

Appendix 1. Sensitivity analysis additionally adjusted for diabetes, hypertension, hypercholesterolemia, PVD and CVD: Association of risk factors with hazard of atrial fibrillation in patients, who received allopurinol with no baseline atrial fibrillation before the index date of allopurinol episode

	Univariate		Multivariable-adjusted (model 1)		Multivariable-adjusted (model 2)	
	HR (95% CI)	P-value	HR (95% CI)	P-value	HR (95% CI)	P-value
Age (in years)						
65 - <75	Ref		Ref		Ref	
75 - <85	1.47 (1.31, 1.66)	<0.0001	1.48 (1.31, 1.67)	<0.0001	1.47 (1.31, 1.67)	<0.0001
≥85	2.12 (1.83, 2.46)	<0.0001	2.10 (1.80, 2.45)	<0.0001	2.10 (1.80, 2.45)	<0.0001
Gender						
Male	Ref		Ref		Ref	
Female	1.04 (0.93, 1.15)	0.50	0.93 (0.84, 1.04)	0.23	0.93 (0.84, 1.04)	0.22
Race						
White	Ref		Ref		Ref	
Black	0.83 (0.70, 0.97)	0.02	0.80 (0.68, 0.94)	0.007	0.79 (0.68, 0.93)	0.005
Other	0.69 (0.58, 0.83)	<0.0001	0.65 (0.54, 0.77)	<0.0001	0.64 (0.54, 0.77)	<0.0001
Diabetes	1.24 (1.12, 1.38)	<0.0001	1.31 (1.17, 1.47)	<0.0001	1.31 (1.17, 1.47)	<0.0001
Hypertension	1.21 (0.97, 1.51)	0.08	1.13 (0.90, 1.42)	0.30	1.13 (0.90, 1.42)	0.28
PVD	1.44 (1.29, 1.62)	<0.0001	1.30 (1.15, 1.47)	<0.0001	1.30 (1.15, 1.47)	<0.0001
CVD	1.24 (1.09, 1.41)	0.0010	1.10 (0.96, 1.26)	0.15	1.10 (0.96, 1.26)	0.16
Hypercholesterolemia	0.90 (0.80, 1.02)	0.09	0.85 (0.74, 0.96)	0.0096	0.85 (0.74, 0.96)	0.009
Statins	0.95 (0.72, 1.24)	0.69	0.88 (0.66, 1.17)	0.37	0.88 (0.66, 1.17)	0.36
Beta blockers	1.58 (1.22, 2.05)	0.0005	1.63 (1.24, 2.14)	0.0005	1.62 (1.24, 2.13)	0.0005
Diuretics	1.09 (0.85, 1.40)	0.51	0.99 (0.75, 1.29)	0.91	0.98 (0.75, 1.29)	0.90
ACE inhibitor	0.99 (0.72, 1.36)	0.95	0.95 (0.69, 1.33)	0.77	0.95 (0.69, 1.32)	0.77
Allopurinol	0.86 (0.77, 0.97)	0.01	0.83 (0.74, 0.93)	0.002	-	-
Allopurinol duration*						
0 days	Ref		-	-	Ref	
1-180 days	0.94 (0.80, 1.11)	0.46	-	-	0.92 (0.78, 1.09)	0.32
181 days -2 years	0.90 (0.77, 1.04)	0.16	-	-	0.86 (0.74, 0.99)	0.04
> 2 years	0.68 (0.54, 0.65)	0.0009	-	-	0.65 (0.52, 0.82)	0.0002

HR, Hazard ratio; CI, confidence interval; Ref, referent category

*based on person day count

Model 1 = Allopurinol Use+ age + race + gender + Diabetes + hypertension + PVD + CVD+ Hypercholesterolemia +beta blockers + diuretics + ACE inhibitors + Statins

Model 2 = Allopurinol duration + age + race + gender + Diabetes + hypertension + PVD + CVD + Hypercholesterolemia + beta blockers + diuretics + ACE inhibitors + Statins