Appendix 1 (as submitted by the authors): Visualization of information provided by NT-

proBNP measurement assessed by analyzing pre- and post-test probabilities.

A) Measures of added predictive value <u>(fraction of new information from NT-proBNP)</u>

pre <- predict(baseline_model, type='fitted') #pre-test prob
post <- predict(marker_model, type='fitted') #post-test prob</pre>

 $a <- baseline_model$stats['Model L.R.']$ $b <- marker_model$stats['Model L.R.']$ c <- a/b d <- 1-c $e <- baseline_model$stats['R2']$ $f <- marker_model$stats['R2']$ g <- var(pre) h <- var(post) i <- g/h j <- 1-i br2 <- function(p) var(p) / (var(p) + sum(p * (1 - p)) / length(p))) k <- br2(pre) l <- br2(post) m <- k/ln <- 1-m

Pre_test_LR_chi2 <- round(a, 2) #105.4 Post_test_LR_chi2 <- round(b, 2) #125.0 Adequacy_of_base_model <- round(c, 2) #0.84

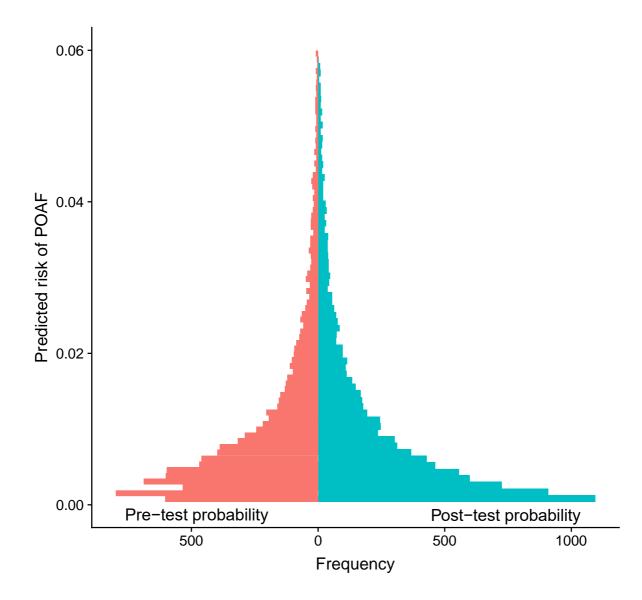
Fraction_of_new_information_from_BNP <- round(d, 2) #0.16</pre>

Pre_test_R2 <- round(e, 3) #0.093 Post_test_R2 <- round(f, 3) #0.11 Variance_of_pre_test_risk <- round(g, 6) # 0.000166 Variance_of_post_test_risk <- round(h, 6) # 0.000195 Relative_explained_variation_1 <- round(i, 2) #0.85

<u>Fraction_of_new_information_discrimination <-round(j, 2)</u> **#0.15** Pre_test_fraction_explained_risk <- round(k, 4) #0.015 Post_test_fraction_explained_risk <- round(l, 4) #0.0177 Relative_explained_variation_2 <-round(m, 2) #0.85

Fraction_of_new_information <- round(n, 2) #0.15</pre>

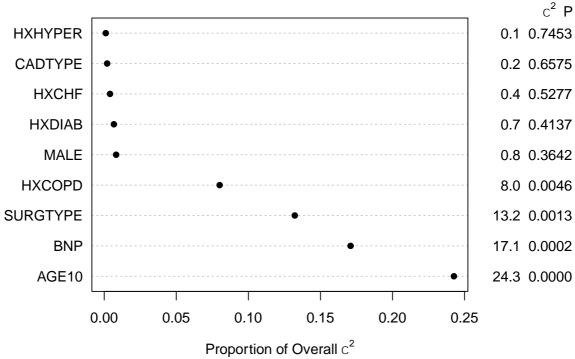
Appendix to: Szczeklik W, LeManach Y, Fronczek J, et al. Preoperative levels of natriuretic peptides and the incidence of postoperative atrial fibrillation after noncardiac surgery: a prospective cohort study. *CMAJ* 2020. doi: 10.1503/cmaj.200840. Copyright © 2020 The Author(s) or their employer(s). To receive this resource in an accessible format, please contact us at cmajgroup@cmaj.ca. **B**) Histograms of predicted probabilites before (pre-test) and after (post-test) adding NT-proBNP to the baseline logistic regression model for POAF



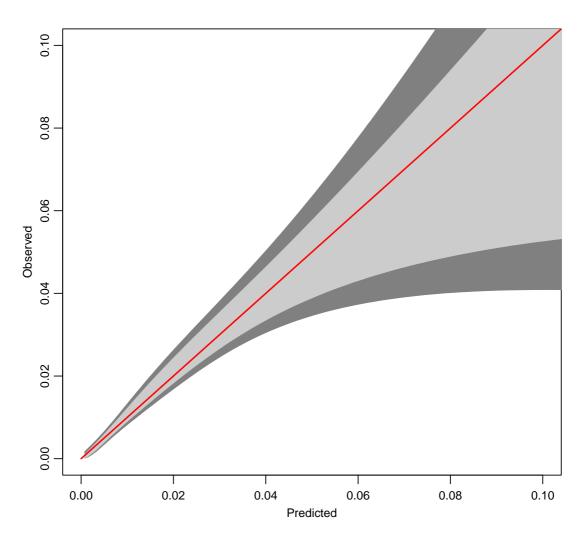
Index	Original	Training	Test	Optimism	Corrected	\overline{n}
	Sample	Sample	Sample		Index	
D_{xy}	0.5872	0.6080	0.5663	0.0417	0.5455	500
R^2	0.1102	0.1210	0.1000	0.0210	0.0892	500
Intercept	0.0000	0.0000	-0.4235	0.4235	-0.4235	500
Slope	1.0000	1.0000	0.8908	0.1092	0.8908	500
E_{\max}	0.0000	0.0000	0.1184	0.1184	0.1184	500
D	0.0127	0.0139	0.0115	0.0024	0.0102	500
U	-0.0002	-0.0002	0.0002	-0.0004	0.0002	500
Q	0.0129	0.0141	0.0113	0.0028	0.0101	500
B	0.0109	0.0108	0.0109	-0.0001	0.0109	500
g	1.4576	1.5704	1.3833	0.1871	1.2705	500
g_p	0.0126	0.0130	0.0121	0.0010	0.0117	500

C) Biomarker model internal validation (bootstrap, n=500)

D) Relative importance of predictors in the biomarker model

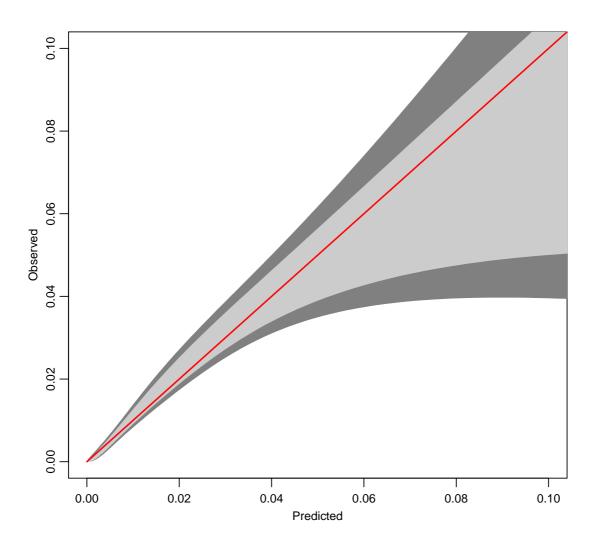


E) Calibration belt of the baseline model (without NT-proBNP)



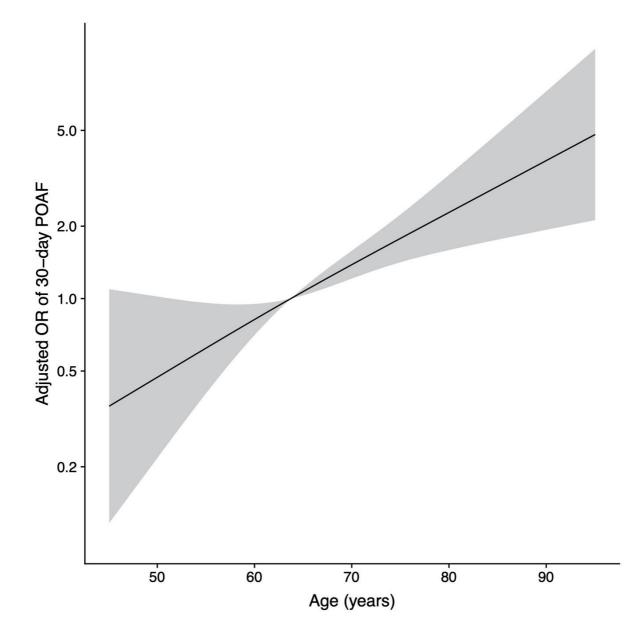
Footnote: The bisector (red line) represents perfect calibration; light grey area represents 80% confidence intervals; dark grey area represents 95% confidence intervals.

F) Calibration belt of the biomarker model (after adding NT-proBNP to the baseline model)



Footnote: The bisector (red line) represents perfect calibration; light grey area represents 80% confidence intervals; dark grey area represents 95% confidence intervals.

G) Adjusted ORs for different values of age



ONLINE TABLES

Supplementary Table 1. Literature review of studies describing incidence of POAF in

patients undergoing non-cardiac surgeries

Study	Design	Population	Inclusion of urgent/emergent procedures	Exclusion of patients with previous history of AF	Number of events	Incidence		
		Non	-cardiac, thoracic					
Amar, 2000(1)	Prospective, interventional	163	Not specified	Yes	40	24.5% *		
Passman, 2005(2)	Retrospective	856	No	Yes	147	17.2%		
Roselli, 2005(3)	Retrospective	604	Not specified	No	113	18.7%		
Cardinale, 2007(4)	Prospective	400	No	Yes	72	18.0%		
Riber, 2012(5)	Prospective, interventional	120	No	Yes	38	31.7%		
Rao, 2012(6)	Retrospective	997	Not specified	Yes	209	21.0%		
Ciszewski, 2013	Prospective	117	Not specified	Yes	19	16.2%		
Berry, 2014(7)	Retrospective	1 412	Not specified	Yes	232	16.4%		
Gialdini, 2014(8)	Registry	35 992	Not specified	Yes	1337	3.7%		
Cardinale, 2016 [‡] (9)	Prospective, interventional	911	Not specified	Yes	75	8.2%		
Lee, 2016(10)	Retrospective	4 662	No	Yes	555	11.9%		
Chin, 2016(11)	Retrospective	583	No	Yes	63	10.8%		
Butt, 2018 (12)**	Registry	1 520 109	Not specified	Yes	6048	0.4%		
Nielsen, 2004(13)	Retrospective	200	No	No	78	39.0%		
Imperatori, 2012(14)	Prospective	454	No	Yes	45	9.9%		
Non-cardiac, nonthoracic								
Brathwaite, 1998(15)	Prospective	404	Not specified	Yes	31	7.7%		
Polanczyk, 1998(16)	Prospective	4 181	No	No	256	4.1%		
Sohn, 2009(17)	Retrospective	7 756	Yes	Yes	30	0.4%		
Gialdini,	Registry	1 606 951	Not specified	Yes	11537	0.7%		

2014(8)

Blackwell, 2015(18)	Registry	4 345	No	Yes	210	4.8%
Kothari, 2016(19)	Registry	15 148	Yes	Yes	554	3.7%
Nassoiy, 2016(20)	Registry	5 065	Not specified	Yes	408	8.1%
Bhave, 2012+ (21)	Registry	363 092	Not specified	Yes	3602	1.0%
POISE Study Group, 2008 [†] (22) (23)	Prospective, interventional	4177	Yes	No	120	2.9%
POISE-2 Study Group, 2014 ^{†(24)}	Prospective, interventional	10 010	Yes	No	203	2.0%
Xia, 2015(25)	Retrospective	1387	Not specified	No	102	7.4%
Leibowitz, 2017(26)	Retrospective	410	Not specified	Yes	15	3.7%

Footnotes: In interventional studies (Amar, Cardinale, Riber, POISE) patients randomized to intervention groups are not presented in this table.

* number includes atrial fibrillation and atrial flutter;

‡ patients with elevated BNP levels that were assigned to the intervention line with

antiarrythimc treatmens (n= 205) were excluded;

+2.2% of patients underwent thoracic surgery;

[†] study included patients who underwent thoracic surgeries;

** study includes 31,192 patients undergoing thoracic surgery among whom POAF incidence was 2.7%

References to Supplementary Table 1:

- 1. Amar D, Roistacher N, Rusch VW et al. Effects of diltiazem prophylaxis on the incidence and clinical outcome of atrial arrhythmias after thoracic surgery. J Thorac Cardiovasc Surg 2000;120:790-8.
- 2. Passman RS, Gingold DS, Amar D et al. Prediction rule for atrial fibrillation after major noncardiac thoracic surgery. Ann Thorac Surg 2005;79:1698-703.
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- 15. Brathwaite D, Weissman C. The new onset of atrial arrhythmias following major noncardiothoracic surgery is associated with increased mortality. Chest 1998;114:462-8.
- 16. Polanczyk CA, Goldman L, Marcantonio ER, Orav EJ, Lee TH. Supraventricular arrhythmia in patients having noncardiac surgery: clinical correlates and effect on length of stay. Ann Intern Med 1998;129:279-85.
- 17. Sohn GH, Shin DH, Byun KM et al. The incidence and predictors of postoperative atrial fibrillation after noncardiothoracic surgery. Korean Circ J 2009;39:100-4.
- 18. Blackwell RH, Ellimoottil C, Bajic P et al. Postoperative Atrial Fibrillation Predicts Long-Term Cardiovascular Events after Radical Cystectomy. J Urol 2015;194:944-9.
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Continent	Country	Center
		Juravinski Hospital and Cancer Centre, Hamilton
		Saint Joseph's Healthcare, Hamilton
		Hamilton General Hospital, Hamilton
North	Canada	McMaster University Medical Centre, Hamilton
North America		Health Sciences Centre Winnipeg, Winnipeg
America		Walter C. MacKenzie Health Sciences Centre, Edmonton
		Victoria Hospital, London
	United States	Cleveland Clinic, Cleveland
	United States	Washington University School of Medicine, St. Louis
	China	Prince of Wales Hospital, Hong Kong
Asia	India	St. John's Medical College Hospital, Bangalore
Asia	mula	Christian Medical College, Ludhiana
	Malaysia	University Malaya Medical Centre, Kuala Lumpur
		Barts And The London, London
	United	University College Hospital, London
	Kingdom	Leeds Teaching Hospitals, Leeds
Europa		Royal Liverpool University Hospital, Liverpool
Europe	Spain	Hospital de Sant Pau, Barcelona
	Span	Hospital Gregorio Maranon, Madrid
	Poland	Jagiellonian University Medical College, Kraków
	France	Pitie-Salpetriere Hospital, Paris
	Brazil	Hospital do Coracao, São Paulo
South		Hospital de Clinicas de Porto Alegre, Porto Alegre
America	Colombia	Hospital Universitario de Santander, Bucaramanga
AIIICIICA	Peru	Foundation CardioInfanil, Bogota
	i ciu	Hospital Nacional Cayetano Heredia, Lima
Africa	South Africa	Inkosi Albert Luthuli Hospital, Durban
Australia	Australia	Westmead Hospital, Sydney

Supplementary Table 2. List of participating centers.

Supplementary Table 3. STROBE Statement Checklist of items that should be included in reports of cohort studies.

	Item No	Recommendation	Page/Table/ Figure
Title and abstract	1	(<i>a</i>) Indicate the study's design with a commonly used term in the title or the abstract	p. 1
		(b) Provide in the abstract an informative and balanced summary of what was done and what was found	p. 2
Introduction			
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	p. 3
Objectives	3	State specific objectives, including any prespecified hypotheses	p. 3
Methods			
Study design	4	Present key elements of study design early in the paper	р. 3-4
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	p. 4
Participants	6	(<i>a</i>) Give the eligibility criteria, and the sources and methods of selection of participants. Describe methods of follow-up	p. 4
		(b) For matched studies, give matching criteria and number of exposed and unexposed	NA
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable	p. 3-5
Data sources/ measurement	8*	For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe comparability of assessment methods if there is more than one group	p. 4-5
Bias	9	Describe any efforts to address potential sources of bias	p. 5-6
Study size	10	Explain how the study size was arrived at	p. 4
Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen and why	p. 5-6
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding	p. 5-6
		(b) Describe any methods used to examine subgroups and interactions	р. 5-6
		(c) Explain how missing data were addressed	p. 5-6
		(d) If applicable, explain how loss to follow-up was addressed	p. 5-6
		(<i>e</i>) Describe any sensitivity analyses	p. 5-6
Results			
Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for eligibility, confirmed eligible, included in the study, completing follow-up, and analysed	p. 6-7
		(b) Give reasons for non-participation at each stage	p. 6-7
		(c) Consider use of a flow diagram	Fig.1
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders	p. 6-7 Tab. 1
		(b) Indicate number of participants with missing data for each variable of interest	p. 6-7 Tab. 1
		(c) Summarise follow-up time (eg, average and total amount)	р. 4 Fig. 1
Outcome data	15*	Report numbers of outcome events or summary measures over time	<u>р.</u> 7
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted	p. 6-8
		estimates and their precision (eg, 95% confidence interval). Make clear which confounders were adjusted for and why they were included	Tab. 1-2 Fig. 3
		(b) Report category boundaries when continuous variables were categorized	NA

		(<i>c</i>) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	NA
Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses	Supplement
Discussion			
Key results	18	Summarise key results with reference to study objectives	p. 8
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias	p. 10
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence	p. 11
Generalisability	21	Discuss the generalisability (external validity) of the study results	p. 8-9
Other information			
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based	p. 1

Supplementary Table 4. Demographic and clinical characteristics stratified for occurrence of new onset clinically important POAF in patients who had NT-proBNP measured before surgery– a univariable analysis.

	Patients with NT- proBNP measurement and without POAF within 30 days after surgery N=9680		Patients with NT-proBNP measurement and with POAF within 30 days after surgery N=109		p-value
	n	%	n	%	
Demographics					
Age	64.2	(10.9)	73.	3 (10.5)	< 0.001
Female	4909	49.3%	50	45.9%	0.363
Frail	366	3.8%	6	5.5%	0.495
Smoking history					
History of smoking	8162	84.3%	93	85.3%	
Current smoker	4698	48.6%	40	36.7%	0.017
Former smoker	3464	35.8%	53	48.6%	
Medical History					
Coronary artery disease	1250	12.9%	27	24.8%	< 0.001
Recent high-risk CAD	76	0.8%	2	1.8%	0.22
Congestive heart failure	219	2.3%	5	4.6%	0.197
Aortic stenosis	87	0.9%	5	4.6%	0.001
Peripheral vascular disease	684	7.1%	13	11.9%	0.076
History of CVE	593	6.1%	9	8.3%	0.47
History of DVT/PE	360	3.7%	1	0.9%	0.19
Hypertension	5013	51.8%	70	64.2%	0.013
Diabetes	1916	19.8%	27	24.8%	0.24
Chronic obstructive pulmonary disease	724	7.5%	22	20.2%	< 0.001
Obstructive sleep apnea	607	6.3%	7	6.4%	1.0
Pre-operative medication	ıs ≤24h befo	ore surgery			
Beta-blockers	1619	16.7%	23	21.1%	0.278
Rate controlling calcium channel blocker	203	2.1%	3	2.8%	0.891
Pre-operative medication	ns >24h to 7	days before	surgery		
Beta-blockers	1808	18.7%	27	24.8%	0.135
Rate controlling calcium channel blocker	233	2.4%	1	0.9%	0.485
Surgery					
Urgent/Emergent	415	4.3%	9	8.3%	0.074
Thoracic surgery	255	2.6%	5	4.6%	< 0.001

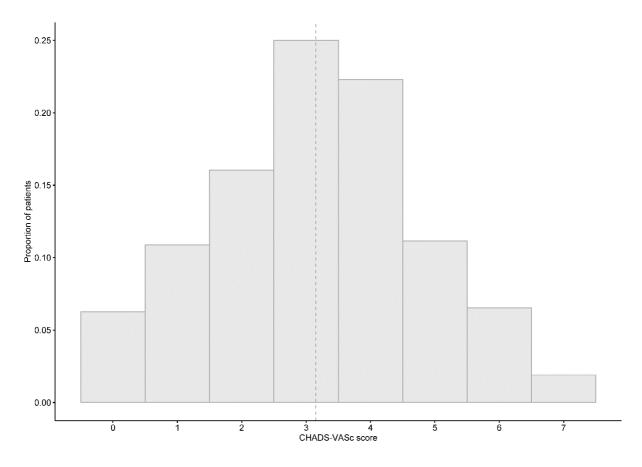
Major nonthoracic	6447	66.6%	94	86.2%	
surgery	0447	00.0%	94	80.2%	
Other surgeries	2978	30.8%	10	9.2%	
Type of anesthesia					
General anesthesia	5649	58.4%	56	51.4%	
Regional anesthesia	2706	28.0%	27	24.8%	0.009
Combined general and	1318	13.6%	26	23.9%	0.007
regional anesthesia	1310	13.070	20	23.970	
Hemoglobin, g/L					
Mean (SD)	132.9	9 (17.7)	128	.4 (18.7)	
Median (IQR)	132 (1	20-143)	128	(113-140)	0.02
eGFR, ml/minute/1.73m2					
Mean (SD)	79.4	(22.0)	69.	7 (23.1)	
Median (IQR)	83.0 (6	7.0-94.9)	76.4 (5	51.8 - 87.1)	< 0.001

Supplementary Table 5. Demographic and clinical characteristics of patients with and without NT-proBNP measurement– a univariable analysis.

	Patients with NT- proBNP measurement N=9789		Patients without NT- proBNP measurement N=27875		p-value	
	n	%	n	%		
Demographics						
Age		3 (11.0)	63.2 (,	< 0.001	
Female	4830	49.3%	13941	50.0%	0.258	
Frail	372	3.8%	1716	6.2%	< 0.001	
Smoking history						
History of smoking	8255	84.3%	24055	86.3%		
Current smoker	4738	48.5%	15474	55.6%	< 0.001	
Former smoker	3517	36.0%	8581	30.8%		
Medical History						
Coronary artery disease	1277	13.0%	3041	10.9%	< 0.001	
Recent high-risk CAD	78	0.8%	235	0.8%	0.66	
Congestive heart failure	224	2.3%	688	2.5%	0.339	
Peripheral vascular disease	697	7.1%	2106	7.6%	0.165	
History of CVE	602	6.1%	1570	5.6%	0.062	
Hypertension	5083	51.9%	13361	47.9%	< 0.001	
Diabetes	1943	19.8%	5787	20.8%	0.056	
COPD	746	7.6%	1999	7.2%	0.147	
Surgery						
Urgent/Emergent	424	4.3%	3430	12.3%	< 0.001	
Thoracic surgery	260	2.7%	828	3.0%		
Major nonthoracic surgery	6541	66.8%	16267	58.4%	< 0.001	
Other surgeries	2988	30.5%	10780	38.7%		
Type of anesthesia						
General anesthesia	5705	58.3%	17443	62.7%		
Regional anesthesia	2733	27.9%	7487	26.9%	< 0.001	
Combined anesthesia	1344	13.7%	2900	10.4%		
Hemoglobin, g/L						
Median (IQR)	134.0 (123.0-145.0)		131.0 (118.0-143.0)		< 0.001	
eGFR, ml/minute/1.73m2						
Median (IQR)	82.9 (66.9-94.9)		85.3 (67.8-97.9)		< 0.001	
Outcomes						
POAF	109	1.1%	260	0.9%	0.133	
Vascular death in 30 days	44	0.4%	219	0.8%	0.001	
Myocardial infarction in 30 days	330	3.4%	861	3.1%	0.180	
Non-fatal cardiac arrest in 30 days	9	0.1%	31	0.1%	0.747	

ONLINE FIGURES

Supplementary Figure 1. Histogram of CHA₂DS₂-VASc scores among patients who developed perioperative atrial fibrillation.



Footnote: Dashed line represents median CHA2DS2-VASc score.