

**Supplementary Table 1. PubMed IDs for sex effects on outcome risk reported in Figure 2**

<b>Index Condition – Outcome Pair</b>	<b>First Author</b>	<b>Publication Year</b>	<b>PubMed ID</b>
<b>CHF – Mortality</b>	Bouvy <sup>1</sup>	2003	12748212
	Adlam <sup>2</sup>	2005	15769783
	Alehagen <sup>3</sup>	2005	15820281
	Pocock <sup>4</sup>	2006	16219658
	Levy <sup>5</sup>	2006	16534009
	Rassi <sup>6</sup>	2006	16928995
	Miura <sup>7</sup>	2008	18296827
	O'Connor <sup>8</sup>	2012	22114101
<b>Revascularization – Mortality</b>	Hannan <sup>9</sup>	1992	1433740
	O'Connor <sup>10</sup>	1992	1591830
	Wong <sup>11</sup>	1999	10519495
	Moscucci <sup>12</sup>	2001	11457742
	Zaroff <sup>13</sup>	2002	12088769
	Wu <sup>14</sup>	2004	14752428
	Wu <sup>15</sup>	2006	16458151
	Hannan <sup>16</sup>	2006	16458152
	Mehta <sup>17</sup>	2008	18250266
	de Mulder <sup>18</sup>	2011	21345854
	Hamburger <sup>19</sup>	2009	19681116
<b>ACS – Mortality</b>	Grzybowski <sup>20</sup>	2000	11265730
	Zaroff <sup>13</sup>	2002	12088769
	Williams <sup>21</sup>	2006	16498154
	de Boer <sup>22</sup>	2011	21392604
	Boersma <sup>23</sup>	2000	10840005
	Mehran <sup>24</sup>	2010	20513595
	Roe <sup>25</sup>	2011	22093204

**Supplementary Table 2.** Study- and model-level characteristics of CVD-related clinical prediction models including sex effects

PubMed ID	First Author	Pub. Year	Inclusion of Sex	Effect of Female Sex	Population	Outcome(s)	Covariates	Cohort Sample Size	% Female in Cohort	No. of Events	Mean Age (SD)	Follow-Up Duration
<b>CAD – MORTALITY</b>												
1433740	Hannan <sup>9</sup>	1992	Covariate	Harmful	Patients undergoing percutaneous transluminal coronary angioplasty	In-hospital mortality	Shock, Hemodynamic myocardial infarction c instability, Sex, LVEF	5827	29.6%	37	NR	Hospitalization period
1591830	O'Connor <sup>10</sup>	1992	Covariate	Harmful	Patients undergoing isolated coronary artery bypass grafting	In-hospital mortality	Prior CABG, Priority at Surgery, Comorbidities, Sex, Left Ventricular End Diastolic Pressure, Ejection Fraction Score, Age, Bovine Serum Albumin	3055	NR	132	NR	Hospitalization period
10519495	Wong <sup>11</sup>	1999	Covariate	Harmful	Patients undergoing CABG surgery who received fast-track cardiac anesthesia (FTCA) technique, consisting of low-dose fentanyl (10-15 mg/kg) and propofol infusion	Mortality	Left Ventricle Grade 4, Emergency Surgery, Sex	885	24%	23	60.8 (11.3)	30 days
10840005	Boersma <sup>23</sup>	2000	Covariate	Protective	Patients presenting within 24 hours of an episode of ischemic chest pain (>10 min) either transient ST-segment elevation (>0.5 mm), transient or persistent ST-segment depression (>0.5 mm), T-wave inversion (>1.0 mm), or elevation of creatinine kinase	Mortality	Age, Signs of Heart Failure (Rales), ST-Depression on EKG, HR, SBP, CCS Class, Sex	9461	35%	342	64 (NR)	30 days
11265730	Grzybowski <sup>20</sup>	2000	Covariate	Harmful	Patients with suspected AMI transported by emergency medical services	Mortality	Age, Respiratory Rate, SBP, Sex	244	39.8%	36	66.6 (14.8)	7 days
11457742	Moscucci <sup>12</sup>	2001	Covariate	Harmful	Patients undergoing percutaneous coronary interventions	In-hospital mortality	Shock, Creatinine, Cardiac Arrest, MI, Age, Sex, Thrombus, LVEF, PVD, Number of Diseased Vessels	10729	31.8%	168	62.6 (11.9)	Hospitalization period

11686666	Marchioli <sup>26</sup>	2001	Stratified	NA	Recent MI and without unfavorable short-term prognosis	Mortality	Age, Left Ventricular Dysfunction, Leukocytes, Claudication, Cholesterol, Fibrinogen, Smoking, HR, Electrical Instability, Diabetes, Arterial HTN, Residual Myocardial Ischemia	11248	14.6%	M: 904 F: 167	59 (11)	4 years
12088769	Zaroff <sup>13</sup>	2002	Covariate	Harmful	Patients with acute MI who underwent CABG	In-hospital mortality	Killip Class, Age, Previous CABG, Q-wave vs. non-Q-wave Acute MI, Sex, CHF, Diabetes, Weight, Previous Acute MI	71774	29.6%	3948	NR	Hospitalization period
14752428	Wu <sup>14</sup>	2004	Covariate	Harmful	Patients undergoing coronary artery bypass grafting (CABG)	Operative mortality	Previous Cardiac Surgery, Renal Insufficiency, Chronic Lung Disease, Urgency, LVEF, Cerebrovascular Disease, History of PVD, Age, Canadian Cardiovascular Society Class, Sex, Left Main Stenosis, NYHA Class	11063	24%	255	65 (11)	Hospitalization period
15135693	Elhendy <sup>27</sup>	2004	Covariate	Protective	Patients with known or suspected CAD	All-cause mortality	Exercise Wall Motion Score Index, Age, Sex, Metabolic Equivalents	2840	42%	151	62 (12)	4 years (median: 3 years)
16098306	Cortigiani <sup>28</sup>	2005	Covariate	Protective	Patients with known or suspected coronary artery disease who underwent dipyridamole stress echocardiography	All-cause mortality	Age, Peak Wall Motion Score Index, Diabetes, Sex	3969	32%	324	60 (NR)	median: 48 months
16210253	Clayton <sup>29</sup>	2005	Covariate	Protective	Patients with stable symptomatic angina requiring treatment, preserved left ventricular function, and either previous MI or proved angiographic CAD	Mortality	Diabetes, Age, Smoking, QT Interval, Lipid Lowering Therapy, Angina, LVEF, Sex, SBP, Previous Angiography, Previous MI, Glucose, Angina Medication, Creatinine, WBC	7311	20.6%	569	63.5 (9.2)	mean: 4.9 years
16427855	Rexius <sup>30</sup>	2006	Covariate	Protective	Patients accepted for elective coronary artery bypass grafting (CABG) awaiting surgery	Pre-surgical mortality	UA, Left Main Stenosis, LVEF, Perioperative Risk, Aortic Valve Disease Requiring Surgery, Sex	5167	24%	42	66 (9)	median: 47 days
16458151	Wu <sup>15</sup>	2006	Covariate	Harmful	Patients in population-based percutaneous coronary intervention (PCI) registry	In-hospital mortality	Hemodyn Acute MI C Static, MI with Stent Thrombosis, Renal Failure, CHF, LVEF, Peripheral Arterial	46090	32.1%	321	NR	Hospitalization period

							Disease, Sex, Age					
16458152	Hannan <sup>16</sup>	2006	Covariate	Harmful	Patients who underwent isolated CABG surgery	In-hospital mortality	MI, Hemodynamically unstable, Renal Failure, LVEF, Previous Open Heart Operations, Sex, Extensively Calcified Ascending Aorta, Peripheral Arterial Disease, COPD, Age	16120	NR	366	NR	Hospitalization period
16498154	Williams <sup>21</sup>	2006	Covariate	Harmful	Patients with acute MI	Mortality	Killip Class, ST Depression, SBP, Sex, Age, Creatinine, QRS Duration, MI Location	809	60%	87	70.7 (NR)	30 days
17223413	Anderson <sup>31</sup>	2007	Covariate	Protective	Patients who underwent coronary angiography	All-cause mortality	WBC, Age, Hematocrit, MCV, Red Cell Distribution, MCH, Platelet Count, Sex	17787	38%	709	61.1 (14.7)	30 days (mean: 4.9 years)
17223413	Anderson <sup>31</sup>	2007	Covariate	Protective	Patients who underwent coronary angiography	All-cause mortality	Age, Red Cell Distribution, MCH, Hematocrit, WBC, Platelet Count, MCV, Sex	17787	38%	709	61.1 (14.7)	30 days (mean: 4.9 years)
17499864	Yap <sup>32</sup>	2008	Covariate	Protective	Patients with recent documented acute MI with at least one high-risk feature	All-cause mortality	Age, Diabetes, HR, HTN, NYHA Class, Non-Q Wave Infarct on EKG, Previous MI or Angina, Sex, SBP	2707	21%	480	NR	2 years
17499864	Yap <sup>32</sup>	2008	Covariate	Protective	Patients with recent documented acute MI with at least one high-risk feature	Arrhythmic cardiac mortality	Age, Diabetes, HR, HTN, NYHA Class, Non-Q Wave Infarct on EKG, Previous MI or Angina, Sex, SBP	2707	21%	480	NR	2 years
17499864	Yap <sup>32</sup>	2008	Covariate	Protective	Patients with recent documented acute MI with at least one high-risk feature	Non-arrhythmic cardiac mortality	Age, Diabetes, HR, HTN, NYHA Class, Non-Q Wave Infarct on EKG, Previous MI or Angina, Sex, SBP	2707	21%	480	NR	2 years
17550750	Singh <sup>33</sup>	2007	Covariate	Harmful	Patients undergoing PCI	Mortality	Age, LVEF, Creatinine, Shock, MI, CHF, Peripheral Artery Disease, Diabetes, Sex	7457	31%	136	66.9 (12.1)	Hospitalization period
18250266	Mehta <sup>17</sup>	2008	Covariate	Harmful	Patients with and without cardiogenic shock undergoing CABG	Post-operative in-hospital mortality	CABG and Ventricular Septal Repair, Creatinine, Operative Status, Prior Cardiac Surgery, CABG and Aortic Valve, CABG and Mitral Valve, Immunosuppressive Therapy, Sex, LVEF, IABP, Preoperative Resuscitation, MI, Age	708593	28.9%	24092	NR	Hospitalization period
19559824	Shahian <sup>34</sup>	2009	Covariate	Harmful	Patients who underwent single aortic or mitral valve surgical	Mortality	BSA, Dialysis, Priority Status, Reoperation, Endocarditis, Shock,	101661	35.5%	6919	NR	30 days

					procedures combined with coronary artery bypass grafting		Sex x BSA, MI, Creatinine, CHF and NYHA IV, IABP or Inotropes, Sex, Immunosuppressive Treatment, Diabetes, PVD, Tricuspid Insufficiency, CHF without NYHA IV, CVD and Prior CVA, AF, CLD Function, MVR, Number of Diseased Vessels, Left Main Disease, UA, Mitral Stenosis, Age, HTN, Race, Mitral Valve Repair					
19681116	Hamburger <sup>19</sup>	2009	Covariate	Protective	Population-based registry of patients who underwent PCI in British Columbia	30-day mortality	STEMI, Critical Preprocedural State, LVEF, ACS, Left Main Disease, Emergency, NYHA Class, Dialysis or Creatinine, Triple-Vessel Disease, Age, Sex	26350	26.8%	500	64.3 (11.5)	30 days
20513595	Mehran <sup>24</sup>	2010	Covariate	Protective	Patients with ACS including NSTEMI, STEMI, and biomarker negative ACS	Mortality	Diabetes, Biomarkers, Smoking, Age, STEMI, Previous MI, WBC, Creatinine, Sex, UFH/Enox + GPI, Glycoprotein Inhibitor, Bivalirudin, Hemoglobin	17421	28.7%	660	62.1 (11.7)	1 year
20923988	Singh <sup>35</sup>	2010	Covariate	Protective	Patients who underwent PCI	All-cause mortality	Age, BMI, CAD Comorbidity Index, LVEF, History of CHF, MEDS, Troponin T, Ventricular Arrhythmia, Prior MI, Smoking Status, Cardiogenic Shock, Prior CABG, Sex, Stable Angina, Procedural Failure, Prior PCI, Three-Vessel Disease, UA, Stent Use, Glycoprotein Use, Cholesterol	9165	29%	1243	66.8 (12.2)	Hospitalization period
21345854	De Mulder <sup>18</sup>	2011	Covariate	Harmful	Patients in the Euro Heart Registry who underwent percutaneous intervention for different indications	In-hospital mortality	Hemodynamic Instability, Indication for PCI, Age, Left Main Disease, Diabetes, BMI, Prior Stroke, Valve Disease, Bifurcation Lesion, Sex, Proximal Left Anterior Descending	23032	26%	339	64 (NR)	Hospitalization period

							Diseased, TIMI, Type-C Lesion, Smoking, Number of Diseased Vessels, Prior CABG					
21392604	de Boer <sup>22</sup>	2011	Covariate	Harmful	Patients who participated in 22 randomized clinical trials of PCI vs. fibrinolysis	All-cause mortality	Age, HR, SBP, Anterior Infarct Location, Prior MI, Time to Treatment, Diabetes, Sex	5421	24.4%	363	NR	30 days
22093204	Roe <sup>25</sup>	2011	Covariate	Protective	NSTEMI	Long-term mortality	Signs of Heart Failure on Presentation, Prior Heart Failure, Prior Stroke, Age, Diabetes, Race, Sex, Prior PAD, Smoking, Hematocrit, Creatinine, Prior MI, HR, SBP, Weight, EKG Changes, Prior CABG, Troponin Ratio, HTN, Prior PCI, Family History of CAD, Hyperlipidemia	33640	47.7%	NR	NR	median: 453 days
<b>CHF – MORTALITY</b>												
12748212	Bouvy <sup>1</sup>	2003	Covariate	Protective	Admission due to heart failure	Mortality	History of Renal Insufficiency, Beta Blockers, Ankle Edema, History of Diabetes, BP, Sex, Age, Weight	152	34.2%	51	69.7 (NR)	18 months
15769783	Adlam <sup>2</sup>	2005	Covariate	Protective	Primary care population prescribed loop diuretics for CHF	All-cause mortality	Age, BNP, CVA, Diabetes, EKG, Sex	528	59%	190	75 (10)	5 years (mean: 6.4 years)
15820281	Alehagen <sup>3</sup>	2005	Covariate	Protective	Patients who attended primary care because of symptoms and/or signs that might be attributed to heart failure	Cardiovascular mortality	NYHA Class, Age, Sex, Diabetes	510	47.8%	71	73 (6)	6 years
16219658	Pocock <sup>4</sup>	2006	Covariate	Protective	Patients with CHF	All-cause mortality	Diabetes, Age, NYHA Class, Prior Heart Failure Hospitalization, Pulmonary Edema, Smoking, CHF, Cardiomegaly, Bundle Branch Block, Previous MI, Dependent Edema, Pulmonary Crackles, Mitral Regurgitation, LVEF, Dyspnea, AF, DBP, HR, BMI, Candesartan vs. Placebo, Sex	7599	68.4%	1831	65.5 (10.9)	median: 3.17 years
16534009	Levy <sup>5</sup>	2006	Covariate	Protective	Patients with heart failure (ejection fraction < 30% and NYHA functional class IIIB to IV)	Mortality	Cholesterol, NYHA Class, Allopurinol Use, Ischemic Etiology, Hemoglobin, Diuretic Dose, Age, Sex, Uric	1125	24%	395	65 (11)	1 year

							Acid, Sodium, LVEF, Lymphocytes, SBP, Statin Use					
16928995	Rassi <sup>6</sup>	2006	Covariate	Protective	Patients with Chagas heart disease	Mortality	NYHA Class, Cardiomegaly, Wall Motion Abnormality, Nonsustained VT, Low QRS Voltage, Sex	424	41.7%	130	47 (11)	10 years (mean: 7.9 years)
18296827	Miura <sup>7</sup>	2008	Covariate	Protective	Patients with idiopathic dilated cardiomyopathy (IDC)	All-cause mortality	Sex, Left Ventricular Diameter Index, Age, NYHA Class, LVEF	1554	27.4%	420	NR	5 years
22114101	O'Connor <sup>8</sup>	2012	Covariate	Protective	Patients enrolled in the HF-ACTION study with a NYHF class II-IV, LVEF < 35%	All-cause mortality	Serum Urea Nitrogen, Exercise Duration on CPX Test, BMI, Sex	387	28%	387	59 (NR)	median: 2.5 years
<b>CAD – MORBIDITY</b>												
1739359	Hubbard <sup>36</sup>	1992	Covariate	Protective	Patients with symptomatic CAD who underwent exercise equilibrium radionuclide angiography and coronary angiography within 6 months	Three vessel or left main CAD	Previous MI, Sex, Diabetes, Angina, Age	680	27.6%	215	58.6 (NR)	NA (case control)
9217616	Morise <sup>37</sup>	1997	Covariate	Protective	Patients with suspected CAD and normal resting EKGs who presented for exercise testing	Severe CAD (presence of $\geq 2$ vessel with $\geq 70\%$ luminal diameter narrowing)	Family History, Hyperlipidemia, Obesity, HTN, Smoking, BMI, Diabetes, Symptoms, Estrogen Status, Age, Sex	915	48%	170	54 (1)	Diagnosis
9313022	Selker <sup>38</sup>	1997	Covariate	Protective	Patients who had acute MI and ST-segment elevation on EKG	Thrombolysis-Related Major Bleed Component	Sex, Age, History of HTN, HR with SBP, Thrombolytic Agents, SBP	2525	NR	NR	NR	30 days
10519495	Wong <sup>11</sup>	1999	Covariate	Harmful	Patients undergoing CABG surgery who received fast-track cardiac anesthesia (FTCA) technique, consisting of low-dose fentanyl (10-15 mg/kg) and propofol infusion	Delayed extubation (defined a priori as extubation time > 10 hours)	Postoperative Excess Bleeding, IABP, Age, Sex, Atrial Arrhythmia, Inotropes	885	24%	221	60.8 (11.3)	10 hours (median: 7 hours)
10519495	Wong <sup>11</sup>	1999	Covariate	Harmful	Patients undergoing CABG surgery who received fast-track cardiac anesthesia (FTCA) technique, consisting of low-dose fentanyl (10-15 mg/kg) and propofol infusion	Prolonged ICU length of stay (defined as stays > 48 hours)	IABP, Renal Insufficiency, Age, Atrial Arrhythmia, Inotropes, Sex, MI, Postoperative Excessive Bleeding	885	24%	221	60.8 (11.3)	48 hours (mean: 1 day)
12473877	West <sup>39</sup>	2002	Covariate	Protective	Patients in the LIPID study with previous ACS and baseline total cholesterol of 4-7 mmol/l	Non-hemorrhagic stroke	Age, AF, Stroke at Baseline, History of Diabetes, SBP, Smoking, BMI, Sex,	9014	16.8%	388	NR	mean: 6 years

					(randomly assigned to receive either 40 mg of pravastatin or placebo)		History of HTN, Creatinine Clearance, HDL Cholesterol, Triglycerides, Anterior MI, Total Cholesterol, UA, Pravastatin					
12902080	Charlesworth <sup>40</sup>	2003	Covariate	Harmful	Patients undergoing isolated CABG surgery with or without extracorporeal circulation	Perioperative stroke (a new focal neurologic deficit that appears and is still at least partially evident more than 24 hours after its onset, occurring during or after the CABG procedure and established before discharge)	Age, Emergency Priority, Vascular Disease, Renal Failure or Creatinine, Diabetes, LVEF < 40%, Urgent Priority, Sex	33062	28.3%	532	NR	Hospitalization period
15312862	Sadanandan <sup>41</sup>	2004	Covariate	Protective	Patients with UA and NSTEMI	In-hospital CABG	Troponin, Prior Angina, ST-Segment Deviation, History of PAD, Sex, History of CABG	2219	34.4%	362	61.3 (11.7)	Hospitalization period
16159846	Fowler <sup>42</sup>	2005	Covariate	Harmful	Patients in the Society of Thoracic Surgeons National Cardiac Database who had CABG (isolated or in combination)	Major infection (mediastinitis, thoracotomy or vein harvest site infection, or septicemia)	BMI, Dialysis, Shock, Age, Other Procedures, Immunosuppressive Therapy, Valve Surgery, Renal Failure, Diabetes, Chronic Lung Disease, CHF, Other Cardiac Surgery, PVD, LVEF, Previous CABG, NYHA Class, Previous MI, Anticoagulants, CVA, Smoking, Valvular Insufficiency, HTN, Cerebrovascular Disease, Race, Sex, Hypercholesterolemia, Cardiac Device	165715	NR	11633	NR	Hospitalization period
16210253	Clayton <sup>29</sup>	2005	Covariate	Protective	Patients with stable symptomatic angina requiring treatment, preserved left ventricular function, and either previous MI or proved angiographic CAD	MI	Diabetes, Previous Angiography, Previous Stroke, Age, Smoking, Sex, Angina, Angina Medication, Previous MI, LVEF, Lipid Lowering Therapy, SBP, QT Interval, Creatinine, Glucose, WBC	7311	20.6%	495	63.5 (9.2)	mean: 4.9 years
16210253	Clayton <sup>29</sup>	2005	Covariate	Protective	Patients with stable symptomatic angina requiring treatment, preserved left ventricular function, and either previous MI or proved	Stroke	Previous Stroke, Smoking, Diabetes, Age, SBP, QT Interval, LVEF, Angina Medication, Angina, Previous Angiography,	7311	20.6%	495	63.5 (9.2)	mean: 4.9 years



					angiographic CAD		Lipid Lowering Therapy, Glucose, Creatinine, Previous MI, WBC, Sex					
19243970	Antunes <sup>43</sup>	2009	Covariate	Harmful	Patients undergoing CABG	In-hospital CVA	Cerebrovascular Disease, PVD, Left Ventricular Dysfunction, Non- Elective Surgery, Sex, Age	4567	11.8%	114	60.7 (9.3)	Hospitalization period
19332461	Subherwal <sup>44</sup>	2009	Covariate	Harmful	High-risk patients with non-ST-elevation acute coronary syndromes	In-hospital major bleed	Hematocrit, Sex, SBP, CHF, PVD, Diabetes, Creatinine Clearance, HR	71277	39.8%	6700	NR	Hospitalization period
20031896	Mehta <sup>45</sup>	2009	Covariate	Protective	Patients undergoing CABG	Reoperation for bleeding in patients undergoing CABG	Dialysis, Urgency of Surgery, BSA, Sex, Glycoprotein, Age, Creatinine, PVD, Thienopyridines, Immunosuppressive Therapy, Previous Cardiovascular Interventions, IABP, Shock, PCI, Diabetes, Race, 3-Vessel CAD, Cerebrovascular Disease, Chronic Lung Disease, CHF	422949	27.1%	10122	NR	Hospitalization period
20513595	Mehran <sup>24</sup>	2010	Covariate	Harmful	Patients with ACS including NSTEMI, STEMI, and biomarker negative ACS	Non-coronary artery bypass graft surgery (CABG)-related major bleeding	Sex, Anemia, STEMI, NSTEMI-Raised Biomarkers, Age, WBC, Creatinine, Biomarker Negative ACS, UFH/Enox + GPI, Glycoprotein Inhibitor, Bivalirudin	17421	28.7%	744	62.1 (11.7)	30 days
21700252	Mehran <sup>46</sup>	2011	Covariate	Harmful	Patients undergoing PCI from 3 randomized trials of bivalirudin vs. heparin plus glycoprotein lib/IIIa inhibitors (REPLACE-2, ACUITY, HORIZONS-AMI)	30 day risk of TIMI Bleed	STEMI, Sex, Smoking, NSTEMI-Raised Biomarkers, Bivalirudin + Glycoprotein Inhibitor, Age, Creatinine, WBC, Normal Biomarkers, Heparin/Enoxaparin + Glycoprotein Inhibitor, Bivalirudin	17034	25.6%	267	62.2 (11.4)	30 days
22064650	Podolecki <sup>47</sup>	2012	Covariate	Harmful	Patients with acute MI who were screened with coronary angiography and underwent PCI	Stroke (ischemic or hemorrhagic)	Previous Stroke/TIA, Sex, GFR, Nephropathy, Prior Acute MI, Smoking	2520	30%	52	62 (NR)	median: 25.5 months
<b>CAD – MORBIDITY AND MORTALITY</b>												
8890808	Magovern <sup>48</sup>	1996	Covariate	Harmful	Patients who underwent CABG surgery	Postoperative morbidity and mortality	Shock, Urgency of Operation, Age, Catheterization-Induced Coronary	1567	30%	NR	65 (9)	Hospitalization period

												Closure, Cardiomegaly, LVEF, CHF, Renal Insufficiency, Diabetes, Renal Dysfunction, Anemia, BUN, PVD, Cerebrovascular Disease, Sex, BMI, Atrial Arrhythmia, COPD, Reoperation, Albumin
8890808	Boersma <sup>23</sup>	2000	Covariate	Harmful	Patients presenting within 24 hours of an episode of ischemic chest pain (>10 minutes) either transient ST-segment elevation (>0.5 mm), transient or persistent ST-segment depression (>0.5 mm), T-wave inversion (>1.0 mm), or elevation of creatinine kinase	Composite of death or myocardial (re)infarction	Age, Signs of Heart Failure (Rales), Worst CCS-Class, Sex, ST-Depression on EKG, HR, SBP	9461	35%	1417	64 (NR)	30 days
11728352	Fortescue <sup>49</sup>	2001	Covariate	Harmful	Patients who underwent coronary artery bypass grafting and not involving valvular or other surgeries	In-hospital major adverse event (death, renal failure, reinfarction, cardiac arrest, CVA, or coma)	Creatinine, Age, Shock, Emergent Operation, Prior CABG, LVEF, History of Liver Disease, History of Stroke or TIA, History of COPD, HTN, Urgent Operation	6237	28.1%	408	64.8 (10.6)	Hospitalization period
15135693	Elhendy <sup>27</sup>	2004	Covariate	Protective	Patients with known or suspected CAD	Cardiac events	Exercise Wall Motion Score Index, Chest Pain, Metabolic Equivalents, Sex	2840	42%	143	62 (12)	4 years (median: 3 years)
16159983	Grayson <sup>50</sup>	2006	Covariate	Harmful	Patients undergoing adult percutaneous coronary intervention (PCI)	In-hospital MACE (in-hospital deaths, Q wave MI, emergency CABG surgery, and CVA)	Shock, Left Main Stem Lesion Treated, Age, Emergent PCI, Graft Lesion Treated, Sex, Cerebrovascular Disease, Urgent PCI	9914	29.1%	129	NR	Hospitalization period
16210253	Clayton <sup>29</sup>	2005	Covariate	Protective	Patients with stable symptomatic angina requiring treatment, preserved left ventricular function, and either previous MI or proved angiographic CAD	Death, MI or stroke	Diabetes, Smoking, Age, Previous Stroke, Previous Angiography, QT Interval, Angina, Lipid Lowering Therapy, SBP, LVEF, Sex, Previous MI, Angina Medication, Creatinine, Glucose, WBC	7311	20.6%	1063	63.5 (9.2)	mean: 4.9 years
17443658	Zhang <sup>51</sup>	2007	Covariate	Protective	Patients with STEMI who did not have emergency intervention therapy within 24hours of onset	Cardiogenic shock within 72 hours after admission	MI, Family History of CAD, Multivessel Disease, BMI, Previous MI, Results of Thrombolytic Therapy, Sex, Age, Killip Class	2077	28.3%	184	76.0 (7.8)	72 hours

17550750	Singh <sup>33</sup>	2007	Covariate	Harmful	Patients undergoing percutaneous coronary intervention	MACE (in hospital death, Q wave MI, urgent or emergent CABG, cerebrovascular event)	LVEF, Creatinine, Age, Cardiogenic Shock, MI, PAD, CHF, Diabetes, Sex	7457	31%	291	66.9 (12.1)	Hospitalization period
19936453	Santos <sup>52</sup>	2009	Covariate	Protective	Patients admitted to the emergency department with non-ST-segment elevation acute coronary syndrome	Death or (re)infarction risk within 30 days	Biomarkers, Prior Stroke, Diabetes, Creatinine, Sex, Age, Previous Use of ACE Inhibitor	1027	42.6%	54	61.6 (0.35)	30 days
20630456	Chen <sup>53</sup>	2010	Covariate	Harmful	Patients with unprotected left main stenosis who underwent percutaneous coronary intervention (patients from DISTAL study)	MACE	Acute MI, Shock, Previous MI, IABP, Ultrasound Guidance, Stent, Biomarkers, Incomplete Revascularization, Diabetes, Cholesterol, PVD, LVEF, Age, HTN, Smoking, Creatinine, Culotte, Sex, Non-Left Main Total Occlusion Failure, Statin Use, Stroke, GI Bleeding	337	21.1%	NR	66.6 (10.5)	MACE
20923988	Singh <sup>35</sup>	2010	Covariate	Protective	Patients who underwent percutaneous coronary intervention	Mortality or MI	LVEF, Age, BMI, CAD Comorbidity Index, Shock, Troponin, Prior CABG, CHF, Ventricular Arrhythmia, Three-Vessel Disease, MI, Smoking Status, Stable Angina, Prior PCI, Sex, UA, Stent Use, Procedural Failure, Glycoprotein Use, Cholesterol, MEDS	9165	29%	1939	66.8 (12.2)	Hospitalization period
<b>POPULATION SAMPLE – MORBIDITY</b>												
1985385	Anderson <sup>54</sup>	1991	Covariate	NA <sup>a</sup>	Members of the Framingham Heart Study and Framingham Offspring Study cohorts, who ranged in age from 30 to 74 years who were free of cardiovascular disease and cancer	Myocardial infarction (including silent and unrecognized MI)	Age x Sex, Diabetes x Sex, Diabetes, LVH x Sex, Smoking Status, Cholesterol, SBP, Age	5573	NR	NR	NR	12 years
1985385	Anderson <sup>54</sup>	1991	Covariate	NA <sup>b</sup>	Members of the Framingham Heart Study and Framingham Offspring Study cohorts, who ranged in age from 30 to 74 years who were free of cardiovascular disease and cancer	Stroke, including transient ischemia	Sex, Cholesterol, LVH, Diabetes x Sex, Diabetes, Smoking, Age, SBP	5573	NR	NR	NR	12 years
1985385	Anderson <sup>54</sup>	1991	Covariate	NA <sup>c</sup>	Members of the	Cardiovascular	Age x Sex, Diabetes x	5573	NR	NR	NR	12 years

					Framingham Heart Study and Framingham Offspring Study cohorts, who ranged in age from 30 to 74 years who were free of cardiovascular disease and cancer	disease	Sex, Diabetes, LVH, Smoking, Cholesterol, Sex, SBP, Age					
2003301	Wolf <sup>55</sup>	1991	Stratified	NA	Subjects in the Framingham Cohort aged 55-84 years and free of stroke	Incident stroke	LVH, Age, AF, CVD, Smoking, Diabetes, SBP, Antihypertensive Therapy	5734	58.6%	M: 213 F: 259	65.8 (NR)	10 years
8266381	D'Agostino <sup>56</sup>	1994	Stratified	NA	Framingham cohort participants aged 55 to 84 years, free of stroke	Incident stroke	Antihypertensive Therapy, LVH, AF, CVD, Smoking, Age, Diabetes, SBP	5734	58.6%	NR	NR	10 years
9603539	Wilson <sup>57</sup>	1998	Stratified	NA	Population based cohort from the Framingham Heart Study who were free of CVD	Incident coronary heart disease (CHD)	Baseline Survival Function, Total Cholesterol, HTN, Smoking, Age, HDL Cholesterol, Diabetes	5345	53.4%	NR	M: 48.6 (11.7) F: 49.8 (12)	12 years
10371227	Kannel <sup>58</sup>	1999	Stratified	NA	Members of the Framingham Heart Study, a population-based cohort aged 45-95 years old, free of heart failure, but predisposed by CAD, hypertension, or valvular heart disease	Heart failure	Congenital Heart Disease, Valve Disease, LVH, Cardiomegaly, Age, Diabetes, HR, SBP, Vital Capacity	15267	58.4%	M: 252 F: 234	M: 62 (NR) F: 63.8 (NR)	4 years
11296578	Qureshi <sup>59</sup>	2001	Covariate	Protective	Unselected volunteers without previous stroke, TIA, or carotid artery surgery	Asymptomatic carotid artery stenosis > 60% by duplex ultrasound	Age, CAD, Smoking, Hypercholesterolemia, Sex	887	69%	170	66.4 (12.9)	Diagnosis
11809350	Lumley <sup>60</sup>	2002	Stratified	NA	Population-based cohort study of men and women age 65 and older at study entry	Stroke	M: Diabetes, LVH, History of CVD, AF, Age, Impaired Fasting Glucose, SBP, Creatinine, 15-ft Walk Time F: AF, Age, History of CVD	5888	57.6%	NR	73 (NR)	5 years (median: 6.3 years)
11822347	Russell <sup>61</sup>	2002	Stratified	NA	Medicare recipients hospitalized for deep venous thrombosis or pulmonary embolism	Diagnosis of venous thromboembolism (deep vein thrombosis or pulmonary embolism)	M: Secondary Neoplasm x Age, Renal Failure x Age, Age, Cellulitis, Varicose Veins, Major Orthopedic Procedure, Primary Neoplasm, Corticosteroids, Pneumonia, Arthritis, Warfarin, Brain and/or Spinal Cord Injury, Blood Disorder x Age, COPD x Age, Abnormal Respiration, Diabetes, Potassium	2823353	52.9%	M: 600 F: 730	NR	Diagnosis

													Chloride, Ischemic Heart Disease, Prior DVT and/or PE, Secondary Neoplasm x Age F: Prior DVT and/or PE, Renal Failure x Age, Secondary Neoplasm x Age, Other CVD x Age, Age, Cellulitis, Major Orthopedic Procedure, Corticosteroids, Anemia x Age, Primary Neoplasm, Diabetes, Varicose Veins, Abnormal Respiration, Brain and/or Spinal Cord Injury, Warfarin, Blood Disorder x Age, IBS, COPD x Age, Neoplasm, Arthritis, Potassium Chloride, Hormone Replacement Therapy, Pneumonia, Ischemic Heart Disease, Pregnancy
12086074	Ciampi <sup>62</sup>	2001	Stratified	NA	Population based cohort of 18 to 74 years of age surveyed with respect to their cardiovascular status	Presence of CHD as either a history of MI, aorto-coronary bypass , other forms of specifically defined ischemic cardiopathies or cardiac complications before or after surgery	Family History, Diabetes, Smoking Status, Cholesterol, Age	1147	100%	NR	39.4 (16.6)	Diagnosis	
17088464	Wu <sup>15</sup>	2006	Stratified	NA	Men and women aged 35 to 59 years in Beijing and Guangzhou (patients from the USA-PRC study cohort)	Ischemic stroke	Diabetes, BMI, Cholesterol, Smoking, SBP, Age	9903	50.6%	M: 158 F: 108	46 (6)	11 years (mean: 15.1 years)	
17586511	Jee <sup>63</sup>	2008	Stratified	NA	Koreans aged 30-84 years insured by the National Health Insurance Corporation (NHIC)	Stroke	Diabetes, Smoking, Cholesterol, Alcohol Intake, Age, Physical Activity, BMI, SBP	1223740	36.5%	M: 29216 F: 18017	M: 46.6 (11) F: 49.4 (12.1)	10 years (mean: 13 years)	
18036028	Assmann <sup>64</sup>	2007	Covariate	Protective	Adult employees in the PROCAM study, which excluded subjects with a history of angina pectoris, MI, or stroke	Cerebral ischemic events (ischemic stroke or TIA)	Smoking, Diabetes, Age, SBP, Sex	26975	27.4%	85	45.7 (6.8)	10 years (mean: 12 years)	
18430204	Yang <sup>65</sup>	2008	Stratified	NA	Patients with type 2 diabetes	Heart failure	Urinary Albumin to Creatinine Ratio, CHD, Glycated Hemoglobin, BMI, Age, Blood	3456	54.8%	131	NR	median: 5.52 years	

							Hemoglobin						
19249635	Schanbel <sup>66</sup>	2009	Covariate	Protective	Framingham cohort participants free of atrial fibrillation, between ages 45 and 95 years	Atrial fibrillation	Age, BMI, Age x Murmur, SBP, Murmur, Sex x Age, Prevalent Heart Failure, Sex, Antihypertensive Therapy, Age x Prevalent Heart Failure, Recalibration for Baseline Survival	4764	55%	457	60.9 (9.9)	10 years	
20001655	Kjeldsen <sup>67</sup>	2009	Covariate	Protective	Patients with HTN and electrocardiographic LVH randomized to losartan-based versus atenolol-based therapy in the Losartan Intervention For Endpoint reduction in hypertension (LIFE) study	Incident myocardial infarction	History of Angina, Sex, History of Diabetes, Urinary Albumin to Creatinine Ratio, Cholesterol, Pulse Pressure	8624	54%	331	66.9 (NR)	mean: 4.8 years	
20535515	Wu <sup>68</sup>	2011	Stratified	NA	Case-Control: Patients admitted for stroke at community hospitals in Chongqing China	Stroke	M: Age, HTN, CHD, Family History of Stroke, Hyperlipidemia, DBP, Family History of HTN, Education Level, Diabetes, High Salt Consumption, Physical Exercise F: Age, HTN, Family History of Stroke, Hyperlipidemia, Diabetes, DBP, Family History of HTN, Education Level, Alcohol Intake, High Salt Consumption, BMI	NR	NR	NR	NR	NA (Case-Control)	
20671251	Chien <sup>69</sup>	2010	Covariate	Protective	Stroke-free at baseline	Stroke	AF, Family History of Stroke, Diabetes, Age, DBP, SBP, Sex	3513	52.7%	240	54.6 (NR)	10 years (mean: 15.9 years)	
21167488	Ramos <sup>70</sup>	2011	Covariate	Harmful	Patients free of cardiovascular disease	Peripheral arterial disease (ABI <0.9)	Smoking, Diabetes, Sex, Age, Pulse Pressure	4046	52%	235	63.4 (8.6)	NA (Case-Control)	
21177699	McGorrian <sup>71</sup>	2011	Covariate	NA <sup>d</sup>	Case-Control: patients with incident acute MI admitted to coronary care or equivalent unit were matched to at least one sex- and age-matched (within 5 years) control, who had no history of heart disease	Incident MI	Lipoprotein, Diabetes, Smoking, HTN, Secondhand Smoke Exposure, Age x Sex	12772	24.7%	5349	57.46 (12.13)	NA (Case-Control)	
<b>POPULATION SAMPLE – MORBIDITY AND MORTALITY</b>													
1985385	Anderson <sup>54</sup>	1991	Covariate	NA <sup>e</sup>	Members of the Framingham Heart	Coronary heart disease (MI and	Sex, Age x Sex, Diabetes, Diabetes x	5573	NR	NR	NR	12 years	

					Study and Framingham Offspring Study cohorts, who were free of CVD and cancer	CHD death plus angina pectoris and coronary insufficiency)	Sex, Smoking, LVH, Cholesterol, SBP, Age					
11724655	Stevens <sup>72</sup>	2001	Covariate	Protective	Patients presenting with newly diagnosed diabetes, a fasting plasma glucose >6 mmol/L (108 mg/dL) on 2 further occasions, and no recent history of MI, angina or heart failure	CHD (occurrence of fatal or non-fatal MI or sudden death)	Cholesterol, Smoking, HbA1c, SBP, Duration of Diabetes, Age at Diagnosis of Diabetes, Sex, Ethnicity	4540	42.9%	NR	52 (8.8)	10 years (median: 10.7 years)
14514579	Folsom <sup>73</sup>	2003	Stratified	NA	Adults initially free of CHD in the Atherosclerosis Risk in Communities study	Incident coronary heart disease (CHD)	Waist-to-Hip Ratio, Intima Media Thickness, Failure Time Function Constant, Cholesterol, Race, LVH, Smoking, Antihypertensive Therapy, WBC, Creatinine, Age, Keys Score, Factor VIII, SBP, BMI, Sports Activity, Serum Albumin	1722	50%	M: 212 F: 95	NR	10 years (mean: 10.2 years)
15173150	Liu <sup>74</sup>	2004	Stratified	NA	Chinese Multi-provincial Cohort Study aged 35 to 64 years free of CVD	CHD (coronary death and myocardial infarction)	HDL Cholesterol, Total Cholesterol, HTN, Smoking, Diabetes, Age	5251	53.6%	M: 195 F: 78	49 (NR)	10 years
16732001	Donnan <sup>75</sup>	2006	Covariate	Protective	Subjects with a diagnosis of type 2 diabetes residing in Tayside, Scotland	Major CHD events defined as fatal or nonfatal AMI or CHD death	Smoking, Treated HTN, Height, SBP x Treated HTN, HbA1c, Sex, SBP, Age at Diagnosis, Total Cholesterol, Duration of Diabetes, HbA1c x Follow Up	4569	47.4%	243	59.5 (12.1)	5 years (median: 4.1 years)
17088464	Wu <sup>15</sup>	2006	Stratified	NA	Men and women in Beijing and Guangzhou (patients from the USA-PRC study cohort)	Ischemic cardiovascular disease (ICVD) events	SBP, Smoking, Total Cholesterol, BMI, Diabetes, Age	9903	50.6%	M: 224 F: 147	46 (6)	11 years (mean: 15.1 years)
17088464	Wu <sup>15</sup>	2006	Stratified	NA	Men and women in Beijing and Guangzhou (patients from the USA-PRC study cohort)	Incident CHD event (defined as MI, possible acute MI and coronary death, ischemic cardiac arrest with successful resuscitation, excluded silent MI)	BMI, Diabetes, Total Cholesterol, Smoking, Age, SBP	9903	50.6%	M: 66 F: 39	46 (6)	11 years (mean: 15.1 years)
17090561	Woodward <sup>76</sup>	2007	Stratified	NA	Patients without CHD and whose prior diagnosis was not a TIA	10-year cardiovascular risk (cardiovascular death, hospitalization for CHD,	Diabetes, Family History, Total Cholesterol, Scottish Index of Multiple Deprivation Score, Age, Smoking, SBP,	13297	50.8%	M: 743 F: 422	M: 48.9 (0.1) F: 48.8 (0.1)	10 years

						cerebrovascular disease, or any coronary artery intervention)	HDL Cholesterol					
17478150	Mainous <sup>77</sup>	2007	Stratified	NA	Population-based cohort of adults in Atherosclerosis Risk in Communities Study (ARIC) who were free of cardiovascular disease at baseline	Incident CHD	M: 10-year CHD Risk, Age, History of Diabetes, Smoking, Hypercholesterolemia, Family History of CAD, HTN, Physical Activity F: 10-year CHD Risk, Age, History of Diabetes, Smoking, HTN, BMI, Hypercholesterolemia	14343	56.5%	NR	54.4 (5.7)	10 years
17615182	Hippisley-Cox <sup>78</sup>	2007	Stratified	NA	Population based cohort ages 35-74 free of cardiovascular disease	Cardiovascular disease (MI, CHD, stroke, TIA)	Age, Antihypertensive Therapy, Smoking, Family History of CVD, Townsend Score of Output Area, BMI, SBP, Cholesterol	1283174	49.6%	M: 37843 F: 27828	NR	10 years
18036028	Assmann <sup>64</sup>	2007	Covariate	Protective	Adult employees in the PROCAM study which excluded subjects with a history of angina pectoris, myocardial infarction, or stroke	Major coronary events	Fasting Blood Glucose or Diabetes, HDL Cholesterol, LDL Cholesterol, SBP, Family History, Triglycerides, Smoking	26975	31.6%	511	45.7 (6.8)	10 years (mean: 12 years)
18212285	D'Agostino <sup>79</sup>	2008	Stratified	NA	Healthy participants free of CVD in the original Framingham Heart Study and the Framingham Offspring Study	CVD events and death	Age, SBP x Treatment, Total Cholesterol, Smoking, Diabetes, HDL Cholesterol	8491	53.3%	M: 718 F: 456	M: 48.5 (10.8) F: 49.1 (11.1)	10 years
18591403	Cederholm <sup>80</sup>	2008	Covariate	Protective	Type 2 diabetics, defined treatment with 1) diet only, 2) oral hypoglycemic agents only, or 3) insulin only or combined with oral agents, and onset age of diabetes 40 years	First incident fatal or nonfatal CVD	Sex, Smoking, Lipid Lowering Therapy, Antihypertensive Therapy, HbA1c, Diabetes Duration, Onset Age, BMI, SBP	11646	43.1%	990	50.7 (9.8)	5 years
18591432	Wilson <sup>81</sup>	2008	Covariate	Protective	Men and women who were the offspring and spouses of the participants in the original Framingham Heart Study, who were free from CHD and CVD	First CHD Event (MI, angina pectoris, coronary insufficiency, and CHD-related death)	Smoking, Diabetes, Age, Cholesterol, BMI, SBP, Sex	4780	51.6%	492	36.7 (9.75)	24 years
18762707	Teramoto <sup>82</sup>	2008	Covariate	Protective	Patients with hypercholesterolemia (total cholesterol 220-270 mg/dl) and no history of ischemic heart disease or stroke, assigned either to diet alone or diet in	CHD	Glucose x HTN, HTN, Age, Sex, HDL Cholesterol, Smoking, LDL Cholesterol, Diet, Diet + Pravastatin	7832	68.5%	138	58.3 (7.2)	5 years (mean: 5.3 years)



					combination with pravastatin (10-20 mg/day)							
19157168	Maracy <sup>83</sup>	2008	Covariate	Protective	Type 2 diabetics seen at a metabolism clinic	Ischemic heart disease	Age, Sex, BMI, Duration, HTN, BMI	2101	56%	773	NR	NA (cross-sectional)
19763133	Arima <sup>84</sup>	2009	Covariate	Protective	Healthy individuals living in the town of Hisayama, a suburb of Fukuoka City in Southern Japan	Cardiovascular disease	Sex, Diabetes, Smoking, Age, SBP, LDL Cholesterol, HDL Cholesterol	1756	57%	216	59 (12)	14 years
19958966	Sacco <sup>85</sup>	2009	Covariate	Protective	Community participants free of stroke and coronary artery disease	Combined cardiovascular outcomes (stroke, myocardial infarction, or vascular death)	Race, SBP, Physical Activity, Smoking, Alcohol Intake, Total Cholesterol, PVD, DBP x Antihypertensive Therapy, Waist, DBP, Age, FBS	2737	63.2%	NR	68.8 (10.4)	median: 9 years
20001655	Kjeldsen <sup>67</sup>	2009	Covariate	Protective	Patients with HTN and electrocardiographic LVH randomized to losartan-based vs. atenolol-based therapy in the Losartan Intervention For Endpoint reduction in hypertension (LIFE) study	Cardiovascular death, stroke, or MI	Sex, History of AF, History of Stroke or TIA, Smoking, History of Diabetes, Age, History of Ischemic Heart Disease, Urinary Albumin to Creatinine Ratio, LVH, Total Cholesterol, Physical Activity	9193	54%	1096	66.9 (NR)	mean: 4.8 years
20236721	Zgibor <sup>86</sup>	2010	Stratified	NA	Participants in the Pittsburgh Epidemiology of Diabetes Complications Study (EDC) cohort who had childhood type 1 diabetes diagnosis at least 6 years prior to enrollment	Hard CHD event (first nonfatal MI that was confirmed by medical records or Q-waves on ECG's according to Minnesota codes 1.1 or 1.2) and fatal CHD	M: HDL Cholesterol, Diabetes Duration, WBC, Microalbuminuria F: SBP, Antihypertensive Therapy, Diabetes Duration, Non-HDL Cholesterol, Waist-to-Hip Ratio	603	49.6%	NR	M: 27.1 (7.6) F: 27.3 (7.9)	15 years
21719139	Zethelius <sup>87</sup>	2011	Covariate	Protective	Patients in a population based registry for type II diabetes	Fatal/nonfatal CVD	Previous CVD, Onset Age, Diabetes Duration, SBP, BMI, Macroalbuminuria, AF, Sex, Smoking, Microalbuminuria, Cholesterol, HbA1c	24288	40%	2488	62 (9)	5 years (mean: 4.8 years)
22374565	Kang <sup>88</sup>	2012	Covariate	NA <sup>f</sup>	Population-based cohort with an equal portion of rural and urban participants from Jiangsu province of China, who were followed for incident diabetes and CVD	Incident coronary heart disease (CHD) and stroke	Age, BP, Triglycerides, Sex x Waist Circumference, HDL Cholesterol, Fasting Plasma Glucose	3598	59.7%	87	50.2 (NR)	median: 6.3 years
<b>CARDIAC SURGERY – MORTALITY</b>												
7828293	Tu <sup>89</sup>	1995	Covariate	Harmful	Patients undergoing cardiac surgery	In-hospital mortality	Urgency of Surgery, Age, Type of Surgery, Left Ventricular	6213	NR	229	NR	Hospitalization

							Function, Repeat Operation, Sex, Elective Surgery, CABG					period
10456395	Nashef <sup>90</sup>	1999	Covariate	Harmful	Patients undergoing cardiac surgery	Early mortality	Postinfarct Septal Rupture, Endocarditis, Critical Preoperative State, Left Ventricular Dysfunction, Previous Cardiac Surgery, Surgery on Thoracic Aorta, Emergency Operation, Neurological Dysfunction, Operation Other Than CABG, Pulmonary HTN, Recent MI, Creatinine, UA, Arteriopathy, Chronic Pulmonary Disease, Sex, Age	13302	NR	NR	NR	NR
10750767	Bernstein <sup>91</sup>	2000	Covariate	Harmful	Patients who had open-heart surgery evaluated with planar thallium myocardial perfusion imaging. Surgical procedures included aortocoronary bypass (ACB), valve replacement or repair (VLV), and combinations of the two (COM).	In-hospital mortality	Reoperation, Dialysis, Cirrhosis, Cardiogenic Shock, Ideopathic Thrombopenic Purpura, Age, Sex, Pulmonary HTN, LVEF, Endocarditis, Bypass Surgery, COPD, Failed Angioplasty or Catheterization, Neurologic Disorder, Tricuspid Valve Replacement, Substance Abuse, Mitral Valve Replacement, Intubated, Acute MI, Preoperative IABP, PVD, Renal Failure, Diabetes, HTN, CHF, Cardiomegaly, Left Main Disease, Carotid Disease, Aneurysm, Asthma, VT or VF, BMI, Pacemaker Dependency, Neoplasm, Cold Agglutinins, Blood Products Refused, Immunodeficiency	8593	NR	455	NR	Hospitalization period
14583307	Huijskes <sup>92</sup>	2003	Covariate	Harmful	Patient who underwent a CABG and/or heart valve operation	In-hospital mortality	MI, Prior Cardiac Surgery, Creatinine, Emergency Procedure, LVEF, CABG, Sex, Age	4855	27%	171	NR	Hospitalization period
15172248	Nowicki <sup>93</sup>	2004	Covariate	Harmful	Patients undergoing	In-hospital mortality	Emergent Priority, Age,	3150	48.4%	296	66.7	Hospitaliz

					mitral valve surgery		Prior Stroke, NYHA Class, CAD, Urgent Priority, Valve Replacement, Creatinine, CHF, Diabetes, Sex, Year			(NR)		ation period
15998680	Ambler <sup>94</sup>	2005	Covariate	Harmful	Patients who underwent aortic and/or mitral heart valve surgery (included both repair and replacement surgeries)	In-hospital mortality	Priority of Surgery, Renal Failure, Age, Operation Sequence, LVEF, Concomitant Tricuspid Surgery, Valve Operation, Preoperative Arrhythmia, CABG, BMI, HTN, Diabetes, Sex, Year	16679	41.6%	1067	64.8 (12.3)	Hospitalization period
16039188	Jin <sup>95</sup>	2005	Covariate	Harmful	Patients aged 30 to 95 years who underwent aortic valve replacement (AVR) or mitral valve replacement or repair (MVRR)	Operative Mortality	Priority of Operation, Creatinine, History of Cardiac Surgery, BMI, Pulmonary HTN, COPD, PVD, AF, CABG, NYHA Class, Sex, Age	1596	NR	96	NR	Hospitalization period
17307434	Hannan <sup>96</sup>	2007	Covariate	Harmful	Patients who underwent isolated cardiac valve surgery (valve surgery without CABG) or cardiac valve surgery along with concomitant CABG	In-hospital mortality	Shock, Hepatic Failure, Dialysis, MI, Multiple Valve Repair or Replacement, Endocarditis, Creatinine, Previous Open Heart Operation, PAD, COPD, Mitral Valve Replacement or Repair, Sex, Age	10702	39%	472	NR	Hospitalization period
17307434	Hannan <sup>96</sup>	2007	Covariate	Harmful	Patients who underwent isolated cardiac valve surgery (valve surgery without CABG) or cardiac valve surgery along with concomitant CABG	In-hospital mortality	Shock, Dialysis, Mitral Valve Repair or Replacement or CABG, MI, Creatinine, Endocarditis, LVEF, Previous Open Heart Operation, Sex, Extensively Calcified Ascending Aorta, Cerebrovascular Disease, Diabetes, Age	8823	39%	784	NR	Hospitalization period
19523840	Nissinen <sup>97</sup>	2009	Covariate	Harmful	Patients who underwent cardiac surgery	Postoperative mortality	Age, LVEF, CKD, Sex, Number of Procedures, Ventricular Septal Defect, Thoracic Aorta Surgery, Critical Preoperative Status, Neurological Dysfunction, Pulmonary Disease, Reoperation, Extracardiac Arteriopathy	3613	22.5%	90	67.7 (10.3)	30 days

19559823	O'Brien <sup>98</sup>	2009	Covariate	NA <sup>g</sup>	Patients who underwent isolated single aortic or mitral valve surgery	Mortality	BSA, Level of Urgency, Dialysis, Sex x BSA, Reoperation, Mitral Valve Repair, Endocarditis, CHF x NYHA Class, Diabetes, Shock, Creatinine, IABP or Inotropes, Immunosuppressive Treatment, Chronic Lung Disease, PVD, Stenosis, UA, AF, Left Main Disease, MI, HTN, Mitral Valve Replacement, Age, LVEF, CVD, Prior CVA, Mitral Insufficiency, Tricuspid Insufficiency, Number of Diseased Vessels, Race	109759	44.6%	3706	NR	30 days
<b>STROKE – MORBIDITY AND MORTALITY</b>												
1527533	Hankey <sup>99</sup>	1992	Covariate	Protective	Transient ischemic attack patients referred to a university hospital without prior stroke	Stroke, myocardial infarction, or vascular death	PVD, Carotid and Vertebral-Basilar TIAs, Brain TIA, Sex, Neurological Signs, LVH, Age, Number of TIAs	469	32.4%	118	62.1 (12)	mean: 4.1 years
1527533	Hankey <sup>99</sup>	1992	Covariate	Protective	Transient ischemic attack patients referred to a university hospital without prior stroke	Coronary event	Carotid and Vertebral-Basilar TIAs, Sex, Ischemic Heart Disease, PVD, Neurological Signs, LVH, Age, Number of TIAs	469	32.40%	58	62.1 (12)	mean: 4.1 years
8290048	Lisk <sup>100</sup>	1994	Covariate	Harmful	Patients with intracerebral hemorrhage	Poor outcome (Rankin scale 5 and death)	Admission Interval, Age, DBP, Hemorrhage Location, Surgery, SBP, Sex, Pupil Abnormality, Mental Status, Mass Effect, Hemorrhage Size, Hyperventilation, Subarachnoid Blood, Ventricular Extension, GCS	75	59%	35	58.6 (16.4)	mean: 18 days
10382694	Rothwell <sup>101</sup>	1999	Covariate	Harmful	Patients were recruited to ECST if they had had a carotid distribution TIA, minor ischemic stroke, non-disabling major ischemic stroke, or a retinal infarction in the previous 6 months, and had evidence of ipsilateral carotid stenosis	Any major stroke (fatal or lasting longer than 7 days) or death from any other cause within 30 days of surgery	PVD, SBP, Sex	1208	NR	117	NR	30 days

17068305	Kent <sup>102</sup>	2006	Covariate	Protective	Patients with acute stroke being evaluated for thrombolysis	Normal/Near-Normal outcome (mRS $\leq$ 1)	rt-PA Treatment, Sex, Prior Stroke, Age, Symptom Onset to Treatment Time, Age x NIHSS, Treatment x Time, SBP, NIHSS, Treatment x SBP, Treatment x Sex, Diabetes, Treatment x Prior Stroke	2131	45.3%	773	65.9 (11.4)	Hospitalization period
18823637	Koopman <sup>103</sup>	2009	Covariate	Protective	Cerebral venous thrombosis (CVT) patients aged older than 15 years who were evaluated in the hospital	Poor outcome (mRS > 2)	CNS Infection, CVT, Malignancy, GCS, Age, Mental Status Disorder, Intracranial Hemorrhage, Sex	90	78%	16	36.2 (NR)	mean: 1.58 years
19420921	Ferro <sup>104</sup>	2009	Covariate	Protective	Patient who had cerebral vein and dural sinus thrombosis	mRS > 2	Malignancy, Coma, DVT, Mental Status Disturbance, Sex, Intracranial hemorrhage	624	74%	19	NR	median: 1.3 years
<b>STROKE – MORTALITY</b>												
9645975	Arboix <sup>105</sup>	1998	Covariate	Protective	Patients with cardioembolic cerebral infarction	In-hospital mortality	Altered Consciousness, Limb Weakness, CHF, Sex, Age	231	62.8%	63	NR	Hospitalization period
18004645	Roquer <sup>106</sup>	2007	Covariate	Protective	Patients with acute ischemic stroke	In-hospital mortality	Age, Sex, NIHSS, Glycemia	1527	49.50%	197	73 (12)	Hospitalization period
21300951	Saposnik <sup>107</sup>	2011	Covariate	Protective	Patients presenting with an acute ischemic stroke	Mortality	Stroke Severity, Stroke Subtype, Renal Dialysis, Glucose, Preadmission Disability, AF, Cancer, CHF, Sex, Age	8223	47.4%	1004	72.04 (13.86)	30 days
21300951	Saposnik <sup>107</sup>	2011	Covariate	Protective	Patients presenting with an acute ischemic stroke	Mortality	Stroke Severity, Renal Dialysis, Stroke Subtype, Preadmission Disability, Cancer, CHF, Glucose, Smoking, AF, Previous MI, Sex, Age	8223	47.4%	1853	72.04 (13.86)	1 year
<b>VTE – MORBIDITY</b>												
10074953	Kuijjer <sup>108</sup>	1999	Covariate	Harmful	Patients with confirmed venous thromboembolism	All bleeding complications	Malignancy, Age, Sex	241	45.6%	63	60.7 (17)	3 months
10102459	Kahn <sup>109</sup>	1999	Covariate	Protective	Patients suspected of deep vein thrombosis	Deep vein thrombosis	Orthopedic Surgery, Superficial Venous Dilation, Sex, Temperature	271	50.60%	73	57.1 (16.9)	6 months
12637130	Miniati <sup>110</sup>	2003	Covariate	Protective	Patients who were referred for suspected pulmonary embolism	Pulmonary embolism (diagnosis was based on angiographic or autopsy)	Amputation of Hilar Artery, Oligemia, Consolidation, EKG Signs of Acute Right Ventricular Overload,	1100	55%	440	NR	Diagnosis

						documentation)	Dyspnea, Age, Hemoptysis, Sex, Thrombophlebitis, Chest Pain, Pre-Existing Disease, Fever, Pulmonary Edema					
14563499	Constans <sup>111</sup>	2003	Covariate	Protective	Outpatients with suspected deep venous thrombosis	In-hospital deep venous thrombosis	Unilateral Pain, Lower Limb Enlargement, Lower Limb Paralysis, Confinement to Bed, Sex, Other Diagnosis	444	66.2%	126	61 (18)	Hospitalization period
16607053	Chen <sup>112</sup>	2006	Covariate	Harmful	Patients with acute dyspnea presenting in the emergency room	Pulmonary embolism diagnosis at admission	Leg Edema, EKG RV Strain Signs, High Alveolar-Arterial Oxygen Gradient, Clear Chest Roentgenogram, Cough, Sex, Unclear Breath Sounds, Chest Tightness	148	45.3%	56	66.3 (17)	NA (case control)
18436792	Miniati <sup>113</sup>	2008	Covariate	Protective	Patients referred from a department or another hospital for suspected pulmonary embolism	Pulmonary embolism confirmed by angiography or autopsy findings	Dyspnea, Acute Cor Pulmonale, Age, Chest Pain, Hemoptysis, Leg Swelling, Syncope, DVT, Sex, Orthopnea, Immobilization, CVD, Crackles, Pulmonary Disease, Wheezes, Fever	1100	55%	440	NR	Diagnosis
<b>AORTIC DISEASES – MORTALITY</b>												
16146566	Urbonavicius <sup>114</sup>	2005	Covariate	Harmful	Patients with a diagnosis of AAA who were treated surgically	Post-operative mortality	Previous MI, Sex, Clinical Course of Abdominal Aortic Aneurysm	69	24.6%	8	NR	30 days
19249184	Giles <sup>115</sup>	2009	Covariate	Harmful	Propensity score matched Medicare beneficiaries undergoing elective open or endovascular abdominal aortic aneurysm repair	Perioperative mortality	Open Repair, Age, Renal Disease, Chronic Renal Insufficiency, CHF, Vascular Disease, Sex	22830	19.7%	1621	NR	30 days
19782526	Egorova <sup>116</sup>	2009	Covariate	Harmful	Medicare Patients > 65 years old hospitalized after undergoing Endovascular aneurysm repair (EVAR)	Mortality	Renal Failure, Lower Extremity Ischemia, Age, CHF, Liver Disease, Sex, Neurological Risk Factor, COPD, Hospital Annual Volume, Surgeon Endovascular Aneurysm Repair Volume	44630	17.1%	NR	NR	30 days
21412997	Grant <sup>117</sup>	2011	Covariate	Harmful	Patients undergoing elective abdominal aortic aneurysm repair surgery	30-day all-cause mortality	Open Surgery, Antiplatelet Therapy, Sex, Diabetes, Respiratory Disease,	1936	17.7%	31	NR	30 days

Age, Creatinine												
POPULATION SAMPLE – MORTALITY												
1985385	Anderson <sup>54</sup>	1991	Covariate	NA <sup>b</sup>	Members of the Framingham Heart Study and Framingham Offspring Study cohorts, who ranged in age from 30 to 74 years who were free of cardiovascular disease and cancer	Death from CHD (sudden or non-sudden)	Sex, Diabetes, LVH, Smoking, Diabetes x Sex, Cholesterol, SBP, Age	5573	NR	NR	NR	12 years
1985385	Anderson <sup>54</sup>	1991	Covariate	NA <sup>b</sup>	Members of the Framingham Heart Study and Framingham Offspring Study cohorts, who ranged in age from 30 to 74 years who were free of cardiovascular disease and cancer	Cardiovascular death	Sex, Diabetes, Smoking, Diabetes x Sex, LVH, Cholesterol, SBP, Age	5573	NR	NR	NR	12 years
9425461	Knuiman <sup>118</sup>	1997	Stratified	NA	General population health survey of people living in Western Australia	Mortality	Smoking, Cholesterol, BMI, Age, SBP	3891	50.6%	M: 116 F: 71	M: 56.4 (9.7) F: 55.8 (9.3)	10 years
11006890	Kornitzer <sup>119</sup>	2000	Stratified	NA	Free of coronary heart disease (CHD) at baseline	CHD mortality	HDL Cholesterol, Total Cholesterol, Age, Smoking, SBP	6066	29.0%	M: 84 F: 30	NR	10 years
11451781	Pocock <sup>120</sup>	2001	Stratified	NA	Patients included in anti-hypertensive drug trials	Cardiovascular death	Age, Smoking, Creatinine, Diabetes, Height, History of MI, History of Stroke, LVH, SBP, Total Cholesterol	NR	NR	M: 27987 F: 20802	NR	mean: 5.2 years
12788299	Conroy <sup>121</sup>	2003	Stratified	NA	General population	Fatal cardiovascular disease	Cholesterol, Sex, Smoking, SBP	NR	0%	NR	NR	10 years
15662552	Balkau <sup>122</sup>	2004	Stratified	NA	Population-based studies or large studies in occupational groups in Europe	Mortality	Cholesterol, Smoking, SBP, Diabetes, Fasting Plasma Glucose, Impaired Glucose Regulation, Age	25413	35.0%	M: 301 F: 56	NR	5 years
15662552	Balkau <sup>122</sup>	2004	Stratified	NA	Population-based studies or large studies in occupational groups in Europe	Mortality	M: Fasting Plasma Glucose, Diabetes, Age, Impaired Glucose Regulation F: Age, Fasting Plasma Glucose, Diabetes, Diabetes	25413	35.0%	M: 632 F: 159	NR	10 years
15933390	Porock <sup>123</sup>	2005	Covariate	Protective	Long-term-care residents in nursing home	Mortality	Cancer, Admission, Deteriorated, Age, Cancer x Age, Admission x Deteriorated, Shortness of Breath, Renal Disease, Sex, Loss of Appetite, Dehydrated, Weight Loss, CHF, ADLs,	32599	73.7%	7473	NR	6 months

													Cognitive Performance Scale, Dementia
20001655	Kjeldsen <sup>67</sup>	2009	Covariate	Protective	Patients with HTN and electrocardiographic LVH randomized to losartan-based versus atenolol-based therapy in the Losartan Intervention For Endpoint reduction in hypertension (LIFE) study	Cardiovascular death	Age x Smoking, Smoking, History of AF, History of Stroke or TIA, Sex, Urinary Albumin to Creatinine Ratio, Prior Ischemic Heart Disease, LVH, Creatinine, Glucose, Exercise	9193	54%	438	66.9 (NR)	mean: 4.8 years	
<b>ARRHYTHMIA – MORBIDITY</b>													
2330896	Cabin <sup>124</sup>	1990	Covariate	Harmful	Patients without mitral stenosis or prosthetic valves who were referred to the echocardiography laboratory with atrial fibrillation	ST segment elevation MI c embolization	Structural Heart Disease, Sex, Left Atrial Size	272	9.9%	27	68 (NR)	12 months (mean: 2.75 years)	
10356104	Hart <sup>125</sup>	1999	Covariate	Harmful	Adults with documented sustained or recurrent AF without mitral stenosis or prosthetic cardiac valves who were recruited from inpatient and outpatient facilities, assigned to aspirin 325 mg/d or aspirin plus fixed, low-dose warfarin	Incident ischemic stroke	Sex, Prior Stroke or TIA, SBP, History of HTN, Age, Alcohol Intake	1853	30.40%	101	69 (10)	mean: 2.0 years	
12941677	Wang <sup>126</sup>	2003	Covariate	Harmful	Community-based cohort (FRAMINGHAM) with new-onset AF	Stroke	Diabetes, Sex, Prior Stroke or TIA, Age, SBP	868	47%	111	75 (9)	5 years (mean: 4.3 years)	
17099015	Shireman <sup>127</sup>	2006	Covariate	Harmful	Elderly patient at least 65 year old with atrial fibrillation who were taking warfarin	Major acute bleeding event within 90 days of the hospital discharge	Anemia, Substance Use, Recent Bleeding Event, Age, Antiplatelet Therapy, Sex, Diabetes	19875	52.50%	318	NR	90 days	
19762550	Lip <sup>128</sup>	2010	Covariate	Harmful	Ambulant and hospitalized patients with AF without mitral stenosis or previous heart valve surgery and who did not use either VKA or heparin at discharge, ≥ 18 years old with an ECG or Holter recording showing atrial fibrillation during the qualifying	Stroke and thromboembolism	Age, Stroke/TIA, CHF/LVD, Diabetes, Sex, HTN, Vascular Disease	5333	40.80%	25	66 (14)	1 year	
21324428	Mathews <sup>129</sup>	2011	Covariate	Harmful	Patients with atrial fibrillation who underwent standard two-dimensional, Doppler, and speckle-tracking echocardiography	In-hospital major bleeding associated with acute myocardial infarction	Signs of Heart Failure, Hemoglobin, ST-Segment Changes, Sex, Previous PAD, Diabetes, Warfarin Use, Creatinine, SBP,	72313	35.3%	7810	64 (NR)	Hospitalization period	



HR, Age, Weight												
CARDIAC SURGERY – MORBIDITY												
1623792	Tuman <sup>130</sup>	1992	Covariate	Harmful	Adult patients undergoing cardiac surgery (cardiac surgery combined with ventricular aneurysmectomy, endocardial mapping, carotid endarterectomy, or intracardiac procedures other than valvular surgery were excluded)	Post-operative morbidity (defined as MI, extended pulmonary assistance, renal insufficiency, serious infection, neurologic abnormalities including coma)	Reoperation, Surgical Procedure, Age, Age of Previous MI, Emergency Surgery, CABG + Valve, Multivalve Procedure, Cerebrovascular Disease, Pulmonary HTN, CHF, LVD, Renal Dysfunction, Sex, Mitral Valve Replacement, Aortic Valve Replacement	3156	34.3%	701	NR	NR
12822622	Lim <sup>131</sup>	2003	Covariate	Protective	Patients undergoing mitral valve repair	Coronary Disease (luminal narrowing of 50%)	EKG Evidence of Ischemia, Angina, Sex, Hypercholesterolemia, Age	359	35%	NR	65 (10)	Diagnosis
14583307	Huijskes <sup>92</sup>	2003	Covariate	Harmful	Patient who underwent a CABG and/or heart valve operation	Extended length of stay (ELOS) (care length of stay of at least 3 days or in-hospital death)	Combined CABG/Mitral Valve, Creatinine, LVEF, Recent MI, Prior Cardiac Surgery, Emergency Procedure, Critical Preoperative State, Failed PCI, Pulmonary HTN, Combined CABG/Aortic Valve, Neurological Dysfunction, Sex, Age, Hemoglobin	4855	27%	1001	NR	Hospitalization period
STROKE – MORBIDITY												
1527533	Hankey <sup>99</sup>	1992	Covariate	Protective	Transient ischemic attack patients referred to a university hospital without prior stroke	Stroke	Ischemic Heart Disease, PVD, Residual Neurological Signs, TIA, LVH, Carotid and Vertebral-Basilar TIAs, Sex, Age	469	32.4%	63	62.1 (12)	mean: 4.1 years
OTHER												
2072767	Selker <sup>132</sup>	1991	Covariate	NA <sup>d</sup>	Critical care unit patients whose chief complaint was chest pain or symptoms consistent with possible acute cardiac ischemia who sought medical care	Acute ischemia diagnosis (including acute myocardial infarction or unstable angina)	EKG T-waves Inverted, EKG ST-segment Changes, Chest or Left Arm Pressure/Pain, Sex, Age, EKG Q waves Present	3453	43%	1251	62 (NR)	NR
19751624	Ngako <sup>133</sup>	2009	Covariate	Protective	Patients presenting to the emergency room with suspicion of acute coronary syndrome	Acute coronary syndrome	Typical Left Arm/Jaw/Neck Pain, Typical Chest Pain, Epigastric Pain, History of CAD, Dyspnea, Sex	399	49.1%	124	76.9 (7.4)	1 month
20092615	Gencer <sup>134</sup>	2010	Covariate	NA <sup>d</sup>	Patients who reported any type of chest pain	Diagnosis of CHD (angina pectoris,	Sex x Age, Cardiovascular Risk	661	52.50%	85	55.4	1 year

					during visit. The presence of chest pain was ascertained according to the usual practice of each family practitioner.	unstable angina, myocardial infarction)	Factors, History of CVD, Area of Pain, Absence of Tenderness, Duration of Chest Pain, Precipitating with Exertion				(19)	
20603345	Bosner <sup>135</sup>	2010	Covariate	NA <sup>d</sup>	Patients who presented with chest pain localized on the anterior chest wall in the area between the clavicles who was seen by a primary care physician and subsequently tested for coronary artery disease	Presence of coronary artery disease	Vascular Disease, Pain Worse During Exercise, Pain Not Reproducible By Palpation, Patient Assumes Pain is Cardiac, Age x Sex, CHF, Diabetes, Stinging Pain, Localized Muscle Tension, Cough	1249	43.9%	180	59 (13.9)	NA (case control)
10571089	Anyanwu <sup>136</sup>	1999	Covariate	Harmful	All patients registered on the national waiting list for heart transplantation in the UK	Thirty-day graft survival	Previous Sternotomy, Donor Age, Donor Inotropic Support, Donor with Diabetes, Sex of Donor, History of Donor Drug Abuse, Ischemic time, Sex x BSA, Pre-op Circulatory Support, Pre-op Ventilatory Support, Pulmonary Vascular Resistance, Donor to Recipient BSA Ratio, Recipient Age, Retransplant	377	17%	332	49 (10)	30 days
16219658	Pocock <sup>4</sup>	2006	Covariate	Protective	Patients with chronic heart failure (CHF)	Composite of cardiovascular (CV) death and heart failure (HF) hospitalization	Diabetes, Prior Heart Failure Hospitalization, NYHA Class, Age, Cardiomegaly, CHF, Pulmonary Edema, Bundle Branch Block, Pulmonary Crackles, Dependent Edema, Dyspnea, Mitral Regurgitation, AF, LVEF, Previous MI, DBP, Smoking, HR, BMI, Sex, Candesartan vs Placebo	7599	68.4%	2460	65.5 (10.9)	median: 3.17 years
21537001	Haines <sup>137</sup>	2011	Covariate	Harmful	Patients undergoing implantable cardioverter-defibrillator implantation	In-hospital complication (mortality or any complications)	Previous Implantable Cardioverter-Defibrillator, Defibrillator Type, NYHA Class, Other Admission, Previous Valvular Surgery, Sex, BUN, Chronic Lung Disease, AF or Flutter, Age	134069	25.9%	8589	NR	Hospitalization period

22114101	O'Connor <sup>8</sup>	2012	Covariate	Protective	Patients enrolled in the HF-ACTION study with a NYHF class II-IV, LVEF ≤ 35%	All-cause death or hospitalization	KCCQ Symptom Stability, Serum Urea Nitrogen, Exercise Duration on CPX Test, Sex	2331	28%	1555	NR	median: 2.5 years
22456478	Bouleti <sup>138</sup>	2012	Covariate	NA <sup>h</sup>	Patients who underwent percutaneous mitral commissurotomy with good results	Poor late functional results	Age x Mitral Valve Area, Final Mean Mitral Gradient, Valve Calcification x Sex, AF x NYHA Class	609	81%	309	49 (14)	20 years (mean: 10.7 years)
22155702	Holme <sup>139</sup>	2012	Covariate	Harmful	Patients in the Simvastatin and Ezetimibe in Aortic Stenosis trial with asymptomatic mild-to-moderate aortic stenosis	Mortality	Age, Sex, Smoking, LVMI, Bilirubin, HR, CRP	772	38%	78	67.6 (9.6)	4 years (mean: 4.8 years)
21871277	Weiss <sup>140</sup>	2011	Covariate	Harmful	Patients who underwent orthotopic heart transplantation (OHT)	Mortality	Bilirubin, Race, Creatinine Clearance, HF Etiology, Ventricular Assist Device, Age	17079	23.7%	NR	52.1 (11.9)	1 year
15854958	Passman <sup>141</sup>	2005	Covariate	Protective	Cancer patients who were in sinus rhythm before surgery before undergoing major non-cardiac thoracic surgery	Post-operative atrial fibrillation	Age, Sex, HR	856	45.1%	147	64.6 (10.6)	Hospitalization period
17544079	Rogers <sup>142</sup>	2007	Covariate	Harmful	Patients from Veteran's Administration medical centers and private sector hospitals who underwent major general or vascular procedures	Post-operative deep venous thromboembolism	Respiratory and Hemic vs Endocrine, Thoracoabdominal vs Endocrine, Stomach/Intestines vs Endocrine, Mouth/Palate vs Endocrine, Aneurysm vs Endocrine, Work RVU, ASA Physical Status, Disseminated Cancer, Ventilator Dependent, Hernia vs Endocrine, Chemotherapy for Malignancy, Sodium, Transfusion, Emergency, Wound Class, Sex, Hematocrit, Bilirubin, Dyspnea, Albumin	182783	19.8%	1162	60.3 (14.3)	30 days
20453069	Decousus <sup>143</sup>	2011	Covariate	Protective	Patients who were admitted to the hospital (duration > 3 days) for an acute medical illness	In-hospital bleeding within 14 days of admission	Active Ulcer, Prior Bleeding, Platelet Count, Age, Hepatic Failure, Renal Failure, ICU, Catheter, Cancer, Rheumatic Disease, Sex	9388	50.60%	198	NR	Hospitalization period
21818020	Cook <sup>144</sup>	2011	Covariate	Harmful	Patients 16 years old or older with an ICD-9 code	Blunt cerebrovascular	Cervical Spinal Cord Surgery, Cerebellar or	349121	NR	471	NR	Diagnosis

					consistent with blunt cerebrovascular trauma	injury	Brainstem Injury, Thoracic Spinal Cord Injury, Maxillary or Malar Fracture, Chest Abbreviated Injury Scale, Trauma Mortality Prediction Model, Cervical Spine Fracture/Dislocation, Mandible Fracture, Age, Sex, Traumatic Intracranial Hemorrhage					
14980900	Rohrig <sup>145</sup>	2004	Covariate	Protective	Patients undergoing non-cardiac surgery	Operative cardiovascular events (CVE)	Major Vascular Surgery, Gastrointestinal Surgery, Hematopoietic/Lymphatic Surgery, Neurosurgery, Thoracic Surgery, Valve Disease, Carotid Artery Stenosis, Hypovolemia, Arterial HTN, Chronic Renal Failure, Emergency Surgery, CABG, Arrhythmia, Sex, Age	29437	49.4%	5249	52.7 (18.4)	Hospitalization period
17544078	Davenport <sup>146</sup>	2007	Covariate	Protective	Patients from VA medical centers and private-sector hospitals who underwent major general or vascular procedures	Postoperative cardiac adverse events	ASA Physical Status, Age, Type of Operation, Work RVU, Creatinine, Emergency, WBC, Weight Loss, CHF, Sepsis, Ascites, Bleeding Disorder, Platelet Count, Specialty, Impaired Sensorium, Wound Class, CVA/Stroke with Neurological Deficit, Sex, Dyspnea, Albumin	91403	19.80%	1181	60.3 (14.3)	30 days
21217144	Verheugt <sup>147</sup>	2011	Covariate	Protective	Inception cohort of adults with congenital heart disease followed from the age of 18	Adult-onset infective endocarditis up to 40 or 60 years of age	Pulmonary Atresia, Transposition of Great Arteries, Ventricular Septal Defect, Bicuspid Aortic Valve, Univentricular Heart/Double Inlet Left Ventricle, Infective Endocarditis, Other Congenital Heart Defect, Aortic Stenosis, Atrioventricular Septal Defect, Tetralogy of Fallot, Marfan Syndrome, CVA/TIA, Aortic Coarctation,	10210	51%	233	NR	mean: 18.9 years

							Ebstein Malformation, Multiple Congenital Heart Defects, Pulmonary Stenosis, Sex, Arrhythmia					
21051669	Calvillo-King <sup>148</sup>	2010	Covariate	Harmful	Medicare beneficiaries underwent carotid endarterectomy and were otherwise asymptomatic	Death or nonfatal stroke within 30 days of surgery	Severe Disability, Race, Stenosis, CHF, CAD, VHD, Distant Stroke or TIA, Sex	6553	45.1%	197	74.5 (6.6)	30 days
20728962	Barrett <sup>149</sup>	2011	Covariate	Harmful	Patients presenting in the emergency department with symptomatic atrial fibrillation	30-day adverse event (ED return visit, unscheduled hospitalization, cardiovascular complication, or death)	Smoking, Inadequate Ventricular Rate Control, Dyspnea, Beta Blockers, CHF, Edema, HTN, Age, Sex, COPD, Diuretic Use, Palpitations	832	41.1%	651	63.6 (NR)	30 days
20351233	Eichinger <sup>150</sup>	2010	Covariate	Protective	Patients admitted for first unprovoked venous thromboembolism	Risk of recurrent VTE after discontinuation of anticoagulation	PE vs Distal Thrombosis, Proximal vs Distal Thrombosis, Sex, D-Dimer	929	39.5%	176	NR	43.3 months
18922710	Yamaki <sup>151</sup>	2009	Covariate	Protective	Patients with confirmed cardiopulmonary stable pulmonary embolism who were prospectively evaluated using bilateral lower extremity compression ultrasound	Recurrence of symptomatic venous thromboembolism (VTE) and fatal events	Sex, History of DVT, Cancer, Presence of DVT, Inadequate Anticoagulation, Presence of Proximal DVT, Leg Symptoms	203	61.6%	118	62.8 (15.9)	12 months
19766355	Sun <sup>152</sup>	2009	Covariate	Protective	Patients aged 60 or older with an emergency department discharge diagnosis of syncope	Serious clinical event that occurred during the 30- day period after the initial ED evaluation (death, arrhythmias, MI, a new diagnosis of structural heart disease thought to be related to syncope, PE)	Near-Syncope, Abnormal EKG, Troponin, Age, Arrhythmia, Sex, Triage SBP	2584	54%	173	NR	30 days
19486721	Cruz-Gonzalez <sup>153</sup>	2009	Covariate	Harmful	Patients with moderate or severe mitral stenosis (MS) patients who underwent percutaneous mitral valvuloplasty (PMV)	Percutaneous mitral valvuloplasty (PMV) success	Mitral Valve Area, Mitral Regurgitation, Sex, Echo Score, NYHA Class, Age	800	81.9%	544	54.9 (15.26)	Hospitalization period
19546391	Monin <sup>154</sup>	2009	Covariate	Harmful	Patients with asymptomatic aortic stenosis	Death or aortic valve replacement necessitated by symptoms or by a positive exercise test	Peak Velocity, Sex, BNP, Three-Leaflet Valve, Calcification Score, Transaortic Mean Pressure Gradient, Aortic Valve Area, Indexed Valve Area	104	34%	62	72 (NR)	2 years
18573589	Nishiuchi <sup>155</sup>	2008	Covariate	Protective	Patients who experienced an out-of-	Survival	Witness, CPR to Hospital Arrival Time,	1028	22.8%	202	NR	1 month

					hospital cardiac arrest		Ambulance Call to CPR Time, Sex, Age, Ambulance Call to Defibrillation Time					
16525500	Kim <sup>156</sup>	2006	Covariate	Harmful	Patients with newly diagnosed acute leukemia	Fatal intracranial hemorrhage	Sex, Leukemia, WBC, Prothombin Time, Platelets, Promyelocytic Leukemia, Activated Partial Thromboplastin Time, Age, Hemorrhage Score	792	68.3%	41	42 (17)	mean: 3.8 years
16520853	Ivanov <sup>157</sup>	2006	Covariate	Harmful	Patients who underwent cardiac surgery	Adverse event (operative death, a perioperative MI defined by EKG and enzymatic criteria, low cardiac output syndrome, a perioperative stroke, new postoperative renal failure)	Priority Status, CABG, LVEF, Renal Insufficiency, Other Pathology, Age, Sex, CHF, Recent MI, Left Main Disease, PVD, Triple Vessel Disease, Complex Valve, Diabetes, HTN	12683	27%	1243	61 (12)	Hospitalization period
16308250	Yuan <sup>158</sup>	2006	Covariate	Protective	Population of patients enrolled at routine visit	Vertigo	Triglycerides, SBP, CVD, DBP, Sex, Age, Cholesterol, Fasting Blood Glucose	200	54.5%	100	58.2 (11.26)	NA (case control)
16020800	Aujesky <sup>159</sup>	2005	Covariate	Protective	Patients with pulmonary embolism	Mortality	Age, Altered Mental Status, Cancer, SBP, Pulse, Arterial Oxygen Saturation, Temperature, Respiratory Rate, HF, Chronic Lung Disease, Sex	10354	60.4%	953	NR	30 days
9002560	Karlson <sup>160</sup>	1997	Covariate	Protective	Patients with chest pain/suspected acute myocardial infarction in the emergency department (ED)	Mortality	History of CHF, History of Diabetes, Loss of Consciousness, Unspecific Symptoms, Signs of Acute Ischemia, ST Elevation on EKG, Acute Severe CHF, History of HTN, Age, Sex, Initial Degree of Suspicion of Infarction, Normal Admission EKG	5362	45%	604	63 (NR)	1 year
17380222	McDonald <sup>161</sup>	2007	Covariate	Protective	Patients with non-traumatic chest pain or symptoms suggestive of ischemic heart disease admitted to emergency department	Primary end point: death, nonfatal myocardial infarction or revascularization	ST Deviation on EKG, Sex, Acetylsalicylic Acid Taken Before ER, Risk Factors for CAD, Prior CHF	1054	44%	141	61 (16)	30 days

<sup>a</sup> Directionality of the predictive effect of female sex cannot be determined without considering the following interaction term with sex:  $\log(\text{age})^2 \times \text{female}$ , diabetes  $\times$  female, LVH  $\times$  male,  $\log(\text{age}) \times \text{female}$ .

<sup>b</sup> Directionality cannot be determined for this model without considering the following interaction term with sex: diabetes  $\times$  female.

- <sup>c</sup> Directionality cannot be determined for this model without considering the following interaction terms with sex:  $\log(\text{age}) \times \text{female}$ , diabetes  $\times$  female.
- <sup>d</sup> Directionality cannot be determined for this model because sex is included in this model in an interaction term only: age  $\times$  sex.
- <sup>e</sup> Directionality cannot be determined for this model without considering the following interaction terms with sex:  $\log(\text{age})^2 \times \text{female}$ , diabetes  $\times$  female,  $\log(\text{age}) \times \text{female}$ .
- <sup>f</sup> Directionality cannot be determined for this model because sex is included in this model in an interaction term only: sex  $\times$  waist circumference.
- <sup>g</sup> Directionality cannot be determined for this model without considering the following interaction terms with sex: female  $\times$  BSA function, female  $\times$  mitral valve repair, female  $\times$  mitral valve replacement.
- <sup>h</sup> Directionality cannot be determined for this model because sex is included in this model in an interaction term only: valve calcification  $\times$  sex.

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