

Appendix 2. Sensitivity analysis additionally adjusted for diabetes, hypertension and hypercholesterolemia: 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 (model1)		Multivariable-adjusted (model2)	
	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	1.51 (1.34, 1.70)	<0.0001	1.50 (1.33, 1.70)	<0.0001
≥85	2.12 (1.83,2.46)	<0.0001	2.20 (1.89, 2.57)	<0.0001	2.20 (1.89, 2.57)	<0.0001
Gender						
Male	Ref		Ref		Ref	
Female	1.04 (0.93, 1.15)	0.50	0.93 (0.83, 1.04)	0.19	0.93 (0.83, 1.04)	0.18
Race						
White	Ref		Ref		Ref	
Black	0.83 (0.70, 0.97)	0.02	0.80 (0.68, 0.94)	0.008	0.80 (0.68, 0.94)	0.006
Other	0.69 (0.58, 0.83)	<0.0001	0.64 (0.54, 0.77)	<0.0001	0.64 (0.53, 0.76)	<0.0001
Diabetes	1.24 (1.12, 1.38)	<0.0001	1.35 (1.21, 1.51)	<0.0001	1.35 (1.21, 1.51)	<0.0001
Hypertension	1.21 (0.97, 1.51)	0.08	1.18 (0.94, 1.47)	0.16	1.18 (0.94, 1.48)	0.15
Hypercholesterol emia	0.90 (0.80, 1.02)	0.09	0.87 (0.77, 0.99)	0.0332	0.87 (0.77, 0.99)	0.03
Statins	0.95 (0.72, 1.24)	0.69	0.88 (0.66, 1.17)	0.38	0.88 (0.66, 1.17)	0.37
Beta blockers	1.58 (1.22, 2.05)	0.0005	1.63 (1.24, 2.14)	0.0004	1.63 (1.24, 2.14)	0.0004
Diuretics	1.09 (0.85, 1.40)	0.51	0.98 (0.75, 1.29)	0.90	0.98 (0.75, 1.29)	0.90
ACE inhibitor	0.99 (0.72,1.36)	0.95	0.94 (0.68,1.31)	0.72	0.94 (0.68,1.31)	0.72
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)	0.33
181 days -2 years	0.90 (0.77,1.04)	0.16	-	-	0.85 (0.73, 0.99)	0.04
> 2 years	0.68 (0.54, 0.65)	0.0009	-	-	0.65 (0.52, 0.81)	0.0002

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

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