

## Supplementary Online Content

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**eTable 1.** List of Covariates Included for Calculation of Multivariate Adjusted Hazard Ratios (aHR)

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**eFigure 1.** Relative Risk (Hazard Ratios, Unadjusted and Adjusted) for Study Outcomes in High-Risk AF Patients (CHA<sub>2</sub>DS<sub>2</sub>-VASc Score ≥2) Treated With OAC Plus AP or OAC Alone (Reference) Over 12 Months (Intent-to-Treat Analyses)

**eFigure 2.** Relative Risk (Hazard Ratios, Unadjusted and Adjusted) for Study Outcomes in High-Risk AF Patients (CHA<sub>2</sub>DS<sub>2</sub>-VASc Score ≥2) Treated with OAC Plus AP or OAC Alone (Reference) Over 3 Months (Intent-to-Treat Analyses)

**eAppendix.** Group Information

This supplementary material has been provided by the authors to give readers additional information about their work.

**eTable 1. List of Covariates Included for Calculation of Multivariate Adjusted Hazard Ratios (aHR)**

Patient characteristics
Sex
Age
Race/ethnicity
Pulse rate (bpm)
Diastolic blood pressure (mmHg)
Systolic blood pressure (mmHg)
Medical history
Heart failure
Acute coronary syndromes (MI or unstable angina)
Coronary artery disease
Diabetes mellitus (type 1 or 2)
Bleeding (history)
Carotid occlusive disease
Coronary artery bypass graft
Dementia
Hypercholesterolemia
Hypertension (history)
Pulmonary embolism or deep vein thrombosis
Stroke/transient ischemic attack
Systemic embolization (history)
Cirrhosis
Renal disease severe

Alcohol consumption
Smoker
Medication use at study entry
NSAIDs/COX-2 inhibitor (prior use)
Calcium channel blockers
Proton pump inhibitors
Digoxin
Statin
Beta blocker
Oral antidiabetic agent
Digitalis
Alfa blocker
Angiotensin-converting enzyme inhibitor (ACEi)
Angiotensin II receptor blocker (ARB)
Aldosterone antagonist
Nitrates
Insulin
Loop or other diuretics
Bisoprolol
Year/quarter study entry

**eTable 2. Baseline Clinical Characteristics of Patients at High Risk for Stroke**

**(CHA<sub>2</sub>DS<sub>2</sub>-VASc score ≥2)**

<b>Parameter</b>	<b>OAC + AP (N = 2735)</b>	<b>OAC Alone (17,892)</b>
Sex male, n (%)	1630 (59.6%)	8392 (46.9%)
Age in years, mean (SD)	71.3 (9.7)	72.6 (9.5)
BMI, kg/m <sup>2</sup> , mean (SD)	28.1 (5.6)	27.9 (5.8)
SBP/DBP, mean	133.6/79.8	135.5/80.5
Pulse rate, bpm, mean (SD)	90.9 (26.8)	91.3 (26.6)
LVEF %, mean (SD)	52.6 (13.9)	56.6 (12.5)
CHA <sub>2</sub> DS <sub>2</sub> -VASc, median	4	3
Medical history, n (%)		
CHF	745 (27.2%)	3405 (19.0%)
CAD	1123 (41.1%)	1960 (11.0%)
ACS	650 (23.8%)	877 (4.9%)
Carotid occlusive disease	142 (5.2%)	404 (2.3%)
DVT/PE	101 (3.7%)	541 (3.0%)
CABG	207 (7.6%)	231 (1.3%)
Stroke/TIA	493 (18.0%)	1832 (10.2%)
Bleeding	80 (2.9%)	307 (1.7%)
Hypertension	2342 (85.6%)	14,761 (82.5%)
Hypercholesterolemia	1,407 (51.4%)	6848 (38.3%)
DM, type 1 or 2	905 (33.1%)	4137 (23.1%)
CKD, moderate–severe	395 (14.4%)	1989 (11.1%)

ACS, acute coronary syndrome; CABG, coronary artery bypass graft; CAD, coronary artery disease; CHF, congestive heart failure; CKD, chronic kidney disease; DBP, diastolic blood pressure; DM, diabetes mellitus; DVT, deep vein thrombosis; LVEF, left ventricular ejection fraction; PE, pulmonary embolism; SBP, systolic blood pressure; TIA, transient ischemic attack.

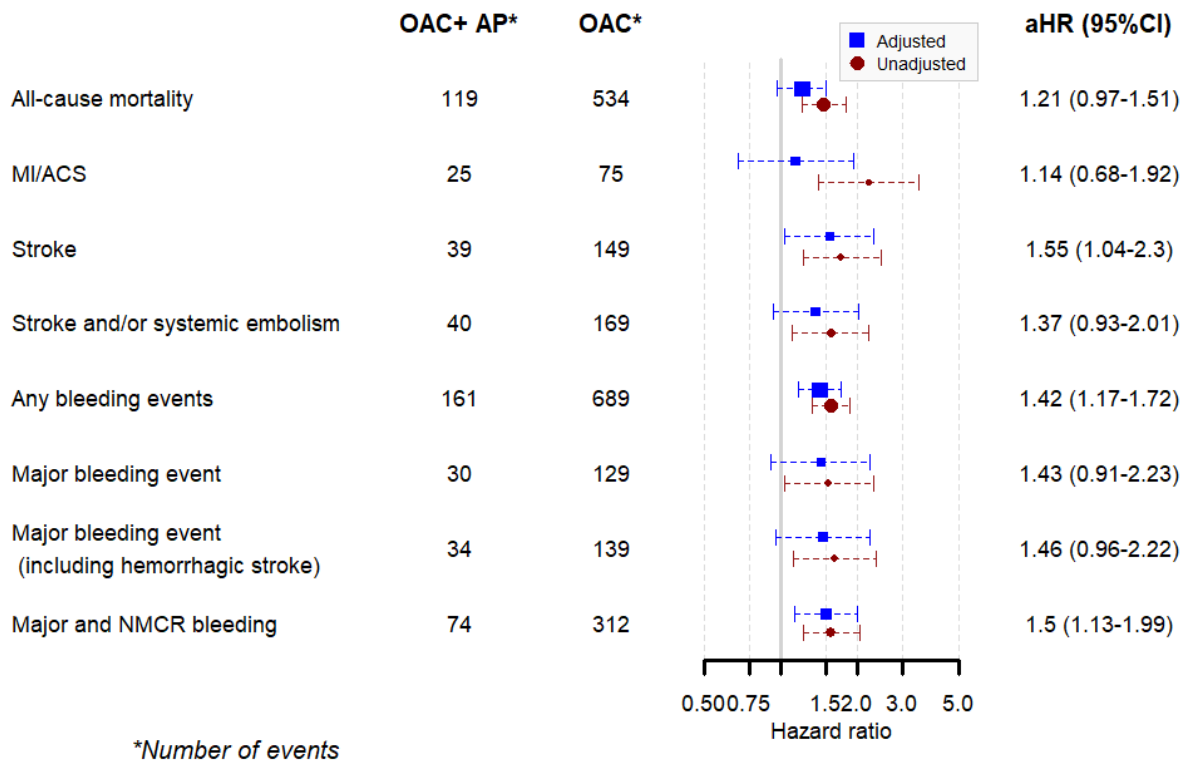
**eTable 3. Falsification Analysis—Non-CV Mortality Calculated over 12-month Follow-up**

	OAC + AP			OAC Alone			Crude HR (95% CI)	aHR (95% CI)
	Events	Subjects	Risk/1000 Pts	Events	Subjects	Risk/1000 Pts		
ITT (multivariate)	24	2541	945	200	17,673	11.32	0.85 (0.55–1.29)	0.76 (0.48–1.22)
ITT (PS matched)	23	2380	966	28	2380	11.76		0.83 (0.48–1.44)

ITT, intent-to-treat analysis with variable follow-up; PS, propensity score.

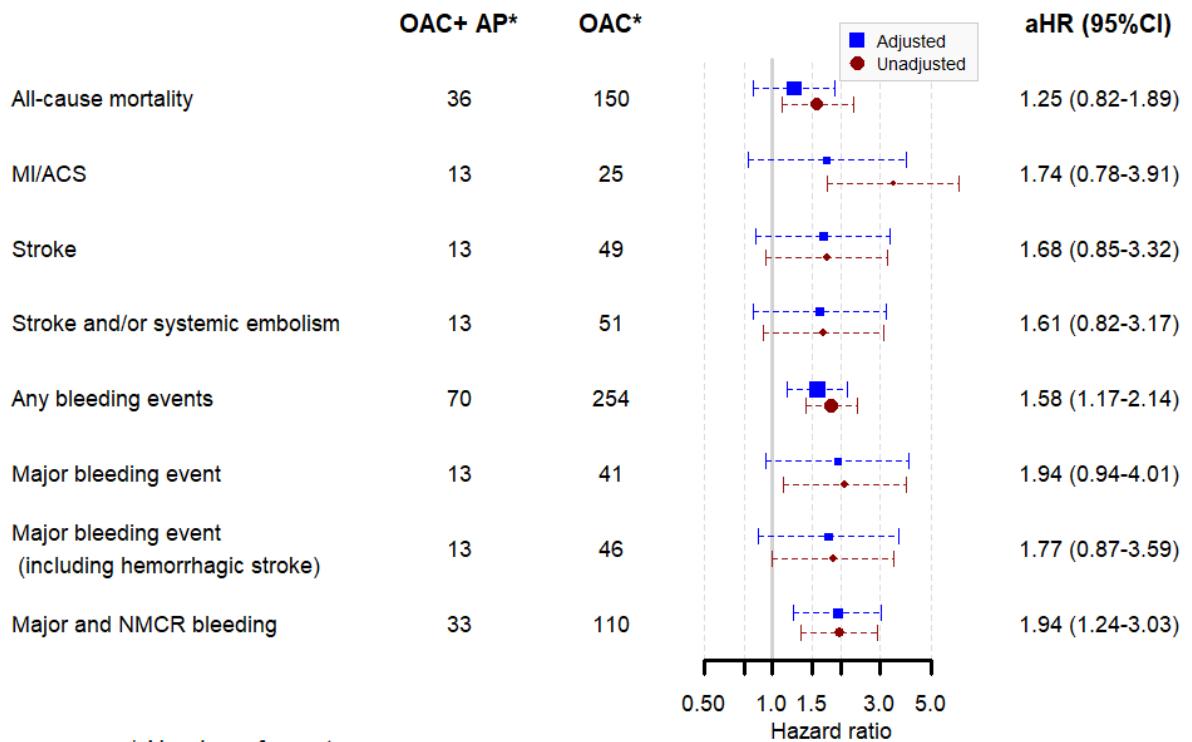
**eFigure 1. Relative Risk (Hazard Ratios, Unadjusted and Adjusted<sup>1</sup>) for Study Outcomes in High-Risk AF Patients (CHA<sub>2</sub>DS<sub>2</sub>-Vasc Score ≥2) Treated with OAC Plus AP or OAC Alone (Reference) over 12 Months (Intent-to-Treat Analyses).**

<sup>1</sup>Adjusted for 40 covariates as shown in Supplementary material, Table S1.



**eFigure 2. Relative Risk (Hazard Ratios, Unadjusted and Adjusted<sup>1</sup>) for Study Outcomes in High-Risk AF Patients (CHA<sub>2</sub>DS<sub>2</sub>-Vasc Score ≥2) Treated with OAC Plus AP or OAC Alone (Reference) over 3 Months (Intent-to-Treat Analyses).**

<sup>1</sup>Adjusted for 40 covariates as shown in Supplementary material, Table S1.





## **eAppendix. Group Information**

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