

SUPPLEMENTARY MATERIAL

Table S1. Distribution of patients by country with information on baseline OAC

Country	N	Patients on OAC (row %)
Japan	2836	80.9
United Kingdom	2172	70.4
Germany	1364	81.9
Russia	1181	57.3
Spain	1152	78.5
Ukraine	1123	52.7
China	1035	29.1
Italy	1035	89.8
Belgium	998	86.3
Poland	986	84.6
Czech Republic	941	78.0
France	870	80.3
Thailand	865	55.8
South Korea	845	65.0
Hungary	768	84.8
India	739	35.1
Chile	592	88.0
Sweden	560	85.5
Argentina	551	58.4
United States	534	72.7
Brazil	507	57.2
Netherlands	481	95.0
Mexico	454	62.3

Canada	408	67.9
Egypt	367	89.7
Turkey	363	69.2
Australia	352	70.2
South Africa	330	77.6
United Arab Emirates	284	60.9
Denmark	213	85.5
Singapore	176	65.9
Austria	164	79.9
Finland	119	88.2
Norway	114	88.6
Switzerland	72	93.1

Table S2. Baseline characteristics by treatment at baseline

Baseline characteristics	Patients treated with OAC			Patients not treated with OAC (N = 7162)
	FXaI (N = 7694)	DTI (N = 2090)	VKA (N = 8605)	
Body mass index, median (Q1; Q3), kg/m²	26.6 (23.7;30.4)	27.3 (24.4;31.6)	27.6 (24.4;31.6)	26.4 (23.5;30.1)
Systolic blood pressure, median (Q1; Q3), mmHg	134.0 (120.0;146.0)	131.0 (120.0;148.0)	134.0 (120.0;147.0)	130.0 (120.0;145.0)
Diastolic blood pressure, median (Q1; Q3), mmHg	80.0 (70.0;88.0)	80.0 (70.0;90.0)	80.0 (70.0;90.0)	80.0 (70.0;88.0)
Pulse, median (Q1; Q3), bpm	84.0 (70.0;106.0)	85.0 (70.0;110.0)	85.0 (72.0;104.0)	83.0 (70.0;105.0)
Care setting specialty at diagnosis, n (%)				
Internal medicine/Neurology/Geriatrics	1483 (19.3)	339 (16.2)	1819 (21.1)	1493 (20.8)
Cardiology	5387 (70.0)	1546 (74.0)	5227 (60.7)	4632 (64.7)
Primary care/general practice	824 (10.7)	205 (9.8)	1559 (18.1)	1037 (14.5)
Care setting location at diagnosis, n (%)				
Hospital	3610 (46.9)	1197 (57.3)	4880 (56.7)	4588 (64.1)
Office/Anticoagulation clinic/Thrombosis centre	3375 (43.9)	700 (33.5)	2711 (31.5)	1795 (25.1)
Emergency room	709 (9.2)	193 (9.2)	1014 (11.8)	779 (10.9)
Heavy alcohol consumption, n (%)	109 (1.7)	33 (1.9)	130 (1.8)	126 (2.1)
Current smoker, n (%)	595 (8.5)	189 (9.8)	681 (8.6)	636 (9.7)

Table S3. Unadjusted and adjusted¹ hazard ratios and corresponding 95% confidence intervals for selected outcomes at two years of follow-up by treatment at baseline

Outcome	Treatment comparison							
	OAC vs No OAC (ref.)				NOAC vs VKA (ref.)			
	Unadjusted		Adjusted ¹		Unadjusted		Adjusted ¹	
	HR (95% CI)	p-value	HR (95% CI)	p-value	HR (95% CI)	p-value	HR (95% CI)	p-value
All-cause mortality	0.74 (0.68-0.81)	<0.0001	0.82 (0.74-0.91)	0.0002	0.74 (0.67-0.82)	<0.0001	0.79 (0.70-0.89)	0.0001
Non-haemorrhagic stroke/SE	0.67 (0.56-0.81)	<0.0001	0.71 (0.57-0.88)	0.0021	0.82 (0.65-1.03)	0.0877	0.96 (0.73-1.25)	0.7360
Major bleeding	1.51 (1.22-1.88)	0.0002	1.46 (1.15-1.86)	0.0022	0.70 (0.57-0.85)	0.0003	0.77 (0.61-0.98)	0.0363

¹Obtained using an overlap-weighted Cox model. Variables included in the weighting scheme are: country and cohort enrolment, sex, age, ethnicity, type of AF, care setting speciality and location, congestive heart failure, acute coronary syndromes, vascular disease, carotid occlusive disease, prior stroke/TIA/SE, prior bleeding, VTE, hypertension, hypercholesterolemia, diabetes, cirrhosis, moderate to severe CKD, dementia, hyperthyroidism, hypothyroidism, current smoking, heavy alcohol consumption, BMI, heart rate, systolic and diastolic blood pressure at diagnosis, and baseline antiplatelet use.

Table S4. Unadjusted and adjusted¹ hazard ratios and corresponding 95% confidence intervals for selected outcomes at two years of follow-up by treatment at baseline

Outcome	Treatment comparison							
	NOAC vs No OAC (ref.)				VKA vs No OAC (ref.)			
	Unadjusted		Adjusted ¹		Unadjusted		Adjusted ¹	
	HR (95% CI)	p-value	HR (95% CI)	p-value	HR (95% CI)	p-value	HR (95% CI)	p-value
All-cause mortality	0.64 (0.58-0.71)	<0.0001	0.67 (0.59-0.77)	<0.0001	0.86 (0.78-0.95)	0.0035	0.93 (0.82-1.04)	0.2088
Non-haemorrhagic stroke/SE	0.61 (0.49-0.76)	<0.0001	0.65 (0.50-0.86)	0.0020	0.75 (0.60-0.93)	0.0084	0.75 (0.58-0.98)	0.0315
Major bleeding	1.26 (0.99-1.61)	0.0627	1.10 (0.82-1.47)	0.5239	1.80 (1.43-2.28)	<0.0001	1.86 (1.42-2.44)	<0.0001

¹Obtained using an overlap-weighted Cox model. Variables included in the weighting scheme are: country and cohort enrolment, sex, age, ethnicity, type of AF, care setting speciality and location, congestive heart failure, acute coronary syndromes, vascular disease, carotid occlusive disease, prior stroke/TIA/SE, prior bleeding, VTE, hypertension, hypercholesterolemia, diabetes, cirrhosis, moderate to severe CKD, dementia, hyperthyroidism, hypothyroidism, current smoking, heavy alcohol consumption, BMI, heart rate, systolic and diastolic blood pressure at diagnosis, and baseline antiplatelet use.

Table S5. Unadjusted and adjusted¹ hazard ratios and corresponding 95% confidence intervals for selected outcomes at two years of follow-up by OAC treatment at baseline. The reference considered is the treatment reported as second

Outcome	Unadjusted HR (95% CI)	Adjusted ¹ HR (95% CI)
Baseline treatment		
All-cause mortality		
DTI vs VKA	0.68 (0.57-0.82)	0.83 (0.71-0.98)
FXaI vs VKA	0.76 (0.68-0.85)	0.77 (0.65-0.92)
FXaI vs DTI	1.11 (0.92-1.35)	0.92 (0.80-1.06)
Non-haemorrhagic stroke/SE		
DTI vs VKA	0.90 (0.62-1.30)	1.01 (0.74-1.39)
FXaI vs VKA	0.80 (0.63-1.02)	0.92 (0.67-1.27)
FXaI vs DTI	0.89 (0.61-1.31)	0.91 (0.58-1.44)
Major bleeding		
DTI vs VKA	0.53 (0.36-0.77)	0.68 (0.47-0.98)
FXaI vs VKA	0.75 (0.61-0.92)	0.84 (0.63-1.12)
FXaI vs DTI	1.42 (0.95-2.11)	1.24 (0.90-1.73)

¹Obtained using an overlap-weighted Cox model. Variables included in the weighting scheme are: country and cohort enrolment, sex, age, ethnicity, type of AF, care setting speciality and location, congestive heart failure, acute coronary syndromes, vascular disease, carotid occlusive disease, prior stroke/TIA/SE, prior bleeding, VTE, hypertension, hypercholesterolemia, diabetes, cirrhosis, moderate to severe CKD, dementia, hyperthyroidism, hypothyroidism, current smoking, heavy alcohol consumption, BMI, heart rate, systolic and diastolic blood pressure at diagnosis, and baseline antiplatelet use.

Figure S1. Propensity score comparing OAC vs no OAC

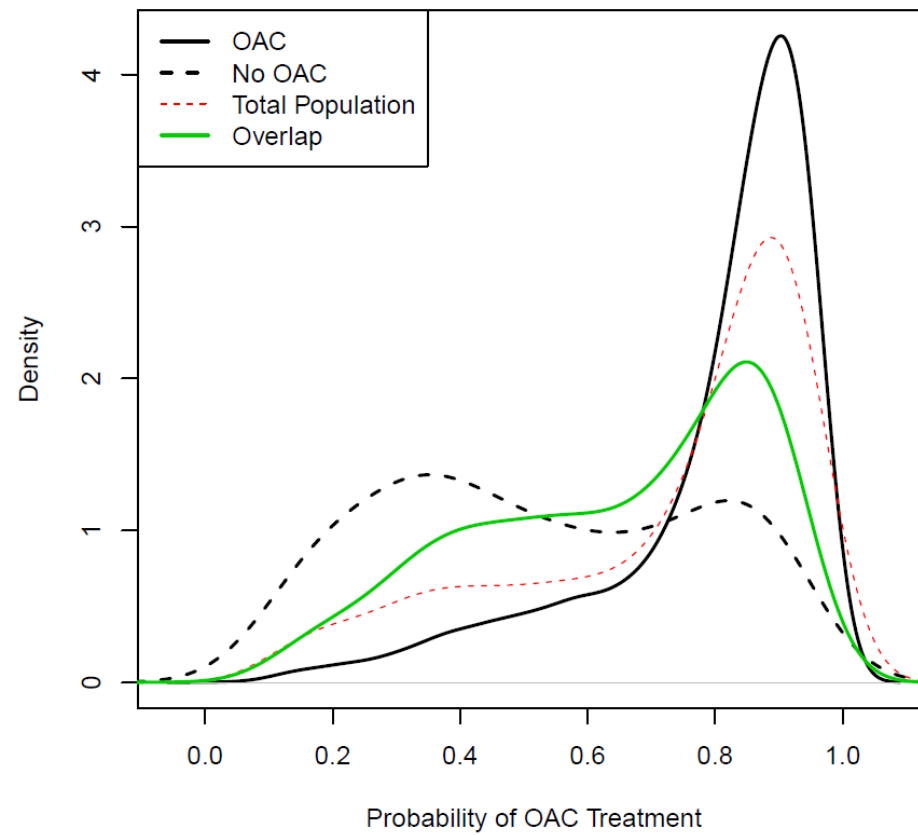


Figure S2. Propensity score comparing NOAC vs VKA

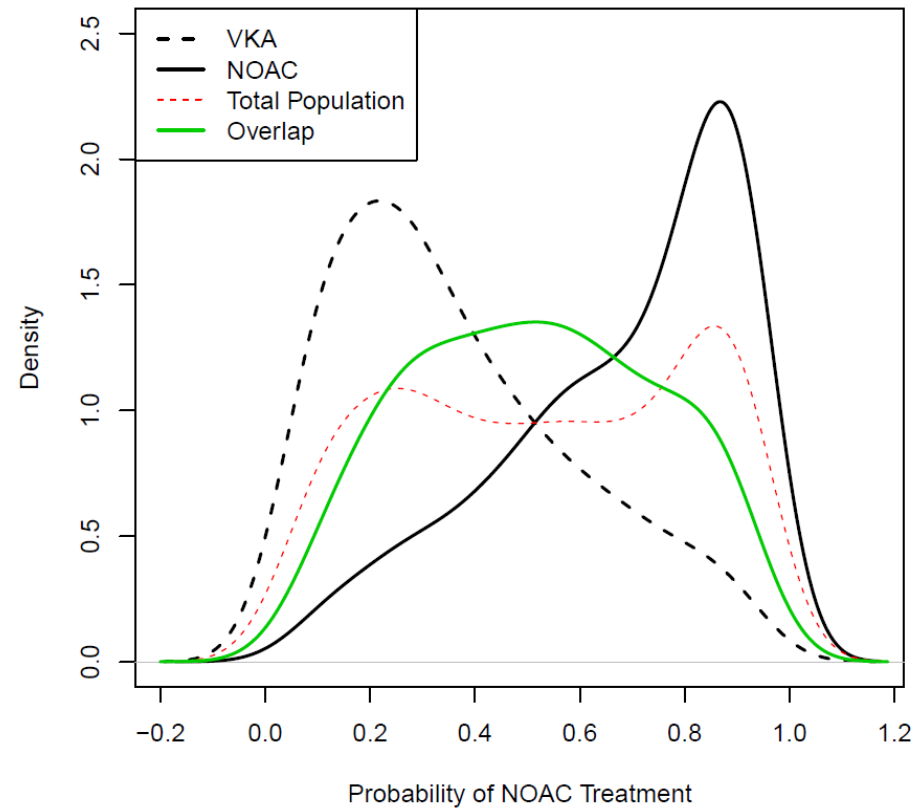


Figure S3. Balance of variables associated with OAC treatment in reference to no OAC treatment at baseline before and after propensity score weighting

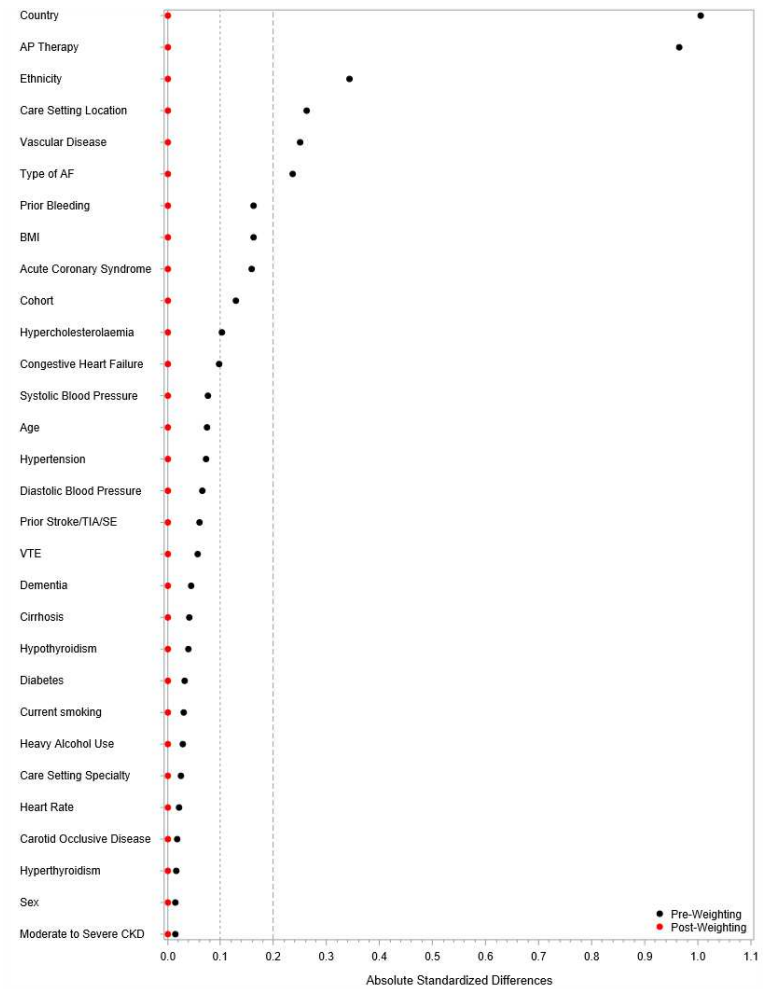


Figure S4. Balance of variables associated with NOAC treatment in reference to VKA treatment at baseline before and after propensity score weighting

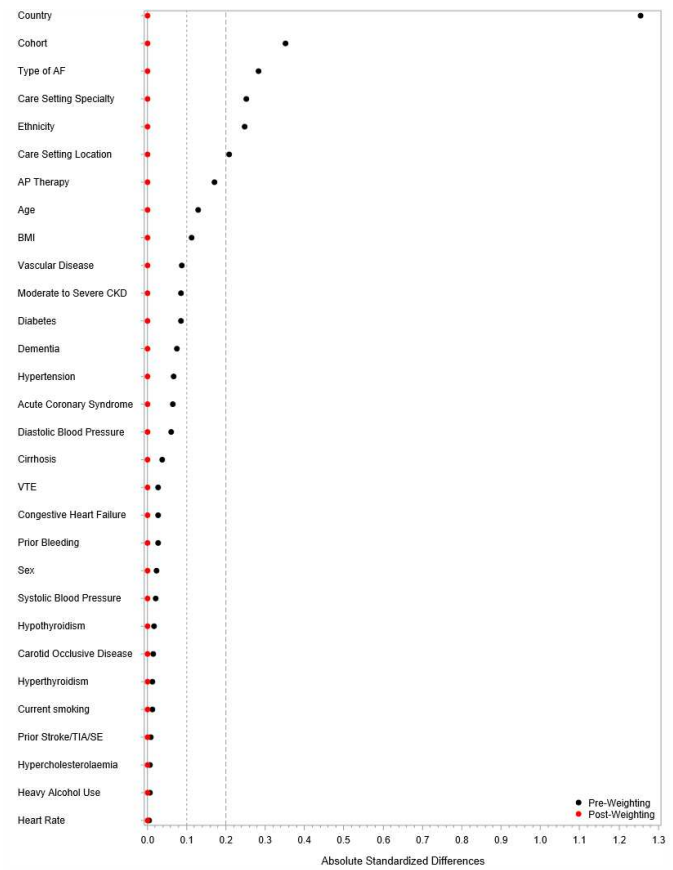


Figure S5. Flowchart for the selection of the study population

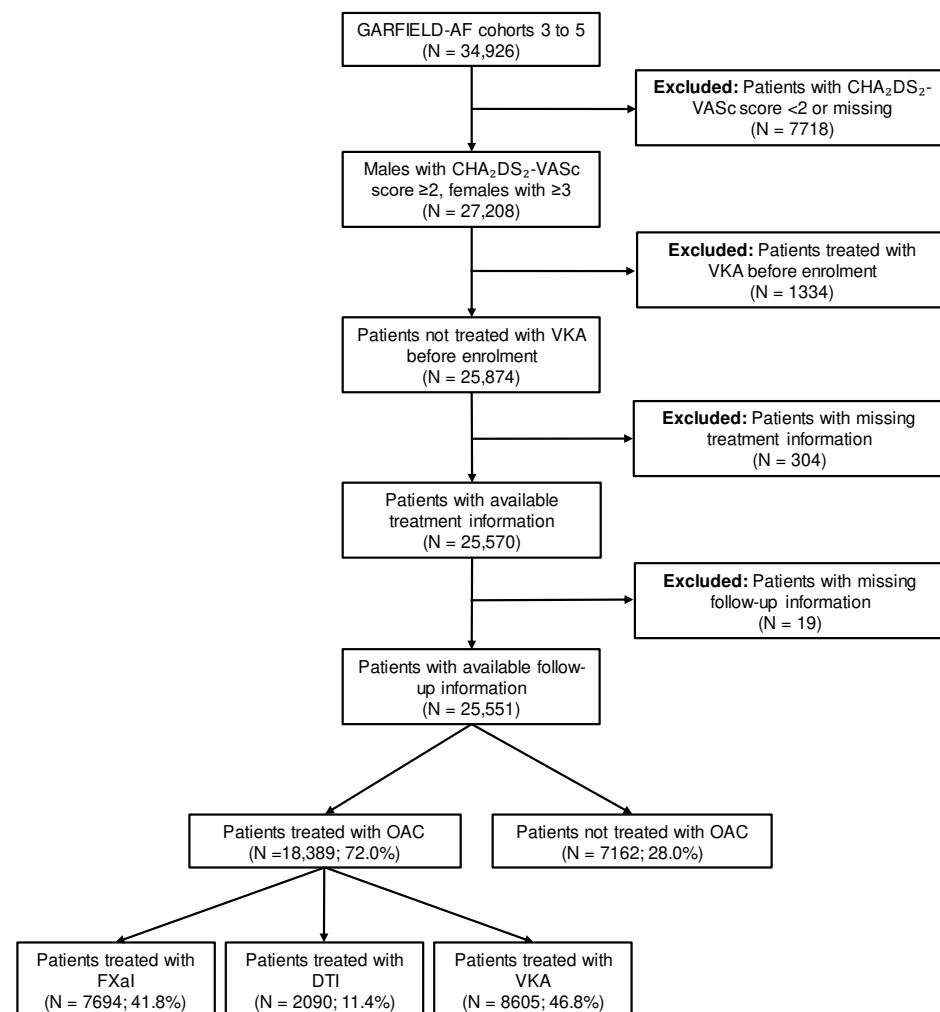
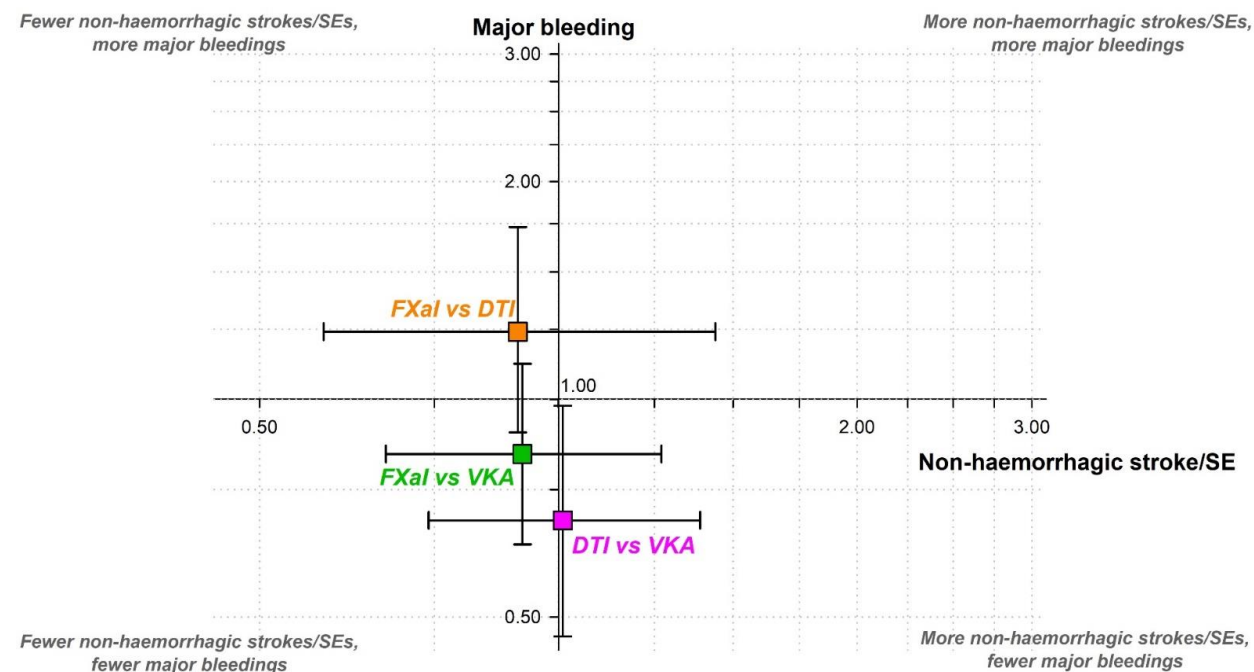


Figure S6. Adjusted¹ hazard ratios and corresponding 95% confidence intervals for selected outcomes at 2 years of follow-up by OAC treatment at baseline. The reference considered is the treatment reported as second



¹Obtained using an overlap-weighted Cox model. Variables included in the weighting scheme are: country and cohort enrolment, sex, age, ethnicity, type of AF, care setting speciality and location, congestive heart failure, acute coronary syndromes, vascular disease, carotid occlusive disease, prior stroke/TIA/SE, prior bleeding, VTE, hypertension, hypercholesterolemia, diabetes, cirrhosis, moderate to severe CKD, dementia, hyperthyroidism, hypothyroidism, current smoking, heavy alcohol consumption, BMI, heart rate, systolic and diastolic blood pressure at diagnosis, and baseline antiplatelet use.

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