,55	0,000
seq77	statins_UVA_Cohort	312	1399	65	0,248	0,186	0,33	0,000
seq78	statins_UVA_Cohort	562	2449	128	0,263	0,214	0,32	0,000
seq79	statins_UVA_Cohort	534	2314	137	0,295	0,241	0,36	0,000
stomach	statins_UVA_Cohort	26	118	4	0,167	0,055	0,51	0,002

		1:1 matched Cohort MEDIAN Odds-Ratio			
Diagnose	Drug	Odds Ratio	Conf-Low	Conf-Up	P-Value
bladder	atorvastatin_charite_Cohort	0,569	0,424	0,76	0,000
brain	atorvastatin_charite_Cohort	0,426	0,251	0,72	0,002
breast	atorvastatin_charite_Cohort	0,367	0,216	0,63	0,000
cancer	atorvastatin_charite_Cohort	0,340	0,311	0,37	0,000
colorectal	atorvastatin_charite_Cohort	0,286	0,208	0,40	0,000
esophagus	atorvastatin_charite_Cohort	0,176	0,078	0,40	0,000
kidney	atorvastatin_charite_Cohort	0,503	0,348	0,73	0,000
liver	atorvastatin_charite_Cohort	0,177	0,105	0,30	0,000
lung	atorvastatin_charite_Cohort	0,444	0,332	0,59	0,000
ovaries	atorvastatin_charite_Cohort	0,198	0,082	0,48	0,000
pancreas	atorvastatin_charite_Cohort	0,366	0,236	0,57	0,000
prostate	atorvastatin_charite_Cohort	0,320	0,236	0,43	0,000
seq77	atorvastatin_charite_Cohort	0,262	0,201	0,34	0,000
seq78	atorvastatin_charite_Cohort	0,200	0,155	0,26	0,000
seq79	atorvastatin_charite_Cohort	0,251	0,196	0,32	0,000
stomach	atorvastatin_charite_Cohort	0,213	0,113	0,40	0,000
bladder	atorvastatin_transatlantic_Cohort	0,397	0,307	0,51	0,000
brain	atorvastatin_transatlantic_Cohort	0,335	0,222	0,51	0,000
breast	atorvastatin_transatlantic_Cohort	0,255	0,168	0,39	0,000
cancer	atorvastatin_transatlantic_Cohort	0,300	0,281	0,32	0,000
colorectal	atorvastatin_transatlantic_Cohort	0,236	0,181	0,31	0,000
esophagus	atorvastatin_transatlantic_Cohort	0,273	0,165	0,45	0,000
kidney	atorvastatin_transatlantic_Cohort	0,370	0,266	0,52	0,000
liver	atorvastatin_transatlantic_Cohort	0,153	0,098	0,24	0,000
lung	atorvastatin_transatlantic_Cohort	0,409	0,326	0,51	0,000
ovaries	atorvastatin_transatlantic_Cohort	0,164	0,086	0,31	0,000
pancreas	atorvastatin_transatlantic_Cohort	0,305	0,213	0,44	0,000
prostate	atorvastatin_transatlantic_Cohort	0,307	0,243	0,39	0,000
seq77	atorvastatin_transatlantic_Cohort	0,219	0,177	0,27	0,000
seq78	atorvastatin_transatlantic_Cohort	0,189	0,155	0,23	0,000
seq79	atorvastatin_transatlantic_Cohort	0,245	0,204	0,30	0,000
stomach	atorvastatin_transatlantic_Cohort	0,153	0,085	0,28	0,000
bladder	atorvastatin_UVA_Cohort	0,249	0,133	0,47	0,000
brain	atorvastatin_UVA_Cohort	0,256	0,130	0,50	0,000
breast	atorvastatin_UVA_Cohort	0,195	0,098	0,39	0,000
cancer	atorvastatin_UVA_Cohort	0,282	0,255	0,31	0,000
colorectal	atorvastatin_UVA_Cohort	0,251	0,153	0,41	0,000
esophagus	atorvastatin_UVA_Cohort	0,561	0,274	1,15	0,114
kidney	atorvastatin_UVA_Cohort	0,266	0,096	0,73	0,011
liver	atorvastatin_UVA_Cohort	0,241	0,088	0,66	0,006
lung	atorvastatin_UVA_Cohort	0,413	0,282	0,61	0,000
ovaries	atorvastatin_UVA_Cohort	0,168	0,064	0,44	0,000
pancreas	atorvastatin_UVA_Cohort	0,327	0,166	0,64	0,001
prostate	atorvastatin_UVA_Cohort	0,342	0,233	0,50	0,000
seq77	atorvastatin_UVA_Cohort	0,224	0,151	0,33	0,000
seq78	atorvastatin_UVA_Cohort	0,210	0,157	0,28	0,000
seq79	atorvastatin_UVA_Cohort	0,256	0,194	0,34	0,000
stomach	atorvastatin_UVA_Cohort	0,080	0,010	0,62	0,016
bladder	fluvastatin_charite_Cohort	0,749	0,353	1,59	0,460
brain	fluvastatin_charite_Cohort	0,000	0,000	nan	nan
breast	fluvastatin_charite_Cohort	0,308	0,063	1,51	0,147
cancer	fluvastatin_charite_Cohort	0,480	0,385	0,60	0,000
colorectal	fluvastatin_charite_Cohort	0,398	0,175	0,91	0,028
esophagus	fluvastatin_charite_Cohort	1,000	0,288	3,47	1,000

Supplementary Table 3
1:1 matched Cohort
Median Odds-Ratio

kidney	fluvastatin_charite_Cohort	1,339	0,599	2,99	0,487
liver	fluvastatin_charite_Cohort	0,319	0,104	0,99	0,046
lung	fluvastatin_charite_Cohort	0,419	0,183	0,96	0,039
ovaries	fluvastatin_charite_Cohort	0,249	0,028	2,23	0,215
pancreas	fluvastatin_charite_Cohort	0,199	0,044	0,91	0,036
prostate	fluvastatin_charite_Cohort	0,569	0,279	1,16	0,121
seq77	fluvastatin_charite_Cohort	0,269	0,134	0,54	0,000
seq78	fluvastatin_charite_Cohort	0,366	0,213	0,63	0,000
seq79	fluvastatin_charite_Cohort	0,454	0,260	0,79	0,006
stomach	fluvastatin_charite_Cohort	0,166	0,020	1,38	0,096
bladder	fluvastatin_transatlantic_Cohort	0,774	0,363	1,65	0,517
brain	fluvastatin_transatlantic_Cohort	0,000	0,000	nan	nan
breast	fluvastatin_transatlantic_Cohort	0,333	0,067	1,65	0,179
cancer	fluvastatin_transatlantic_Cohort	0,514	0,412	0,64	0,000
colorectal	fluvastatin_transatlantic_Cohort	0,499	0,224	1,11	0,089
esophagus	fluvastatin_transatlantic_Cohort	1,000	0,288	3,47	1,000
kidney	fluvastatin_transatlantic_Cohort	1,658	0,704	3,91	0,250
liver	fluvastatin_transatlantic_Cohort	0,443	0,136	1,44	0,177
lung	fluvastatin_transatlantic_Cohort	0,469	0,202	1,09	0,078
ovaries	fluvastatin_transatlantic_Cohort	0,333	0,035	3,20	0,345
pancreas	fluvastatin_transatlantic_Cohort	0,249	0,053	1,18	0,078
prostate	fluvastatin_transatlantic_Cohort	0,544	0,269	1,10	0,090
seq77	fluvastatin_transatlantic_Cohort	0,312	0,153	0,64	0,001
seq78	fluvastatin_transatlantic_Cohort	0,423	0,243	0,73	0,002
seq79	fluvastatin_transatlantic_Cohort	0,455	0,260	0,79	0,006
stomach	fluvastatin_transatlantic_Cohort	0,166	0,020	1,38	0,096
bladder	fluvastatin_UVA_Cohort	nan	nan	nan	nan
brain	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
breast	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
cancer	fluvastatin_UVA_Cohort	0,333	0,035	3,21	0,345
colorectal	fluvastatin_UVA_Cohort	inf	0,062	inf	nan
esophagus	fluvastatin_UVA_Cohort	nan	nan	nan	nan
kidney	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
liver	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
lung	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
ovaries	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
pancreas	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
prostate	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
seq77	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
seq78	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
seq79	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
stomach	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
bladder	pravastatin_charite_Cohort	1,000	0,519	1,93	1,000
brain	pravastatin_charite_Cohort	0,285	0,059	1,37	0,118
breast	pravastatin_charite_Cohort	0,713	0,226	2,25	0,576
cancer	pravastatin_charite_Cohort	0,659	0,545	0,80	0,000
colorectal	pravastatin_charite_Cohort	0,244	0,100	0,60	0,002
esophagus	pravastatin_charite_Cohort	0,462	0,117	1,83	0,274
kidney	pravastatin_charite_Cohort	0,881	0,390	1,99	0,773
liver	pravastatin_charite_Cohort	0,066	0,009	0,50	0,008
lung	pravastatin_charite_Cohort	0,569	0,280	1,16	0,120

Supplementary Table 3
1:1 matched Cohort
Median Odds-Ratio

ovaries	pravastatin_charite_Cohort	1,101	0,343	3,53	0,881
pancreas	pravastatin_charite_Cohort	0,453	0,157	1,31	0,143
prostate	pravastatin_charite_Cohort	0,939	0,530	1,66	0,840
seq77	pravastatin_charite_Cohort	0,570	0,344	0,94	0,028
seq78	pravastatin_charite_Cohort	0,430	0,260	0,71	0,001
seq79	pravastatin_charite_Cohort	0,407	0,235	0,71	0,001
stomach	pravastatin_charite_Cohort	0,534	0,158	1,81	0,317
bladder	pravastatin_transatlantic_Cohort	0,446	0,255	0,78	0,005
brain	pravastatin_transatlantic_Cohort	0,341	0,135	0,86	0,023
breast	pravastatin_transatlantic_Cohort	0,407	0,187	0,89	0,023
cancer	pravastatin_transatlantic_Cohort	0,475	0,416	0,54	0,000
colorectal	pravastatin_transatlantic_Cohort	0,228	0,122	0,43	0,000
esophagus	pravastatin_transatlantic_Cohort	0,318	0,103	0,98	0,046
kidney	pravastatin_transatlantic_Cohort	0,562	0,284	1,11	0,098
liver	pravastatin_transatlantic_Cohort	0,073	0,017	0,31	0,000
lung	pravastatin_transatlantic_Cohort	0,534	0,328	0,87	0,012
ovaries	pravastatin_transatlantic_Cohort	0,748	0,314	1,78	0,523
pancreas	pravastatin_transatlantic_Cohort	0,330	0,148	0,74	0,007
prostate	pravastatin_transatlantic_Cohort	0,818	0,539	1,24	0,350
seq77	pravastatin_transatlantic_Cohort	0,325	0,210	0,50	0,000
seq78	pravastatin_transatlantic_Cohort	0,321	0,225	0,46	0,000
seq79	pravastatin_transatlantic_Cohort	0,311	0,212	0,46	0,000
stomach	pravastatin_transatlantic_Cohort	0,283	0,093	0,86	0,026
bladder	pravastatin_UVA_Cohort	0,000	0,000	nan	nan
brain	pravastatin_UVA_Cohort	0,395	0,123	1,27	0,119
breast	pravastatin_UVA_Cohort	0,330	0,106	1,03	0,055
cancer	pravastatin_UVA_Cohort	0,386	0,320	0,47	0,000
colorectal	pravastatin_UVA_Cohort	0,382	0,148	0,99	0,046
esophagus	pravastatin_UVA_Cohort	0,196	0,023	1,69	0,137
kidney	pravastatin_UVA_Cohort	0,444	0,082	2,39	0,351
liver	pravastatin_UVA_Cohort	0,246	0,027	2,22	0,214
lung	pravastatin_UVA_Cohort	0,644	0,318	1,30	0,224
ovaries	pravastatin_UVA_Cohort	0,349	0,092	1,31	0,121
pancreas	pravastatin_UVA_Cohort	0,369	0,097	1,40	0,144
prostate	pravastatin_UVA_Cohort	0,795	0,424	1,49	0,484
seq77	pravastatin_UVA_Cohort	0,089	0,027	0,29	0,000
seq78	pravastatin_UVA_Cohort	0,320	0,188	0,55	0,000
seq79	pravastatin_UVA_Cohort	0,290	0,169	0,50	0,000
stomach	pravastatin_UVA_Cohort	0,000	0,000	nan	nan
bladder	simvastatin_charite_Cohort	0,926	0,798	1,07	0,315
brain	simvastatin_charite_Cohort	0,672	0,516	0,87	0,003
breast	simvastatin_charite_Cohort	0,512	0,397	0,66	0,000
cancer	simvastatin_charite_Cohort	0,612	0,587	0,64	0,000
colorectal	simvastatin_charite_Cohort	0,506	0,436	0,59	0,000
esophagus	simvastatin_charite_Cohort	0,710	0,534	0,95	0,019
kidney	simvastatin_charite_Cohort	0,830	0,688	1,00	0,052
liver	simvastatin_charite_Cohort	0,406	0,329	0,50	0,000
lung	simvastatin_charite_Cohort	0,794	0,689	0,92	0,002
ovaries	simvastatin_charite_Cohort	0,452	0,320	0,64	0,000
pancreas	simvastatin_charite_Cohort	0,563	0,455	0,70	0,000
prostate	simvastatin_charite_Cohort	0,622	0,539	0,72	0,000
seq77	simvastatin_charite_Cohort	0,538	0,479	0,61	0,000
seq78	simvastatin_charite_Cohort	0,445	0,401	0,50	0,000
seq79	simvastatin_charite_Cohort	0,482	0,431	0,54	0,000

Supplementary Table 3
1:1 matched Cohort
Median Odds-Ratio

stomach	simvastatin_charite_Cohort	0,603	0,469	0,77	0,000
bladder	simvastatin_transatlantic_Cohort	1,055	0,911	1,22	0,486
brain	simvastatin_transatlantic_Cohort	0,660	0,514	0,85	0,001
breast	simvastatin_transatlantic_Cohort	0,538	0,425	0,68	0,000
cancer	simvastatin_transatlantic_Cohort	0,633	0,608	0,66	0,000
colorectal	simvastatin_transatlantic_Cohort	0,597	0,516	0,69	0,000
esophagus	simvastatin_transatlantic_Cohort	0,788	0,597	1,04	0,092
kidney	simvastatin_transatlantic_Cohort	1,030	0,849	1,25	0,777
liver	simvastatin_transatlantic_Cohort	0,508	0,409	0,63	0,000
lung	simvastatin_transatlantic_Cohort	0,891	0,776	1,02	0,101
ovaries	simvastatin_transatlantic_Cohort	0,457	0,332	0,63	0,000
pancreas	simvastatin_transatlantic_Cohort	0,651	0,526	0,80	0,000
prostate	simvastatin_transatlantic_Cohort	0,661	0,576	0,76	0,000
seq77	simvastatin_transatlantic_Cohort	0,604	0,539	0,68	0,000
seq78	simvastatin_transatlantic_Cohort	0,476	0,431	0,53	0,000
seq79	simvastatin_transatlantic_Cohort	0,505	0,454	0,56	0,000
stomach	simvastatin_transatlantic_Cohort	0,718	0,557	0,92	0,010
bladder	simvastatin_UVA_Cohort	0,219	0,089	0,54	0,001
brain	simvastatin_UVA_Cohort	0,555	0,243	1,27	0,163
breast	simvastatin_UVA_Cohort	0,509	0,265	0,98	0,042
cancer	simvastatin_UVA_Cohort	0,428	0,377	0,49	0,000
colorectal	simvastatin_UVA_Cohort	0,267	0,131	0,54	0,000
esophagus	simvastatin_UVA_Cohort	0,559	0,201	1,56	0,269
kidney	simvastatin_UVA_Cohort	0,000	0,000	nan	nan
liver	simvastatin_UVA_Cohort	0,129	0,016	1,05	0,055
lung	simvastatin_UVA_Cohort	0,601	0,368	0,98	0,041
ovaries	simvastatin_UVA_Cohort	0,354	0,138	0,91	0,031
pancreas	simvastatin_UVA_Cohort	0,375	0,144	0,98	0,044
prostate	simvastatin_UVA_Cohort	0,364	0,210	0,63	0,000
seq77	simvastatin_UVA_Cohort	0,364	0,227	0,59	0,000
seq78	simvastatin_UVA_Cohort	0,328	0,228	0,47	0,000
seq79	simvastatin_UVA_Cohort	0,333	0,230	0,48	0,000
stomach	simvastatin_UVA_Cohort	0,487	0,119	2,00	0,322
bladder	statins_charite_Cohort	0,893	0,779	1,02	0,103
brain	statins_charite_Cohort	0,580	0,458	0,73	0,000
breast	statins_charite_Cohort	0,468	0,373	0,59	0,000
cancer	statins_charite_Cohort	0,535	0,515	0,56	0,000
cancersubentities	statins_charite_Cohort	0,558	0,532	0,59	0,000
colorectal	statins_charite_Cohort	0,442	0,386	0,50	0,000
esophagus	statins_charite_Cohort	0,576	0,443	0,75	0,000
kidney	statins_charite_Cohort	0,816	0,689	0,97	0,019
liver	statins_charite_Cohort	0,342	0,280	0,42	0,000
lung	statins_charite_Cohort	0,712	0,625	0,81	0,000
ovaries	statins_charite_Cohort	0,423	0,310	0,58	0,000
pancreas	statins_charite_Cohort	0,494	0,406	0,60	0,000
prostate	statins_charite_Cohort	0,565	0,495	0,64	0,000
seq77	statins_charite_Cohort	0,480	0,432	0,53	0,000
seq78	statins_charite_Cohort	0,389	0,353	0,43	0,000
seq79	statins_charite_Cohort	0,430	0,389	0,48	0,000
stomach	statins_charite_Cohort	0,491	0,389	0,62	0,000
bladder	statins_transatlantic_Cohort	0,831	0,730	0,95	0,005
brain	statins_transatlantic_Cohort	0,527	0,428	0,65	0,000
breast	statins_transatlantic_Cohort	0,426	0,349	0,52	0,000
cancer	statins_transatlantic_Cohort	0,494	0,478	0,51	1.97626258336E-323

Supplementary Table 3
1:1 matched Cohort
Median Odds-Ratio

colorectal	statins_transatlantic_Cohort	0,434	0,382	0,49	0,000
esophagus	statins_transatlantic_Cohort	0,637	0,502	0,81	0,000
kidney	statins_transatlantic_Cohort	0,803	0,681	0,95	0,009
liver	statins_transatlantic_Cohort	0,350	0,289	0,43	0,000
lung	statins_transatlantic_Cohort	0,682	0,606	0,77	0,000
ovaries	statins_transatlantic_Cohort	0,363	0,278	0,48	0,000
pancreas	statins_transatlantic_Cohort	0,486	0,405	0,58	0,000
prostate	statins_transatlantic_Cohort	0,558	0,495	0,63	0,000
seq77	statins_transatlantic_Cohort	0,450	0,407	0,50	0,000
seq78	statins_transatlantic_Cohort	0,369	0,339	0,40	0,000
seq79	statins_transatlantic_Cohort	0,411	0,376	0,45	0,000
stomach	statins_transatlantic_Cohort	0,497	0,396	0,62	0,000
bladder	statins_UVA_Cohort	0,214	0,131	0,35	0,000
brain	statins_UVA_Cohort	0,352	0,222	0,56	0,000
breast	statins_UVA_Cohort	0,300	0,196	0,46	0,000
cancer	statins_UVA_Cohort	0,338	0,315	0,36	0,000
cancersubentities	statins_UVA_Cohort	0,317	0,284	0,35	0,000
colorectal	statins_UVA_Cohort	0,274	0,188	0,40	0,000
esophagus	statins_UVA_Cohort	0,605	0,349	1,05	0,073
kidney	statins_UVA_Cohort	0,186	0,080	0,44	0,000
liver	statins_UVA_Cohort	0,168	0,072	0,39	0,000
lung	statins_UVA_Cohort	0,501	0,378	0,66	0,000
ovaries	statins_UVA_Cohort	0,260	0,146	0,46	0,000
pancreas	statins_UVA_Cohort	0,323	0,192	0,54	0,000
prostate	statins_UVA_Cohort	0,422	0,316	0,56	0,000
seq77	statins_UVA_Cohort	0,244	0,183	0,33	0,000
seq78	statins_UVA_Cohort	0,263	0,213	0,32	0,000
seq79	statins_UVA_Cohort	0,298	0,243	0,37	0,000
stomach	statins_UVA_Cohort	0,149	0,050	0,45	0,001

		1:1 matched Cohort			
		MEAN Relative Risk			
Diagnose	Drug	Relative Risk	Conf-Low	Conf-Up	P-Value
bladder	atorvastatin_charite_Cohort	0,730	0,609	0,88	0,001
brain	atorvastatin_charite_Cohort	0,596	0,413	0,86	0,006
breast	atorvastatin_charite_Cohort	0,545	0,371	0,80	0,002
cancer	atorvastatin_charite_Cohort	0,513	0,481	0,55	0,000
colorectal	atorvastatin_charite_Cohort	0,450	0,352	0,57	0,000
esophagus	atorvastatin_charite_Cohort	0,294	0,148	0,58	0,000
kidney	atorvastatin_charite_Cohort	0,678	0,534	0,86	0,001
liver	atorvastatin_charite_Cohort	0,308	0,199	0,48	0,000
lung	atorvastatin_charite_Cohort	0,629	0,517	0,77	0,000
ovaries	atorvastatin_charite_Cohort	0,338	0,163	0,70	0,004
pancreas	atorvastatin_charite_Cohort	0,544	0,396	0,75	0,000
prostate	atorvastatin_charite_Cohort	0,499	0,400	0,62	0,000
seq77	atorvastatin_charite_Cohort	0,418	0,341	0,51	0,000
seq78	atorvastatin_charite_Cohort	0,342	0,278	0,42	0,000
seq79	atorvastatin_charite_Cohort	0,413	0,340	0,50	0,000
stomach	atorvastatin_charite_Cohort	0,354	0,212	0,59	0,000
bladder	atorvastatin_transatlantic_Cohort	0,575	0,481	0,69	0,000
brain	atorvastatin_transatlantic_Cohort	0,502	0,370	0,68	0,000
breast	atorvastatin_transatlantic_Cohort	0,413	0,298	0,57	0,000
cancer	atorvastatin_transatlantic_Cohort	0,467	0,445	0,49	0,000
colorectal	atorvastatin_transatlantic_Cohort	0,388	0,315	0,48	0,000
esophagus	atorvastatin_transatlantic_Cohort	0,445	0,302	0,66	0,000
kidney	atorvastatin_transatlantic_Cohort	0,555	0,438	0,70	0,000
liver	atorvastatin_transatlantic_Cohort	0,267	0,181	0,40	0,000
lung	atorvastatin_transatlantic_Cohort	0,583	0,497	0,68	0,000
ovaries	atorvastatin_transatlantic_Cohort	0,295	0,171	0,51	0,000
pancreas	atorvastatin_transatlantic_Cohort	0,480	0,366	0,63	0,000
prostate	atorvastatin_transatlantic_Cohort	0,481	0,405	0,57	0,000
seq77	atorvastatin_transatlantic_Cohort	0,365	0,307	0,43	0,000
seq78	atorvastatin_transatlantic_Cohort	0,323	0,276	0,38	0,000
seq79	atorvastatin_transatlantic_Cohort	0,399	0,345	0,46	0,000
stomach	atorvastatin_transatlantic_Cohort	0,271	0,163	0,45	0,000
bladder	atorvastatin_UVA_Cohort	0,410	0,252	0,67	0,000
brain	atorvastatin_UVA_Cohort	0,422	0,250	0,71	0,001
breast	atorvastatin_UVA_Cohort	0,322	0,182	0,57	0,000
cancer	atorvastatin_UVA_Cohort	0,448	0,416	0,48	0,000
colorectal	atorvastatin_UVA_Cohort	0,427	0,292	0,63	0,000
esophagus	atorvastatin_UVA_Cohort	0,728	0,471	1,13	0,155
kidney	atorvastatin_UVA_Cohort	0,432	0,200	0,94	0,033
liver	atorvastatin_UVA_Cohort	0,404	0,185	0,88	0,023
lung	atorvastatin_UVA_Cohort	0,587	0,453	0,76	0,000
ovaries	atorvastatin_UVA_Cohort	0,301	0,135	0,68	0,004
pancreas	atorvastatin_UVA_Cohort	0,520	0,319	0,85	0,009
prostate	atorvastatin_UVA_Cohort	0,519	0,396	0,68	0,000
seq77	atorvastatin_UVA_Cohort	0,378	0,278	0,52	0,000
seq78	atorvastatin_UVA_Cohort	0,357	0,282	0,45	0,000
seq79	atorvastatin_UVA_Cohort	0,417	0,337	0,52	0,000
stomach	atorvastatin_UVA_Cohort	0,154	0,023	1,01	0,052
bladder	fluvastatin_charite_Cohort	0,877	0,576	1,34	0,552

Supplementary Table 3
1:1 matched Cohort
Mean Relative Risk

brain	fluvastatin_charite_Cohort	0,000	0,000	nan	nan
breast	fluvastatin_charite_Cohort	0,517	0,161	1,67	0,273
cancer	fluvastatin_charite_Cohort	0,651	0,562	0,76	0,000
colorectal	fluvastatin_charite_Cohort	0,549	0,305	0,99	0,045
esophagus	fluvastatin_charite_Cohort	1,013	0,557	1,85	0,970
kidney	fluvastatin_charite_Cohort	1,141	0,811	1,61	0,458
liver	fluvastatin_charite_Cohort	0,552	0,246	1,24	0,151
lung	fluvastatin_charite_Cohort	0,609	0,343	1,08	0,090
ovaries	fluvastatin_charite_Cohort	0,503	0,154	2,37	0,330
pancreas	fluvastatin_charite_Cohort	0,365	0,107	1,25	0,108
prostate	fluvastatin_charite_Cohort	0,741	0,475	1,16	0,188
seq77	fluvastatin_charite_Cohort	0,436	0,253	0,75	0,003
seq78	fluvastatin_charite_Cohort	0,538	0,363	0,80	0,002
seq79	fluvastatin_charite_Cohort	0,616	0,419	0,90	0,013
stomach	fluvastatin_charite_Cohort	0,302	0,051	1,80	0,189
bladder	fluvastatin_transatlantic_Cohort	0,909	0,601	1,38	0,665
brain	fluvastatin_transatlantic_Cohort	0,000	0,000	nan	nan
breast	fluvastatin_transatlantic_Cohort	0,484	0,147	1,60	0,235
cancer	fluvastatin_transatlantic_Cohort	0,679	0,587	0,79	0,000
colorectal	fluvastatin_transatlantic_Cohort	0,655	0,384	1,12	0,120
esophagus	fluvastatin_transatlantic_Cohort	1,056	0,618	1,85	0,856
kidney	fluvastatin_transatlantic_Cohort	1,265	0,926	1,73	0,141
liver	fluvastatin_transatlantic_Cohort	0,609	0,270	1,37	0,234
lung	fluvastatin_transatlantic_Cohort	0,665	0,380	1,17	0,154
ovaries	fluvastatin_transatlantic_Cohort	0,532	0,160	2,52	0,376
pancreas	fluvastatin_transatlantic_Cohort	0,436	0,130	1,47	0,180
prostate	fluvastatin_transatlantic_Cohort	0,711	0,452	1,12	0,140
seq77	fluvastatin_transatlantic_Cohort	0,484	0,283	0,83	0,008
seq78	fluvastatin_transatlantic_Cohort	0,611	0,416	0,90	0,012
seq79	fluvastatin_transatlantic_Cohort	0,639	0,437	0,94	0,021
stomach	fluvastatin_transatlantic_Cohort	0,329	0,057	1,92	0,217
bladder	fluvastatin_UVA_Cohort	nan	nan	nan	nan
brain	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
breast	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
cancer	fluvastatin_UVA_Cohort	0,605	0,126	2,98	0,544
colorectal	fluvastatin_UVA_Cohort	1,847	1,644	2,40	0,000
esophagus	fluvastatin_UVA_Cohort	nan	nan	nan	nan
kidney	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
liver	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
lung	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
ovaries	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
pancreas	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
prostate	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
seq77	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
seq78	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
seq79	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
stomach	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
bladder	pravastatin_charite_Cohort	0,989	0,713	1,37	0,952
brain	pravastatin_charite_Cohort	0,486	0,149	1,59	0,235
breast	pravastatin_charite_Cohort	0,798	0,409	1,56	0,519
cancer	pravastatin_charite_Cohort	0,796	0,710	0,89	0,000
colorectal	pravastatin_charite_Cohort	0,411	0,203	0,84	0,014
esophagus	pravastatin_charite_Cohort	0,724	0,304	1,74	0,478

Supplementary Table 3
1:1 matched Cohort
Mean Relative Risk

kidney	pravastatin_charite_Cohort	0,993	0,657	1,50	0,976
liver	pravastatin_charite_Cohort	0,129	0,019	0,85	0,034
lung	pravastatin_charite_Cohort	0,724	0,462	1,14	0,160
ovaries	pravastatin_charite_Cohort	1,083	0,637	1,85	0,782
pancreas	pravastatin_charite_Cohort	0,634	0,310	1,30	0,214
prostate	pravastatin_charite_Cohort	0,959	0,714	1,29	0,793
seq77	pravastatin_charite_Cohort	0,743	0,541	1,02	0,066
seq78	pravastatin_charite_Cohort	0,602	0,424	0,85	0,004
seq79	pravastatin_charite_Cohort	0,587	0,398	0,86	0,007
stomach	pravastatin_charite_Cohort	0,714	0,331	1,55	0,399
bladder	pravastatin_transatlantic_Cohort	0,628	0,428	0,92	0,017
brain	pravastatin_transatlantic_Cohort	0,523	0,265	1,03	0,062
breast	pravastatin_transatlantic_Cohort	0,601	0,350	1,04	0,065
cancer	pravastatin_transatlantic_Cohort	0,645	0,590	0,70	0,000
colorectal	pravastatin_transatlantic_Cohort	0,375	0,226	0,62	0,000
esophagus	pravastatin_transatlantic_Cohort	0,537	0,235	1,23	0,141
kidney	pravastatin_transatlantic_Cohort	0,730	0,474	1,12	0,154
liver	pravastatin_transatlantic_Cohort	0,149	0,039	0,56	0,005
lung	pravastatin_transatlantic_Cohort	0,702	0,512	0,96	0,028
ovaries	pravastatin_transatlantic_Cohort	0,869	0,534	1,42	0,584
pancreas	pravastatin_transatlantic_Cohort	0,494	0,272	0,90	0,021
prostate	pravastatin_transatlantic_Cohort	0,903	0,720	1,13	0,385
seq77	pravastatin_transatlantic_Cohort	0,486	0,350	0,68	0,000
seq78	pravastatin_transatlantic_Cohort	0,491	0,375	0,64	0,000
seq79	pravastatin_transatlantic_Cohort	0,480	0,360	0,64	0,000
stomach	pravastatin_transatlantic_Cohort	0,450	0,191	1,06	0,067
bladder	pravastatin_UVA_Cohort	0,000	0,000	nan	nan
brain	pravastatin_UVA_Cohort	0,591	0,263	1,33	0,206
breast	pravastatin_UVA_Cohort	0,519	0,225	1,20	0,124
cancer	pravastatin_UVA_Cohort	0,563	0,492	0,64	0,000
colorectal	pravastatin_UVA_Cohort	0,579	0,296	1,13	0,110
esophagus	pravastatin_UVA_Cohort	0,369	0,066	2,08	0,261
kidney	pravastatin_UVA_Cohort	0,652	0,216	1,99	0,459
liver	pravastatin_UVA_Cohort	0,424	0,077	2,36	0,330
lung	pravastatin_UVA_Cohort	0,780	0,511	1,19	0,253
ovaries	pravastatin_UVA_Cohort	0,564	0,222	1,44	0,233
pancreas	pravastatin_UVA_Cohort	0,569	0,221	1,47	0,245
prostate	pravastatin_UVA_Cohort	0,915	0,650	1,29	0,623
seq77	pravastatin_UVA_Cohort	0,162	0,054	0,48	0,001
seq78	pravastatin_UVA_Cohort	0,486	0,326	0,73	0,000
seq79	pravastatin_UVA_Cohort	0,472	0,312	0,72	0,000
stomach	pravastatin_UVA_Cohort	0,000	0,000	nan	nan
bladder	simvastatin_charite_Cohort	0,963	0,893	1,04	0,334
brain	simvastatin_charite_Cohort	0,800	0,687	0,93	0,004
breast	simvastatin_charite_Cohort	0,687	0,585	0,81	0,000
cancer	simvastatin_charite_Cohort	0,764	0,746	0,78	0,000
colorectal	simvastatin_charite_Cohort	0,685	0,625	0,75	0,000
esophagus	simvastatin_charite_Cohort	0,849	0,724	1,00	0,043
kidney	simvastatin_charite_Cohort	0,906	0,820	1,00	0,053
liver	simvastatin_charite_Cohort	0,595	0,516	0,69	0,000
lung	simvastatin_charite_Cohort	0,895	0,829	0,97	0,005
ovaries	simvastatin_charite_Cohort	0,638	0,509	0,80	0,000
pancreas	simvastatin_charite_Cohort	0,728	0,639	0,83	0,000

Supplementary Table 3
1:1 matched Cohort
Mean Relative Risk

prostate	simvastatin_charite_Cohort	0,779	0,717	0,85	0,000
seq77	simvastatin_charite_Cohort	0,710	0,661	0,76	0,000
seq78	simvastatin_charite_Cohort	0,627	0,585	0,67	0,000
seq79	simvastatin_charite_Cohort	0,665	0,619	0,71	0,000
stomach	simvastatin_charite_Cohort	0,763	0,658	0,89	0,000
bladder	simvastatin_transatlantic_Cohort	1,030	0,958	1,11	0,431
brain	simvastatin_transatlantic_Cohort	0,800	0,691	0,93	0,003
breast	simvastatin_transatlantic_Cohort	0,706	0,608	0,82	0,000
cancer	simvastatin_transatlantic_Cohort	0,781	0,763	0,80	0,000
colorectal	simvastatin_transatlantic_Cohort	0,753	0,690	0,82	0,000
esophagus	simvastatin_transatlantic_Cohort	0,878	0,755	1,02	0,092
kidney	simvastatin_transatlantic_Cohort	1,009	0,917	1,11	0,864
liver	simvastatin_transatlantic_Cohort	0,676	0,589	0,78	0,000
lung	simvastatin_transatlantic_Cohort	0,945	0,879	1,02	0,123
ovaries	simvastatin_transatlantic_Cohort	0,637	0,515	0,79	0,000
pancreas	simvastatin_transatlantic_Cohort	0,788	0,696	0,89	0,000
prostate	simvastatin_transatlantic_Cohort	0,802	0,741	0,87	0,000
seq77	simvastatin_transatlantic_Cohort	0,755	0,705	0,81	0,000
seq78	simvastatin_transatlantic_Cohort	0,653	0,612	0,70	0,000
seq79	simvastatin_transatlantic_Cohort	0,676	0,631	0,72	0,000
stomach	simvastatin_transatlantic_Cohort	0,853	0,740	0,98	0,027
bladder	simvastatin_UVA_Cohort	0,383	0,186	0,78	0,009
brain	simvastatin_UVA_Cohort	0,686	0,403	1,17	0,165
breast	simvastatin_UVA_Cohort	0,684	0,447	1,05	0,080
cancer	simvastatin_UVA_Cohort	0,606	0,555	0,66	0,000
colorectal	simvastatin_UVA_Cohort	0,427	0,247	0,74	0,002
esophagus	simvastatin_UVA_Cohort	0,734	0,388	1,39	0,348
kidney	simvastatin_UVA_Cohort	0,000	0,000	nan	nan
liver	simvastatin_UVA_Cohort	0,248	0,040	1,53	0,134
lung	simvastatin_UVA_Cohort	0,748	0,553	1,01	0,059
ovaries	simvastatin_UVA_Cohort	0,549	0,279	1,08	0,083
pancreas	simvastatin_UVA_Cohort	0,558	0,284	1,10	0,091
prostate	simvastatin_UVA_Cohort	0,550	0,371	0,82	0,003
seq77	simvastatin_UVA_Cohort	0,535	0,379	0,75	0,000
seq78	simvastatin_UVA_Cohort	0,501	0,383	0,66	0,000
seq79	simvastatin_UVA_Cohort	0,494	0,376	0,65	0,000
stomach	simvastatin_UVA_Cohort	0,716	0,293	1,76	0,474
bladder	statins_charite_Cohort	0,943	0,879	1,01	0,100
brain	statins_charite_Cohort	0,743	0,644	0,86	0,000
breast	statins_charite_Cohort	0,647	0,559	0,75	0,000
cancer	statins_charite_Cohort	0,710	0,694	0,73	0,000
cancersubentities	statins_charite_Cohort	0,729	0,709	0,75	0,000
colorectal	statins_charite_Cohort	0,631	0,579	0,69	0,000
esophagus	statins_charite_Cohort	0,743	0,638	0,87	0,000
kidney	statins_charite_Cohort	0,900	0,823	0,99	0,021
liver	statins_charite_Cohort	0,533	0,466	0,61	0,000
lung	statins_charite_Cohort	0,842	0,783	0,90	0,000
ovaries	statins_charite_Cohort	0,611	0,498	0,75	0,000
pancreas	statins_charite_Cohort	0,672	0,596	0,76	0,000
prostate	statins_charite_Cohort	0,737	0,683	0,79	0,000
seq77	statins_charite_Cohort	0,663	0,620	0,71	0,000
seq78	statins_charite_Cohort	0,578	0,542	0,62	0,000
seq79	statins_charite_Cohort	0,621	0,583	0,66	0,000

Supplementary Table 3
1:1 matched Cohort
Mean Relative Risk

stomach	statins_charite_Cohort	0,676	0,586	0,78	0,000
bladder	statins_transatlantic_Cohort	0,915	0,855	0,98	0,010
brain	statins_transatlantic_Cohort	0,696	0,611	0,79	0,000
breast	statins_transatlantic_Cohort	0,613	0,538	0,70	0,000
cancer	statins_transatlantic_Cohort	0,676	0,662	0,69	0,000
colorectal	statins_transatlantic_Cohort	0,626	0,577	0,68	0,000
esophagus	statins_transatlantic_Cohort	0,780	0,681	0,90	0,000
kidney	statins_transatlantic_Cohort	0,895	0,820	0,98	0,014
liver	statins_transatlantic_Cohort	0,540	0,474	0,62	0,000
lung	statins_transatlantic_Cohort	0,820	0,768	0,88	0,000
ovaries	statins_transatlantic_Cohort	0,549	0,455	0,66	0,000
pancreas	statins_transatlantic_Cohort	0,671	0,599	0,75	0,000
prostate	statins_transatlantic_Cohort	0,735	0,686	0,79	0,000
seq77	statins_transatlantic_Cohort	0,637	0,598	0,68	0,000
seq78	statins_transatlantic_Cohort	0,557	0,525	0,59	0,000
seq79	statins_transatlantic_Cohort	0,599	0,565	0,64	0,000
stomach	statins_transatlantic_Cohort	0,677	0,588	0,78	0,000
bladder	statins_UVA_Cohort	0,376	0,257	0,55	0,000
brain	statins_UVA_Cohort	0,545	0,394	0,75	0,000
breast	statins_UVA_Cohort	0,470	0,343	0,64	0,000
cancer	statins_UVA_Cohort	0,521	0,495	0,55	0,000
cancersubentities	statins_UVA_Cohort	0,501	0,464	0,54	0,000
colorectal	statins_UVA_Cohort	0,451	0,342	0,60	0,000
esophagus	statins_UVA_Cohort	0,760	0,551	1,05	0,094
kidney	statins_UVA_Cohort	0,343	0,175	0,68	0,002
liver	statins_UVA_Cohort	0,327	0,166	0,65	0,001
lung	statins_UVA_Cohort	0,682	0,574	0,81	0,000
ovaries	statins_UVA_Cohort	0,436	0,282	0,67	0,000
pancreas	statins_UVA_Cohort	0,512	0,356	0,74	0,000
prostate	statins_UVA_Cohort	0,599	0,497	0,72	0,000
seq77	statins_UVA_Cohort	0,413	0,332	0,51	0,000
seq78	statins_UVA_Cohort	0,433	0,371	0,51	0,000
seq79	statins_UVA_Cohort	0,472	0,407	0,55	0,000
stomach	statins_UVA_Cohort	0,301	0,122	0,74	0,009

Diagnose	Drug	1:1 matched Cohort			
		Relative Risk	Conf-Low	Conf-Up	P-Value
bladder	atorvastatin_charite_Cohort	0,727	0,606	0,87	0,00066
brain	atorvastatin_charite_Cohort	0,600	0,415	0,87	0,00659
breast	atorvastatin_charite_Cohort	0,540	0,368	0,79	0,00174
cancer	atorvastatin_charite_Cohort	0,512	0,480	0,55	0,00000
colorectal	atorvastatin_charite_Cohort	0,450	0,352	0,58	0,00000
esophagus	atorvastatin_charite_Cohort	0,304	0,154	0,60	0,00065
kidney	atorvastatin_charite_Cohort	0,673	0,528	0,86	0,00133
liver	atorvastatin_charite_Cohort	0,306	0,198	0,47	0,00000
lung	atorvastatin_charite_Cohort	0,619	0,508	0,75	0,00000
ovaries	atorvastatin_charite_Cohort	0,333	0,161	0,69	0,00317
pancreas	atorvastatin_charite_Cohort	0,540	0,393	0,74	0,00016
prostate	atorvastatin_charite_Cohort	0,491	0,393	0,61	0,00000
seq77	atorvastatin_charite_Cohort	0,418	0,342	0,51	0,00000
seq78	atorvastatin_charite_Cohort	0,338	0,275	0,42	0,00000
seq79	atorvastatin_charite_Cohort	0,406	0,334	0,49	0,00000
stomach	atorvastatin_charite_Cohort	0,356	0,213	0,60	0,00009
bladder	atorvastatin_transatlantic_Cohort	0,575	0,481	0,69	0,00000
brain	atorvastatin_transatlantic_Cohort	0,506	0,373	0,69	0,00001
breast	atorvastatin_transatlantic_Cohort	0,410	0,296	0,57	0,00000
cancer	atorvastatin_transatlantic_Cohort	0,468	0,446	0,49	0,00000
colorectal	atorvastatin_transatlantic_Cohort	0,389	0,315	0,48	0,00000
esophagus	atorvastatin_transatlantic_Cohort	0,436	0,296	0,64	0,00003
kidney	atorvastatin_transatlantic_Cohort	0,547	0,431	0,69	0,00000
liver	atorvastatin_transatlantic_Cohort	0,272	0,184	0,40	0,00000
lung	atorvastatin_transatlantic_Cohort	0,586	0,500	0,69	0,00000
ovaries	atorvastatin_transatlantic_Cohort	0,287	0,166	0,50	0,00001
pancreas	atorvastatin_transatlantic_Cohort	0,474	0,361	0,62	0,00000
prostate	atorvastatin_transatlantic_Cohort	0,479	0,403	0,57	0,00000
seq77	atorvastatin_transatlantic_Cohort	0,366	0,308	0,44	0,00000
seq78	atorvastatin_transatlantic_Cohort	0,323	0,276	0,38	0,00000
seq79	atorvastatin_transatlantic_Cohort	0,401	0,347	0,46	0,00000
stomach	atorvastatin_transatlantic_Cohort	0,272	0,164	0,45	0,00000
bladder	atorvastatin_UVA_Cohort	0,410	0,252	0,67	0,00036
brain	atorvastatin_UVA_Cohort	0,415	0,244	0,70	0,00117
breast	atorvastatin_UVA_Cohort	0,333	0,189	0,59	0,00016
cancer	atorvastatin_UVA_Cohort	0,449	0,416	0,48	0,00000
colorectal	atorvastatin_UVA_Cohort	0,412	0,281	0,61	0,00001
esophagus	atorvastatin_UVA_Cohort	0,724	0,465	1,13	0,15360
kidney	atorvastatin_UVA_Cohort	0,428	0,197	0,94	0,03239
liver	atorvastatin_UVA_Cohort	0,399	0,182	0,88	0,02183
lung	atorvastatin_UVA_Cohort	0,593	0,458	0,77	0,00009
ovaries	atorvastatin_UVA_Cohort	0,294	0,131	0,66	0,00308
pancreas	atorvastatin_UVA_Cohort	0,502	0,307	0,82	0,00612
prostate	atorvastatin_UVA_Cohort	0,521	0,397	0,68	0,00000
seq77	atorvastatin_UVA_Cohort	0,375	0,275	0,51	0,00000
seq78	atorvastatin_UVA_Cohort	0,356	0,282	0,45	0,00000
seq79	atorvastatin_UVA_Cohort	0,417	0,337	0,52	0,00000
stomach	atorvastatin_UVA_Cohort	0,153	0,023	1,01	0,05112
bladder	fluvastatin_charite_Cohort	0,856	0,558	1,32	0,48671

Supplementary Table 3
1:1 matched Cohort
Median Relative Risk

brain	fluvastatin_charite_Cohort	0,000	0,000	nan	nan
breast	fluvastatin_charite_Cohort	0,472	0,141	1,58	0,22567
cancer	fluvastatin_charite_Cohort	0,649	0,559	0,75	0,00000
colorectal	fluvastatin_charite_Cohort	0,570	0,317	1,03	0,06003
esophagus	fluvastatin_charite_Cohort	1,000	0,537	1,86	1,00000
kidney	fluvastatin_charite_Cohort	1,144	0,811	1,62	0,45259
liver	fluvastatin_charite_Cohort	0,484	0,206	1,14	0,09544
lung	fluvastatin_charite_Cohort	0,591	0,330	1,06	0,07642
ovaries	fluvastatin_charite_Cohort	0,399	0,069	2,31	0,30909
pancreas	fluvastatin_charite_Cohort	0,332	0,094	1,18	0,08713
prostate	fluvastatin_charite_Cohort	0,726	0,462	1,14	0,16575
seq77	fluvastatin_charite_Cohort	0,424	0,245	0,74	0,00228
seq78	fluvastatin_charite_Cohort	0,536	0,361	0,80	0,00204
seq79	fluvastatin_charite_Cohort	0,625	0,426	0,92	0,01616
stomach	fluvastatin_charite_Cohort	0,285	0,046	1,75	0,17729
bladder	fluvastatin_transatlantic_Cohort	0,872	0,570	1,34	0,53944
brain	fluvastatin_transatlantic_Cohort	0,000	0,000	nan	nan
breast	fluvastatin_transatlantic_Cohort	0,500	0,150	1,66	0,26136
cancer	fluvastatin_transatlantic_Cohort	0,679	0,587	0,79	0,00000
colorectal	fluvastatin_transatlantic_Cohort	0,666	0,390	1,14	0,13635
esophagus	fluvastatin_transatlantic_Cohort	1,000	0,537	1,86	1,00000
kidney	fluvastatin_transatlantic_Cohort	1,247	0,903	1,72	0,18191
liver	fluvastatin_transatlantic_Cohort	0,615	0,272	1,39	0,24529
lung	fluvastatin_transatlantic_Cohort	0,639	0,361	1,13	0,12455
ovaries	fluvastatin_transatlantic_Cohort	0,500	0,091	2,52	0,42092
pancreas	fluvastatin_transatlantic_Cohort	0,399	0,116	1,38	0,14618
prostate	fluvastatin_transatlantic_Cohort	0,705	0,447	1,11	0,13286
seq77	fluvastatin_transatlantic_Cohort	0,476	0,277	0,82	0,00715
seq78	fluvastatin_transatlantic_Cohort	0,594	0,403	0,88	0,00845
seq79	fluvastatin_transatlantic_Cohort	0,625	0,426	0,92	0,01616
stomach	fluvastatin_transatlantic_Cohort	0,285	0,046	1,75	0,17729
bladder	fluvastatin_UVA_Cohort	nan	nan	nan	nan
brain	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
breast	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
cancer	fluvastatin_UVA_Cohort	0,500	0,092	2,73	0,43091
colorectal	fluvastatin_UVA_Cohort	2,001	1,898	2,11	0,00000
esophagus	fluvastatin_UVA_Cohort	nan	nan	nan	nan
kidney	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
liver	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
lung	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
ovaries	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
pancreas	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
prostate	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
seq77	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
seq78	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
seq79	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
stomach	fluvastatin_UVA_Cohort	0,000	0,000	nan	nan
bladder	pravastatin_charite_Cohort	1,000	0,720	1,39	1,00000
brain	pravastatin_charite_Cohort	0,444	0,131	1,51	0,19388
breast	pravastatin_charite_Cohort	0,833	0,426	1,63	0,60544
cancer	pravastatin_charite_Cohort	0,794	0,709	0,89	0,00009
colorectal	pravastatin_charite_Cohort	0,393	0,192	0,80	0,01055
esophagus	pravastatin_charite_Cohort	0,631	0,248	1,61	0,34004

Supplementary Table 3

1:1 matched Cohort

Median Relative Risk

kidney	pravastatin_charite_Cohort	0,936	0,607	1,44	0,77709
liver	pravastatin_charite_Cohort	0,124	0,019	0,83	0,03007
lung	pravastatin_charite_Cohort	0,726	0,462	1,14	0,16575
ovaries	pravastatin_charite_Cohort	1,046	0,601	1,82	0,88290
pancreas	pravastatin_charite_Cohort	0,624	0,301	1,29	0,20578
prostate	pravastatin_charite_Cohort	0,968	0,721	1,30	0,83985
seq77	pravastatin_charite_Cohort	0,726	0,527	1,00	0,04967
seq78	pravastatin_charite_Cohort	0,602	0,424	0,85	0,00447
seq79	pravastatin_charite_Cohort	0,579	0,392	0,86	0,00605
stomach	pravastatin_charite_Cohort	0,696	0,316	1,54	0,37531
bladder	pravastatin_transatlantic_Cohort	0,618	0,421	0,91	0,01404
brain	pravastatin_transatlantic_Cohort	0,509	0,255	1,02	0,05510
breast	pravastatin_transatlantic_Cohort	0,579	0,334	1,00	0,05124
cancer	pravastatin_transatlantic_Cohort	0,645	0,591	0,70	0,00000
colorectal	pravastatin_transatlantic_Cohort	0,373	0,224	0,62	0,00017
esophagus	pravastatin_transatlantic_Cohort	0,483	0,206	1,13	0,09414
kidney	pravastatin_transatlantic_Cohort	0,720	0,466	1,11	0,13937
liver	pravastatin_transatlantic_Cohort	0,137	0,036	0,52	0,00362
lung	pravastatin_transatlantic_Cohort	0,697	0,508	0,96	0,02526
ovaries	pravastatin_transatlantic_Cohort	0,856	0,522	1,41	0,54936
pancreas	pravastatin_transatlantic_Cohort	0,498	0,273	0,91	0,02278
prostate	pravastatin_transatlantic_Cohort	0,900	0,716	1,13	0,37260
seq77	pravastatin_transatlantic_Cohort	0,491	0,354	0,68	0,00003
seq78	pravastatin_transatlantic_Cohort	0,487	0,372	0,64	0,00000
seq79	pravastatin_transatlantic_Cohort	0,475	0,356	0,64	0,00000
stomach	pravastatin_transatlantic_Cohort	0,443	0,186	1,05	0,06493
bladder	pravastatin_UVA_Cohort	0,000	0,000	nan	nan
brain	pravastatin_UVA_Cohort	0,568	0,248	1,30	0,18243
breast	pravastatin_UVA_Cohort	0,497	0,212	1,16	0,10739
cancer	pravastatin_UVA_Cohort	0,558	0,488	0,64	0,00000
colorectal	pravastatin_UVA_Cohort	0,554	0,280	1,10	0,08948
esophagus	pravastatin_UVA_Cohort	0,330	0,055	1,98	0,22732
kidney	pravastatin_UVA_Cohort	0,615	0,194	1,95	0,41655
liver	pravastatin_UVA_Cohort	0,396	0,068	2,30	0,30647
lung	pravastatin_UVA_Cohort	0,784	0,512	1,20	0,26634
ovaries	pravastatin_UVA_Cohort	0,519	0,196	1,38	0,18810
pancreas	pravastatin_UVA_Cohort	0,541	0,205	1,43	0,21604
prostate	pravastatin_UVA_Cohort	0,887	0,626	1,26	0,50939
seq77	pravastatin_UVA_Cohort	0,165	0,056	0,49	0,00115
seq78	pravastatin_UVA_Cohort	0,486	0,325	0,73	0,00047
seq79	pravastatin_UVA_Cohort	0,453	0,298	0,69	0,00023
stomach	pravastatin_UVA_Cohort	0,000	0,000	nan	nan
bladder	simvastatin_charite_Cohort	0,962	0,892	1,04	0,31797
brain	simvastatin_charite_Cohort	0,807	0,692	0,94	0,00609
breast	simvastatin_charite_Cohort	0,685	0,583	0,80	0,00001
cancer	simvastatin_charite_Cohort	0,766	0,747	0,79	0,00000
colorectal	simvastatin_charite_Cohort	0,682	0,623	0,75	0,00000
esophagus	simvastatin_charite_Cohort	0,835	0,712	0,98	0,02670
kidney	simvastatin_charite_Cohort	0,909	0,822	1,00	0,06109
liver	simvastatin_charite_Cohort	0,593	0,514	0,68	0,00000
lung	simvastatin_charite_Cohort	0,887	0,822	0,96	0,00219
ovaries	simvastatin_charite_Cohort	0,631	0,504	0,79	0,00007
pancreas	simvastatin_charite_Cohort	0,728	0,640	0,83	0,00000

Supplementary Table 3
1:1 matched Cohort
Median Relative Risk

prostate	simvastatin_charite_Cohort	0,776	0,714	0,84	0,00000
seq77	simvastatin_charite_Cohort	0,708	0,659	0,76	0,00000
seq78	simvastatin_charite_Cohort	0,628	0,586	0,67	0,00000
seq79	simvastatin_charite_Cohort	0,661	0,617	0,71	0,00000
stomach	simvastatin_charite_Cohort	0,758	0,654	0,88	0,00029
bladder	simvastatin_transatlantic_Cohort	1,026	0,956	1,10	0,49152
brain	simvastatin_transatlantic_Cohort	0,797	0,689	0,92	0,00240
breast	simvastatin_transatlantic_Cohort	0,705	0,608	0,82	0,00001
cancer	simvastatin_transatlantic_Cohort	0,780	0,762	0,80	0,00000
colorectal	simvastatin_transatlantic_Cohort	0,754	0,690	0,82	0,00000
esophagus	simvastatin_transatlantic_Cohort	0,883	0,759	1,03	0,10658
kidney	simvastatin_transatlantic_Cohort	1,015	0,922	1,12	0,77260
liver	simvastatin_transatlantic_Cohort	0,682	0,595	0,78	0,00000
lung	simvastatin_transatlantic_Cohort	0,942	0,877	1,01	0,10404
ovaries	simvastatin_transatlantic_Cohort	0,634	0,512	0,79	0,00003
pancreas	simvastatin_transatlantic_Cohort	0,792	0,700	0,90	0,00025
prostate	simvastatin_transatlantic_Cohort	0,801	0,740	0,87	0,00000
seq77	simvastatin_transatlantic_Cohort	0,758	0,708	0,81	0,00000
seq78	simvastatin_transatlantic_Cohort	0,653	0,612	0,70	0,00000
seq79	simvastatin_transatlantic_Cohort	0,679	0,634	0,73	0,00000
stomach	simvastatin_transatlantic_Cohort	0,838	0,727	0,97	0,01471
bladder	simvastatin_UVA_Cohort	0,365	0,177	0,75	0,00643
brain	simvastatin_UVA_Cohort	0,715	0,422	1,21	0,21364
breast	simvastatin_UVA_Cohort	0,677	0,441	1,04	0,07366
cancer	simvastatin_UVA_Cohort	0,603	0,552	0,66	0,00000
colorectal	simvastatin_UVA_Cohort	0,426	0,246	0,74	0,00244
esophagus	simvastatin_UVA_Cohort	0,719	0,377	1,37	0,32152
kidney	simvastatin_UVA_Cohort	0,000	0,000	nan	nan
liver	simvastatin_UVA_Cohort	0,232	0,037	1,46	0,11928
lung	simvastatin_UVA_Cohort	0,752	0,557	1,02	0,06308
ovaries	simvastatin_UVA_Cohort	0,526	0,265	1,05	0,06660
pancreas	simvastatin_UVA_Cohort	0,549	0,277	1,09	0,08564
prostate	simvastatin_UVA_Cohort	0,539	0,362	0,80	0,00234
seq77	simvastatin_UVA_Cohort	0,538	0,382	0,76	0,00042
seq78	simvastatin_UVA_Cohort	0,498	0,381	0,65	0,00000
seq79	simvastatin_UVA_Cohort	0,504	0,384	0,66	0,00000
stomach	simvastatin_UVA_Cohort	0,658	0,259	1,67	0,38583
bladder	statins_charite_Cohort	0,944	0,881	1,01	0,10547
brain	statins_charite_Cohort	0,741	0,643	0,86	0,00004
breast	statins_charite_Cohort	0,650	0,562	0,75	0,00000
cancer	statins_charite_Cohort	0,710	0,694	0,73	0,00000
cancersubentities	statins_charite_Cohort	0,728	0,708	0,75	0,00000
colorectal	statins_charite_Cohort	0,629	0,579	0,69	0,00000
esophagus	statins_charite_Cohort	0,743	0,637	0,87	0,00016
kidney	statins_charite_Cohort	0,901	0,823	0,99	0,02354
liver	statins_charite_Cohort	0,532	0,466	0,61	0,00000
lung	statins_charite_Cohort	0,837	0,779	0,90	0,00000
ovaries	statins_charite_Cohort	0,607	0,494	0,75	0,00000
pancreas	statins_charite_Cohort	0,675	0,599	0,76	0,00000
prostate	statins_charite_Cohort	0,737	0,683	0,79	0,00000
seq77	statins_charite_Cohort	0,662	0,620	0,71	0,00000
seq78	statins_charite_Cohort	0,578	0,543	0,62	0,00000
seq79	statins_charite_Cohort	0,619	0,580	0,66	0,00000

Supplementary Table 3
1:1 matched Cohort
Median Relative Risk

stomach	statins_charite_Cohort	0,673	0,583	0,78	0,00000
bladder	statins_transatlantic_Cohort	0,910	0,850	0,97	0,00665
brain	statins_transatlantic_Cohort	0,698	0,613	0,80	0,00000
breast	statins_transatlantic_Cohort	0,610	0,534	0,70	0,00000
cancer	statins_transatlantic_Cohort	0,675	0,661	0,69	0,00000
colorectal	statins_transatlantic_Cohort	0,622	0,573	0,67	0,00000
esophagus	statins_transatlantic_Cohort	0,786	0,685	0,90	0,00061
kidney	statins_transatlantic_Cohort	0,893	0,817	0,98	0,01255
liver	statins_transatlantic_Cohort	0,540	0,474	0,62	0,00000
lung	statins_transatlantic_Cohort	0,817	0,764	0,87	0,00000
ovaries	statins_transatlantic_Cohort	0,547	0,454	0,66	0,00000
pancreas	statins_transatlantic_Cohort	0,668	0,596	0,75	0,00000
prostate	statins_transatlantic_Cohort	0,730	0,681	0,78	0,00000
seq77	statins_transatlantic_Cohort	0,635	0,595	0,68	0,00000
seq78	statins_transatlantic_Cohort	0,558	0,526	0,59	0,00000
seq79	statins_transatlantic_Cohort	0,599	0,565	0,64	0,00000
stomach	statins_transatlantic_Cohort	0,676	0,587	0,78	0,00000
bladder	statins_UVA_Cohort	0,376	0,257	0,55	0,00000
brain	statins_UVA_Cohort	0,532	0,384	0,74	0,00017
breast	statins_UVA_Cohort	0,473	0,346	0,65	0,00000
cancer	statins_UVA_Cohort	0,522	0,496	0,55	0,00000
cancersubentities	statins_UVA_Cohort	0,498	0,461	0,54	0,00000
colorectal	statins_UVA_Cohort	0,448	0,339	0,59	0,00000
esophagus	statins_UVA_Cohort	0,762	0,552	1,05	0,09826
kidney	statins_UVA_Cohort	0,332	0,169	0,66	0,00147
liver	statins_UVA_Cohort	0,309	0,156	0,61	0,00079
lung	statins_UVA_Cohort	0,679	0,571	0,81	0,00002
ovaries	statins_UVA_Cohort	0,427	0,276	0,66	0,00015
pancreas	statins_UVA_Cohort	0,507	0,352	0,73	0,00029
prostate	statins_UVA_Cohort	0,611	0,506	0,74	0,00000
seq77	statins_UVA_Cohort	0,409	0,330	0,51	0,00000
seq78	statins_UVA_Cohort	0,432	0,371	0,51	0,00000
seq79	statins_UVA_Cohort	0,475	0,410	0,55	0,00000
stomach	statins_UVA_Cohort	0,277	0,112	0,69	0,00572

Supplementary Table 4

This spreadsheet supplies the results of our analyses of co-medications prescribed with statins. The first tab “Co-medication Counts” provides a list of drugs prescribed together with statins. The total number of counts of those prescriptions is provided (in descending order) for the top 10 comedications. The second tab “Co-medication General” consists of the results (odds-ratios [ORs]) for those 10 drugs provided in tab 1. The ORs are calculated for the transatlantic cohort as well as the 1:1 matched cohort design study. Besides the ORs, p-values are also provided. Drugs with an OR < 1 are highlighted in yellow. The third tab “Exclusive Drug Calculation” consists of the analysis of previously identified drugs (tab 2) with an OR < 1. Those co-medications were analyzed separately by excluding patients taking at least two drugs identified in tab 2. These analyses were performed for the transatlantic as well as the 1:1 matched cohort. In addition, we analyzed the two most prominent statins (simvastatin and atorvastatin) separately to point out differences between statins. As can be seen in the tables, atorvastatin outperformed simvastatin with respect to the ORs in the transatlantic as well as 1:1 matched cohort

Co-medication	
NAME	Drugs prescribed with statins
STATINS	42.682
1 Aspirin	26.800
2 Metoprolol	16.740
3 Pantoprazole	13.652
4 Ramipril	11.179
5 Amlodipine	7.636
6 Torsemide	6.512
7 Levothyroxine sodium	5.394
8 hydrochlorothiazide	5.372
9 Acetaminophen	5.358
10 Furosemide	5.198

ComedicationGeneral

ODDS-RATIO analysis for co-medication

		TRANSATLANTIC COHORT										
Diagnose	Drug	NbDrug	NbDiag	NbDiag/Diag	AllRatio	Conf-Lower	Conf-Upper	P-Value	RelativRisk	Conf-Upper	Conf-Lower	P-Value
cancer	statins	42682	53113	6426	0,716	0,7	0,74	2,1139E-108	0,759	0,777	0,741	1,2698E-101
cancer	aspirin	57943	53158	8305	0,661	0,64	0,68	1,5790E-138	0,71	0,725	0,694	8,0867E-181
cancer	metoprolol	41414	53338	7833	0,977	0,95	1	7,4983E-02	0,981	1,003	0,96	8,5809E-02
cancer	pantoprazole	79764	55830	20265	1,607	1,58	1,64	0,0000E+00	1,453	1,475	1,431	0,0000E+00
cancer	ramipril	27732	53726	5829	1,106	1,07	1,14	1,1013E-09	1,084	1,11	1,058	1,2657E-10
cancer	amlodipine	22105	53466	4545	1,083	1,05	1,12	1,7966E-06	1,066	1,095	1,038	3,7359E-06
cancer	torsemide	16416	53791	4220	1,453	1,4	1,51	1,6900E-74	1,336	1,373	1,301	1,3900E-87
cancer	levothyroxine	21138	53350	4890	1,267	1,22	1,31	1,7838E-35	1,205	1,236	1,174	1,3360E-41
cancer	hydrochlorothiazide	14207	53549	2854	1,037	0,99	1,08	1,0146E-01	1,03	1,065	0,995	8,8010E-02
cancer	acetaminophen	50920	53467	9971	1,04	1,01	1,07	7,7279E-03	1,032	1,052	1,012	1,4967E-03
cancer	furosemide	17553	53126	2789	0,78	0,75	0,81	1,3308E-33	0,815	0,844	0,787	4,6544E-28

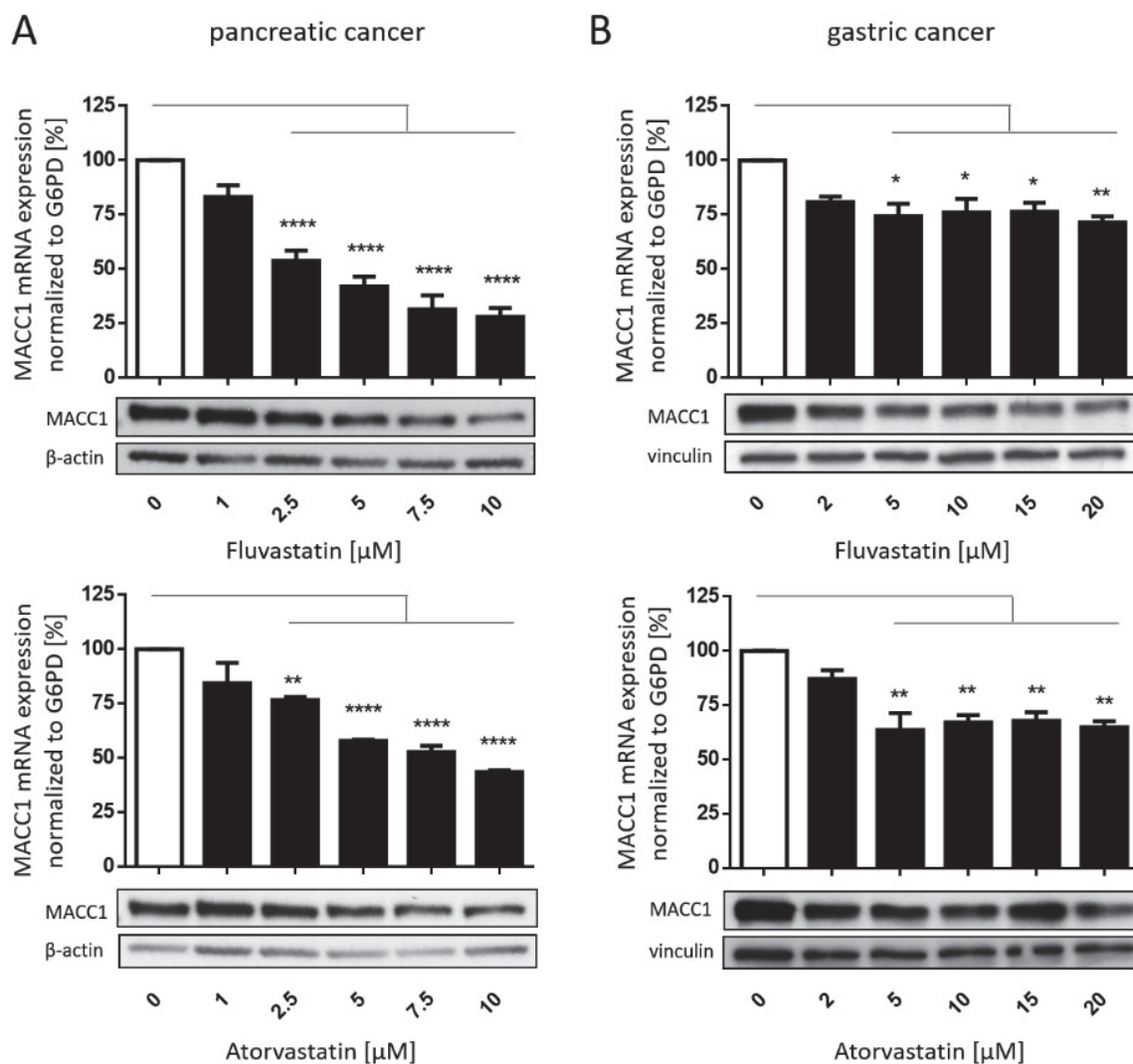
		1:1 MATCHED COHORT										
Diagnose	Drug	NbDrug	NbDiag	NbDiag/Drug	MatchedRatio	Conf-Lower	Conf-Upper	P-Value				
cancer	statins	17859	53113	6425	0,497	0,481	0,514	0,0000E+00				
cancer	aspirin	23299	53158	8304	0,474	0,46	0,489	0,0000E+00				
cancer	metoprolol	18062	53338	7833	0,72	0,697	0,743	2,3898E-80				
cancer	pantoprazole	36013	55830	20264	1,449	1,413	1,486	3,3304E-160				
cancer	ramipril	12719	53726	5829	0,824	0,794	0,855	6,5749E-23				
cancer	amlodipine	10018	53466	4545	0,827	0,793	0,861	2,4136E-18				
cancer	torsemide	8069	53791	4220	1,093	1,044	1,143	1,3400E-04				
cancer	levothyroxine	9506	53350	4890	1,061	1,017	1,106	5,6948E-03				
cancer	hydrochlorothiazide	6365	53549	2854	0,803	0,763	0,844	1,6515E-16				
cancer	acetaminophen	19907	53467	9970	1,009	0,978	1,041	5,8563E-01				
cancer	furosemide	6903	53126	2789	0,639	0,608	0,671	1,4929E-63				

Supplementary Table 4
Odds-Ratios for co-medication

ExclusiveDrugCalculation

ANALYSIS OF CO-MEDICATION														
ANALYSIS OF STATINS AS A GROUP														
					Transatlantic Study					Matched Transatlantic Study				
	Statins	Aspirin	Furosemide	Cancer	Drug Cancer	& Odds Ratio	Lower Bound	Upper Bound	P-Value	Odds Ratio	Lower Bound	Upper Bound	P-Value	
Statins	Yes	No	No		53227	2635	1,025	0,98	1,07	0,2734	0,756	0,718	0,797	<0.0001
Aspirin	No	Yes	No		53245	4332	0,821	0,79	0,85	<0.0001	0,655	0,629	0,683	<0.0001
Furosemide	No	No	Yes		53175	1806	1,154	1,09	1,22	<0.0001	1,057	0,988	1,13	0,1055
Exclusion of co-medication with positive cancer preventive effects														
ANALYSIS OF SEPARATE STATINS														
					Transatlantic Study					Matched Transatlantic Study				
	Simvastatin	Atorvastatin	Aspirin	Furosemide	Cancer	Drug Cancer	& Odds Ratio	Lower Bound	Upper Bound	P-Value	Odds Ratio	Lower Bound	Upper Bound	P-Value
Simvastatin	Yes	No	No	No	53280	1984	1,191	1,13	1,25	<0.0001	0,871	0,819	0,926	<0.0001
Atorvastatin	No	Yes	No	No	53016	403	0,634	0,57	0,7	<0.0001	0,475	0,422	0,536	<0.0001
Aspirin	No	No	Yes	No	53245	4332	0,821	0,79	0,85	<0.0001	0,655	0,629	0,683	<0.0001
Furosemide	No	No	No	Yes	53175	1806	1,154	1,09	1,22	<0.0001	1,057	0,988	1,13	0,1055
Exclusion of co-medication with positive cancer preventive effects														

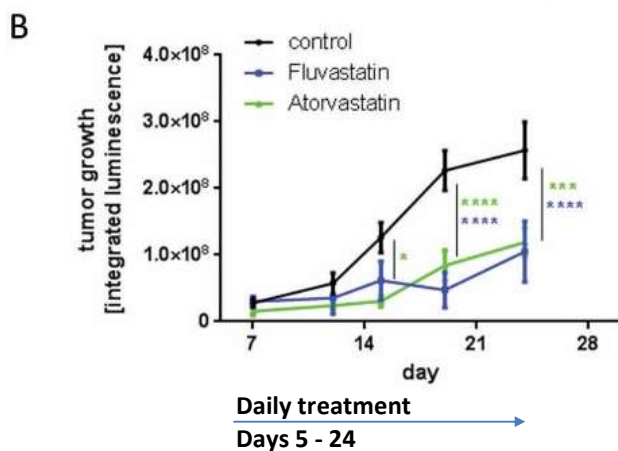
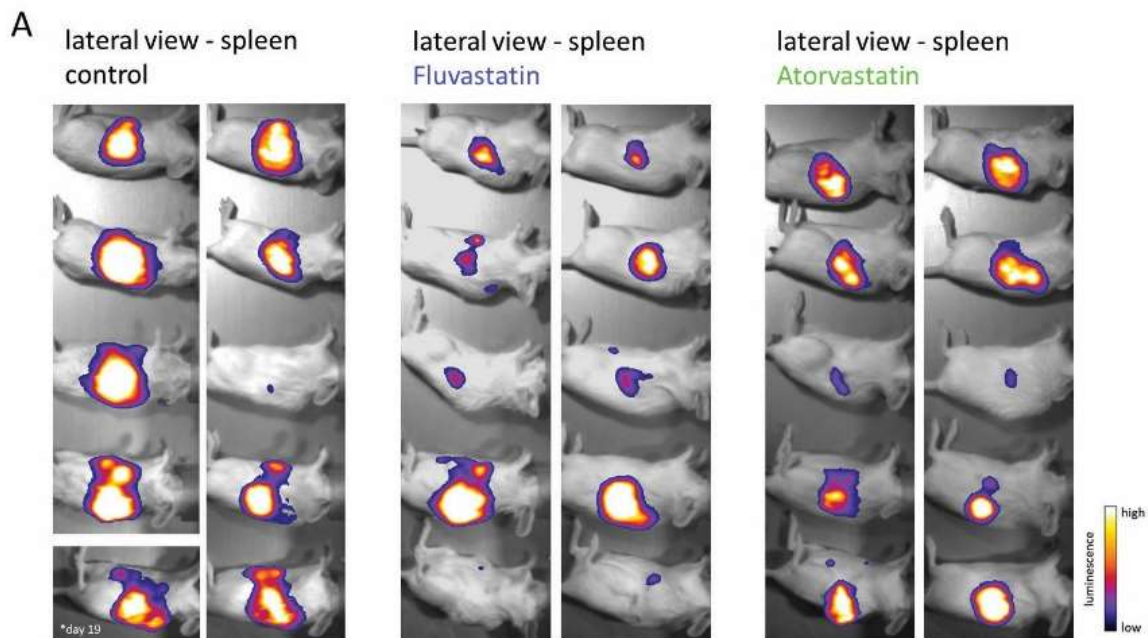
Supplementary Table 4
Analysis of Co-medication



Supplementary Figure 1.

Statins reduce MACC1 expression in different cancer entities.

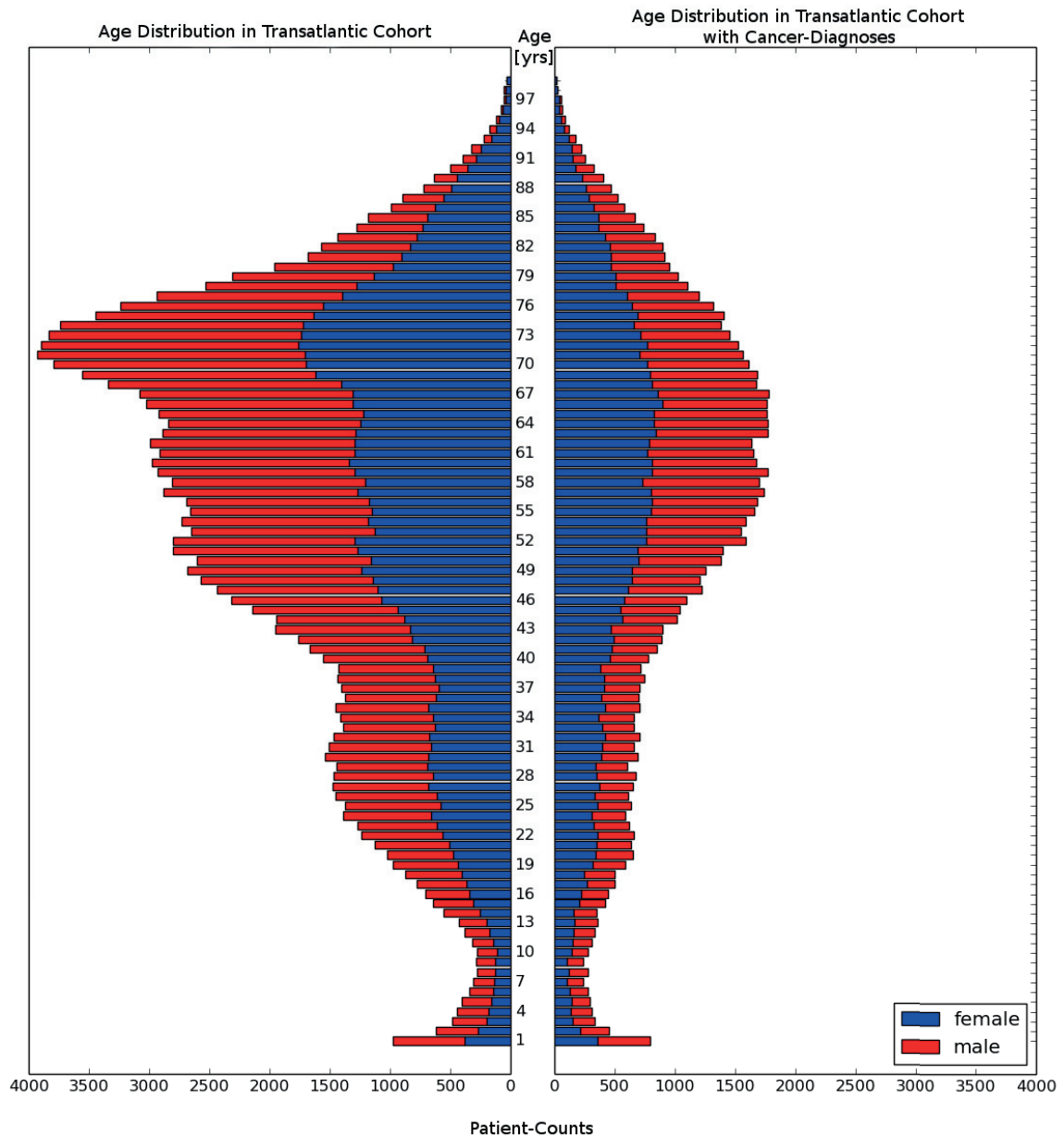
Fluvastatin and atorvastatin decreased MACC1 mRNA and protein expression in pancreatic cancer (BxPC3; Panel A) and gastric cancer (MKN45; Panel B) cells. MACC1 mRNA levels were normalized to G6PD mRNA expression and respective treatment controls (DMSO, indicated with white bars). Results for mRNA represent means + SEMs of three independent experiments, and for WB one representative example of at least two independent experiments is shown. In the WB, β -actin or vinculin served as loading control. Significant results were determined by one-way ANOVA and Dunnett's multiple comparison test with a confidence interval of 95% (* = $p < 0.05$, ** = $p < 0.01$, *** = $p < 0.001$, **** = $p < 0.0001$).



Supplementary Figure 2.

Statin treatment decreases tumor burden in vivo.

Intra-splenically xenografted (HCT116/CMVp-Luc cells) SCID-beige mice were treated either with solvent or daily doses of 13 mg/kg body weight fluvastatin or atorvastatin as indicated. Bioluminescent imaging of animals from a lateral view (A), at day 24 of statin treatment, showed significantly weaker signals, indicating restricted tumor burden in the spleen. Bioluminescent signals from all ten animals over the time course of the experiment were quantified (B). Results represent means \pm SEM and significant results were determined by two-way ANOVA and Dunnett's multiple comparison test with a 95% CI (* = $p < 0.05$, ** = $p < 0.01$, *** = $p < 0.001$, **** = $p < 0.0001$). Data were analyzed by ANOVA and Dunnett's multiple comparison post tests or two-way ANOVA including Tukey's additivity test.

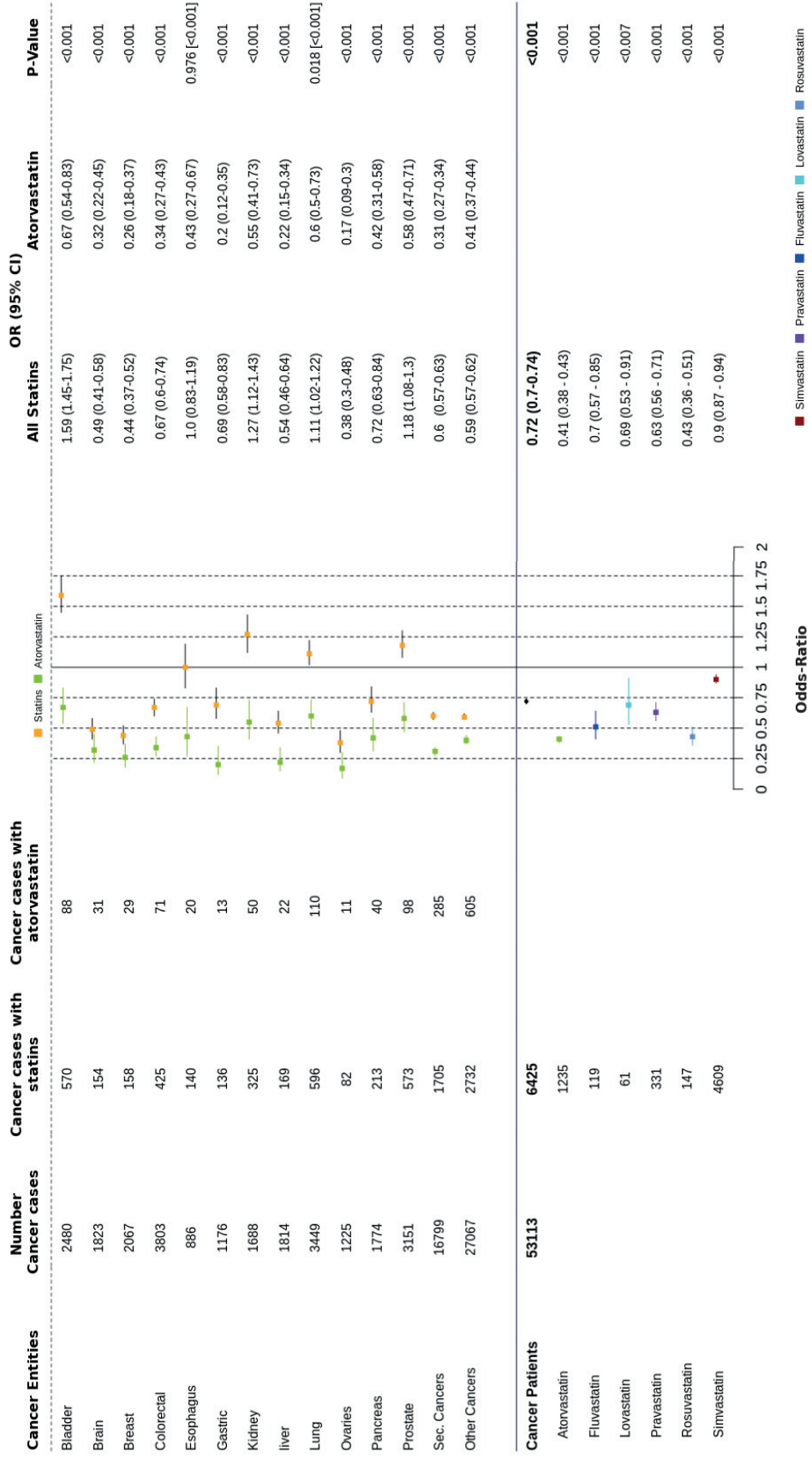


Supplementary Figure 3.

Age and gender distribution in the trans-Atlantic cohort and patients diagnosed with cancer.

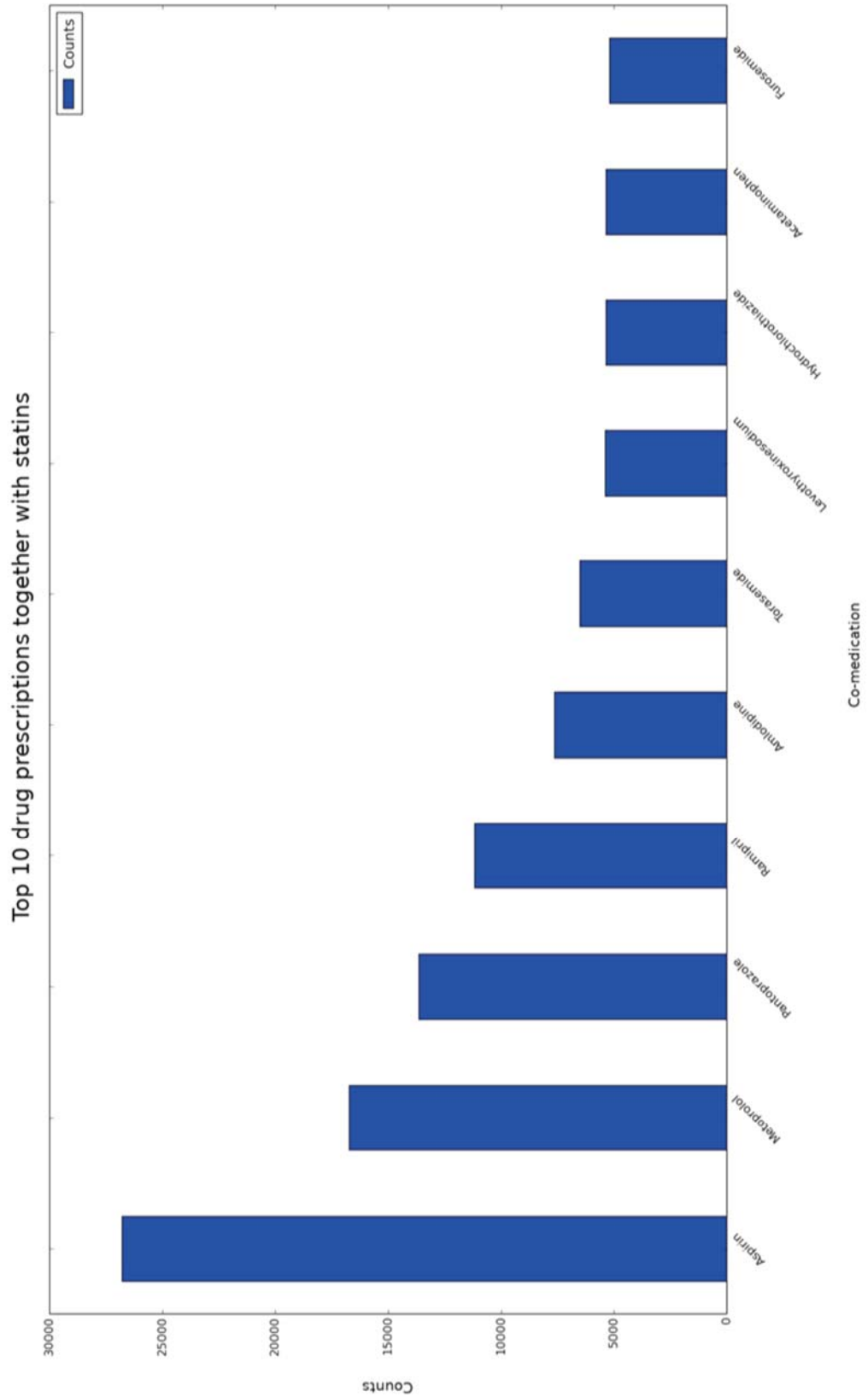
The age distribution for both genders in the full trans-Atlantic cohort is shown (left half), along with the analogous distribution for patients diagnosed with cancer (right); any effects that might stem from the visible differences between these two groups were mitigated by using a 1:1 matched study design.

Transatlantic Cohort



Supplementary Figure 4.

Cancer preventive effect of different statins and different cancers across the transatlantic cohort. The cancer preventive effect of statins as a group, and atorvastatin alone, were calculated for a number of different cancer types. A detailed overview of cancer incidences, cancer diagnoses and prescribed statins is provided. The cancer preventive effect is calculated as an odds-ratios for both, statins and atorvastatin. The 95% confidence intervals p-values are provided. P-values in square brackets are provided for atorvastatin, for any values that differ from the statins at large. An overview of results for all statins prescribed in the study population is presented in the bottom of the figure.



Supplementary Figure 5.

Co-medication with statins. The medications prescribed alongside statins are shown, in decreasing order of the total number of prescriptions of each