

# Management of Anticoagulation in the Peri-Procedural Period (MAPPP) App:

**Overview, Instructions and Case Studies** 

May 12, 2017



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Alex C. Spyropoulos has a consultant relationship with: Janssen, Boehringer Ingelheim, Bristol-Myers Squibb, Pfizer, and Daiichi-Sankyo.



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# Alex C Spyropoulos, MD, FACP, FCCP, FRCPC

Professor of Medicine, Hofstra Northwell School of Medicine Professor, The Merinoff Center for Patient-Oriented Research -The Feinstein Institute for Medical Research System Director, Anticoagulation and Clinical Thrombosis Services Northwell Health at Lenox Hill Hospital

# Jessica Cohen, MD

Division of Hospital Medicine, North Shore-LIJ Department of Medicine

Assistant Professor of Medicine, Hofstra North Shore-LIJSchool of Medicine



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# **Origin of the MAPPP app**



- Evidence based clinician's guide developed by the multidisciplinary members of the Peri-Procedural Task Force of the NYS Anticoagulation Coalition and IPRO, the CMS designated Quality Improvement Organization for NYS
- Task Force Lead: Dr. Alex Spyropoulos
- Members: Darren Triller, Jason Gilleylen, Peter Kouides, Carol Patrick, Katherine Cabral, MaryAnne Cronin, Patrick Meek, Anne Myrka, Susan Wymer

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# Why is Perioperative Anticoagulant **Management Relevant?**



- Perioperative management of patients on chronic warfarin is common...
  - 400,000-500,000 patients per year in North America alone
  - ~1 in 6 to 10 patients receiving long-term warfarin are assessed for periprocedural management annually
  - Every NYS Medicare beneficiary undergoes approximately 2 procedures annually requiring anticoagulant interruption



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Douketis J et al Chest 2012: 141(2):e326S-e350S IPRO analysis of Medicare Fee for Service Claims 8/2014 -7/2015

## The Perioperative Management of Antithrombotic Therapy\*

### American College of Chest Physicians Evidence-Based Clinical Practice Guidelines (8th Edition)

James D. Douketis, MD, FRCP(C); Peter B. Berger, MD, FACP; Andrew S. Dunn, MD, FACP; Amir K. Jaffer, MD; Alex C. Spyropoulos, MD, FACP, FCCP; Richard C. Becker, MD, FACP, FCCP; and Jack Ansell, MD, FACP, FCCP

# <text><text><text><text><text><text>

### Chest 2008;133;299-339

**REVIEW ARTICLE** 

### Periprocedural management of patients receiving a vitamin K antagonist or a direct oral anticoagulant requiring an elective procedure or surgery

A. C. SPYROPOULOS, \* A. AL-BADRI, † M. W. SHERWOOD ‡ and J. D. DOUKETIS§ \*Department of Medicine, Anticoagulation and Clinical Thrombosis Services, Hofstra North Shore/UJ School of Medicine, North Shore/UJ Health System, Manhasat, NY; †Cedars-Sirai Heart Institute, Los Angeles, CA; ‡Durham VA Medical Center, Duke University Medical Center, Duke Clinical Research Institute, Durham, NC, USA; and §Department of Medicine, McMaster University, Hamilton, Ontario, Carada

# <section-header><image><image>

/Thromb Haemost 2016; DOI: 10.1111/jth.13305.



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### **ARTICLE IN PRESS**

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### EXPERT CONSENSUS DECISION PATHWAY

2017 ACC Expert Consensus Decision Pathway for Periprocedural Management of Anticoagulation in Patients With Nonvalvular Atrial Fibrillation

A Report of the American College of Cardiology Clinical Expert Consensus Document Task Force

Periprocedural	John U. Doherty, MD, FACC, Chair
Management of	
Anticoagulation	Ty J. Gluckman, MD, FACC
Writing	William J. Hucker, MD, PHD
Committee	James L. Januzzi, Jr, MD, FACC

Thomas L. Ortel, MD, PhD Sherry J. Saxonhouse, MD, FACC Sarah A. Spinler, PharmD, AACC



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# **Anticoagulant Overview**



Warfarin

# Direct Oral Anticoagulants (DOACs)

- Pradaxa<sup>®</sup> (dabigatran)
- Xarelto<sup>®</sup> (rivaroxaban)
- Eliquis<sup>®</sup> (apixaban)
- Savaysa<sup>®</sup> (edoxaban)

# Common DOAC characteristics

- More consistent effects at fixed doses
- Lack of routine laboratory testing
- Rapid onset of effects (anticoagulation achieved ~2 hrs)

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Rapid loss of activity (e.g. when doses missed)



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# Significance of the MAPPP App



**Clinical Decision Support that guides:** 

- Whether to interrupt anticoagulation for a procedure by balancing:
  - Risk of bleeding from procedure
  - Risk of thrombosis from underlying indication
- Timing for interruption of anticoagulation
- Peri-procedural "bridging" when appropriate
- Clinical monitoring
- Timing and dosing for resumption of anticoagulants

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# **Overview**



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# Perioperative Management of Anticoagulation







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# Suggested Thromboembolic Risk Stratification when Discontinuing VKAs Health\*

### <u>High</u>

### **Atrial Fibrillation**

- recent (<3 months) stroke/TIA</p>
- CHADS score 5-6
- rheumatic heart disease

### **Mechanical Heart Valves**

- any caged-ball or tilting disc valve in mitral/aortic position
- any mitral valve prosthesis
- Recent (within 6 mos) stroke/TIA

### Venous Thromboembolism (VTE)

- VTE within past 3 months
- severe thrombophilia
- deficiency of protein C, protein S or antithrombin
- antiphospholipid antibodies
- multiple thrombophilias

### **Moderate**

### **Atrial Fibrillation**

CHADS score 3-4

### **Mechanical Heart Valves**

bileaflet AVR <u>with</u> major risk factors

### <u>VTE</u>

- VTE within past 3-12 months
- Nonsevere thrombophilia
- Active cancer
- Recurrent VTE

### Low

### **Atrial Fibrillation**

CHADS score 0-2

### **Mechanical Heart Valves**

bileaflet AVR <u>without</u> major risk factors

### <u>VTE</u>

VTE more than 12 months ago



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Douketis J et al Chest 2008; 133:299 339S

# **Suggested Procedural Bleed Risk**



HIGH BLEEDING RISK PROCEDURES (2 day risk of major bleed ≥ 2%)	LOW BLEEDING RISK PROCEDURES (2 day risk of major bleed <2%)	MINIMAL BLEEDING RISK PROCEDURES
Major surgery with extensive tissue injury	Arthroscopy	Minor dermatologic procedures (excision of basal and squamous cell skin cancers, actinic keratoses, and premalignant or cancerous skin nevi)
Cancer surgery	Cutaneous/lymph node biopsies	Cataract procedures
Major orthopedic surgery	Shoulder/foot/hand surgery	Minor dental procedures (dental extractions, restorations, prosthetics, endodontics), dental cleanings, fillings
Reconstructive plastic surgery	Coronary angiography	Pacemaker or cardioverter- defibrillator device implantation*
Urologic or Gastrointestinal surgery	Gastrointestinal endoscopy +/- biopsy	
Transurethral prostate resection, bladder resection or tumor ablation	Colonoscopy +/- biopsy	
Nephrectomy, kidney biopsy	Abdominal hysterectomy	
Colonic polyp resection	Laparoscopic cholecystectomy	
Bowel resection	Abdominal hernia repair	
Percutaneous endoscopic gastrotomy (PEG) placement, endoscopic retrograde cholangiopancreatography (ERCP)	Hemorrhoidal surgery	
Surgery in highly vascular organs (kidneys, liver, spleen)	Bronchoscopy +/- biopsy	
Cardiac, intracranial, or spinal surgery	Epidural injections with INR <1.2	
Any major operation (procedure duration >45 minutes)		

### Minimal Bleed Risk:

Continue OAC

### <u>\_ow Bleed Risk</u>

Allow residual AC effect pre-op (i.e. 2-3 half lives) Restart within 24 hrs

### <u>High Bleed Risk</u>

No residual AC effect (i.e. 4-5 half lives)

Restart within 48-72 hrs



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Spyropoulos AC et al J of Thromb Haemost 2016;14(5):875-85

# **Consequences of Thromboembolism** and Major Bleeding



- Arterial thromboembolism
- 15% case-fatality for heart valve thrombosis
- 70% rate of death or disability in stroke

# Venous thromboembolism

- 6% rate of death or permanent disability for DVT; 25% rate for PE
- Major bleeding
- 8-9% case-fatality



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IPRO Serving New York State Martinelli J et al. Circulation 1991; 84(3) Longstreth JR et al. Neurology 2001: 56:368 75 Douketis JD et al JAMA 1998; 279: 458-62 Linkins L et al Ann Intern Med 2003; 893-900

# Hypercoagulability Associated with Surgery: Newer Concepts

- Surgery increases risk of arterial thromboembolism [Wahl 1998]
- Perioperative arterial thromboembolic and stroke rates (1.6% and 0.6%) 10-fold higher than modeling suggests (~0.1-0.2% for 8d)

[Dunn A et al Arch Intern Med 2003; White RH, JTH, 2007]





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# Three Key Questions Regarding Perioperative Management of Patients on Chronic OACs?

- Should oral anticoagulant therapy be discontinued?
- When VKA is discontinued, should the patient have perioperative "bridging" therapy with heparin (UFH or LMWH)?
- What is the optimal periprocedural management of patients on DOACs needing interruption?

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# Minimal Bleed Risk Procedures



Minor dermatologic, cutaneous, dental, opthalmologic procedures (cataract surgery), pacemaker/cardioverter-defibrillator device implantation

**Do not interrupt OAC**\* (Grade 2C)

\*May consider interrupting DOAC day of procedure

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Douketis J et al Chest 2008;133:299S- 339S

# BRUISE Control Study for Pacemaker or Defibrillator Surgery (N = 681)<sup>1</sup>

Table 3. Primary and Secondary Outcomes.*						
Outcome	Heparin Bridging (N = 338)	Continued Warfarin (N = 343)	Relative Risk (95% CI)	P Value		
Primary outcome						
Clinically significant hematoma — no. (%)	54 (16.0)	12 (3.5)	0.19 (0.10-0.36)	<0.001		
Components of primary outcome						
Hematoma prolonging hospitalization — no. (%)	16 (4.7)	4 (1.2)	0.24 (0.08-0.72)	0.006		
Hematoma requiring interruption of anticoagulation — no. (%)	48 (14.2)	11 (3.2)	0.20 (0.10-0.39)	<0.001		
Hematoma requiring evacuation — no. (%)	9 (2.7)	2 (0.6)	0.21 (0.05–1.00)	0.03		

**COMPARE Trial for Catheter Ablation in AF**  $(N = 1584)^2$ 

Warfarin discontinuation/Heparin Bridging emerged as a strong predictor of periprocedural TE (OR 13; 95% CI, 3.1–55.6; P<0.001).

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Birnie DH et al NEJM 2013; 368(22):2084 93
 Di Biase L et al Circulation 2014; 129(25):2638 44

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# Do We Need To Bridge?

# **Bridging Therapy**



The goal of bridging therapy with parenteral heparin (either UFH or LMWH), usually in therapeutic doses, is to allow for continued anticoagulation during temporary discontinuation of vitamin K antagonist (VKA) therapy, usually for an elective procedure or surgery

# "This makes intuitive sense"



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# Meta-Analysis and Systematic Review of Bridging vs No-Bridging: Thromboembolic Events

	Bridgir	ıg	No bridg	jing	Odds Ratio	Odds Ratio
Study or Subgroup	<b>Events</b>	Total	Events	Total	<b>M-H, Random, 95% Cl</b>	<b>M-H, Random, 95% Cl</b>
Daniels et al., 2009	4	342	1	213	2.51 [0.28, 22.60]	
Garcia et al., 2008	0	108	7	1185	0.72 [0.04, 12.76]	
Jaffer et al., 2010	1	229	3	263	0.38 [0.04, 3.68]	
Marquie et al., 2006	0	114	2	114	0.20 [0.01, 4.14]	
McBane et al., 2010	10	514	6	261	0.84 [0.30, 2.35]	
Tompkins et al., 2010	1	155	6	513	0.55 [0.07, 4.59]	
Varkarakis et al., 2005	0	25	3	762	4.25 [0.21, 84.56]	
Wysokinski et al., 2008	3	204	4	182	0.66 [0.15, 3.01]	
Total (95% CI)		1691		3493	0.80 [0.42, 1.54]	•
Total events	1 <del>9</del>		32			
<b>Heterogeneity:</b> I <sup>2</sup> = 0%						0.005 0.1 1 10 200 Favours bridging Favours no bridging

No risk reduction for TE with heparin bridging; no difference in ATE or VTE risks.

No difference in TE risk between full and intermediate/prophylactic dose LMWH.

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Siegel D et al Circulation 2012;126:1630 39

# Meta-Analysis and Systematic Review of Bridging vs No-Bridging: Major Bleeding



	Bridgir	ıg	No bridg	jing	Odds Ratio	Odds Ratio
Study or Subgroup	<b>Events</b>	Total	<b>Events</b>	Total	<b>M-H, Random, 95% Cl</b>	<b>M-H, Random, 95% Cl</b>
Daniels et al., 2009	15	342	5	213	1.91 [0.68, 5.33]	
Garcia et al., 2008	4	108	2	1185	22.75 [4.12, 125.68]	
Jaffer et al., 2010	13	229	3	263	5.22 [1.47, 18.54]	
McBane et al., 2010	14	514	2	261	3.63 [0.82, 16.08]	
Wysokinski etal., 2008	6	204	4	182	1.35 [0.37, 4.86]	
<b>Total (95% CI)</b>		1397		2104	3.60 [1.52, 8.50]	•
Total events	52		16			
Heterogeneil, 1 <sup>2</sup> = 52%						0.005 0.1 1 10 200 Favours bridging Favours no bridging

### Bridging associated with an increase in major bleeding. Significant heterogeneity noted across studies.

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Siegel D et al Circulation 2012;126:1630-39

# **Periprocedural Bridging vs No-Bridging Studies**

Study (N)	Year	Population	Comparators	30-da (post-pr	y event ocedure)	
				ATE or VTE OR	MB +/- CRNMB	
Background 30d Event Rates in No Bridging Arms: ATE = ~ 0.5 - 1.0% MB = ~ 1.0 - 1.5%						
$\frac{RELY}{(N = 1,415)}$	2014	AF	Bridging vs No Bridging	(0.38, 19.3)	(2.45, 8.72)	
<u>MVR Study</u> (N = 1,777)	2014	MHV	Rx-dose vs Px- dose Bridging	0.90 (0.37, 2.18)	3.23 (1.58, 6.62)	
<u>Kaiser VTE</u> (N = 1,178)	2015	VTE	Bridging vs No Bridging	0 vs 3	17.2 (3.9- 75.1)	

# **BRIDGE - Trial Design**



Douketis JD, Spyropoulos AC et al NEJM 2015; 373(9):823-33

# **BRIDGE Trial - Primary Outcomes**

Outcome No. (%)	No Bridging (N=918)	Bridging (N=895)	P Value
ATE	4 (0.4)	3 (0.3)	0.01 (non- inf) 0.73 (sup)
Stroke	2 (0.2)	3 (0.3)	
ΤΙΑ	2 (0.2)	0 (0)	
Systemic embolism	0 (0)	0 (0)	
Major bleeding	12 (1.3)	29 (3.2)	0.005 (sup)

The mean  $CHADS_2$  score in patients who sustained a thromboembolic event was 2.6 (range, 1-4) The median time to an arterial thromboembolic event was 19.0 days (IQR, 6.0-23.0 days) The median time to a major bleeding event after a procedure was 7.0 days (IQR, 4.0-18.0 days)

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# **Periprocedural DOAC Outcomes in SPAF Trials**



	Study	DOAC	<b>30-day rate (post-procedure)</b>		
			stroke/systemic	Major bleeding	
<ul> <li>Va</li> <li>pi</li> <li>Ma</li> <li>ai</li> <li>Oi</li> </ul>	ast majority of p rocedures ajority of patien nd restarted wit nly minority und	oatients underwo Its (~80%) held l hin 2 days post erwent bridging	ent minor (non-h DOAC 2 - 3 days -procedure J (except RELY)	high bleed risk)	ure
	(11 - 4092)		1.30)	1.49)	
	ARISTOTLE	apixaban	0.60% (0.32-	0.85% (0.61-	

$$(N = 5439)$$

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1.12)

Healey JS et al. Circulation 2012;126:343 8 Sherwood MW et al Circulation 2014; 129(18):1850 9 Garcia D et al Blood 2014; 124(25):3692 8

\* Includes only 150mg non-bridging groups

1.12)

# **General principles of pre-procedure DOAC** discontinuation



Stratify by procedural bleed risk (type, urgency) and renal function

'Low' bleed risk: 2–3 half-lives i.e. 1 – 2 days pre-op 'High' bleed risk: 4-5 half-lives

i.e. 2 or more days pre-op

For moderate renal insufficiency: add 1-2 days pre-op

**Consider coagulation tests in specific situations** aPTT, PT, TT, dTT (e.g. Hemoclot<sup>®</sup>), ECT

Pay special attention in patients on antiplatelet therapy and those requiring neuraxial anaesthesia

### No heparin bridging!

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Spyropoulos AC et al Blood. 2012;120(15):2954-62 New York State Darvis-Kasem S et al Semin Thromb Hemost. 2012(7):652-60

# General principles of post-procedure **DOAC** resumption



Only after good control of hemostasis

Dependent on bleeding risk and type of operation

Wait at least 24 hours after operation to restart NOAC for minor or "lowbleed" risk procedures

Wait 48–72 hrs after operation to restart NOAC for major or "high-bleed" risk procedures Consider initial prophylactic doses of NOAC

No full-dose heparin bridging!

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In patients who cannot tolerate orals consider prophylactic doses of heparin for VTE prevention

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Spyropoulos AC et al Blood. 2012;120(15):2954-62 Darvis-Kasem S et al Semin Thromb Hemost. 2012(7):652-60 34



# Validated Periprocedural VKA, Bridging, and DOAC Protocols



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# Suggested Periprocedural Strategies of VKA and DOACs Based on Procedural Bleed Risk

	HIGH BLEEDING RISK PROCEDURES	LOW BLEEDING RISK PROCEDURES	MINIMAL BLEEDING RISK PROCEDURES
HIGH THROMBOEMBOLIC RISK	DOAC users: Interrupt DOAC. Bridging with LMWH not suggested for DOACs Warfarin users: Interrupt warfarin with LMWH bridging suggested based on clinician judgment and most current evidence* †	DOAC users: Interrupt DOAC. Bridging with LMWH not suggested for DOACs Warfarin users: Interrupt warfarin with LMWH bridging suggested based on clinician judgment and most current evidence*	Do not interrupt anticoagulants**
INTERMEDIATE THROMBOEMBOLIC RISK	DOAC users: Interrupt DOAC. Bridging with LMWH not suggested for DOACs Warfarin users: Consider interrupting warfarin without LMWH bridging based on clinician judgment and most current evidence* †	DOAC users: Interrupt DOAC. Bridging with LMWH not suggested for DOACs Warfarin users: Consider interrupting warfarin without LMWH bridging based on clinician judgment and most current evidence*	Do not interrupt anticoagulants**
LOW THROMBOEMBOLIC RISK	DOAC users: Interrupt DOAC. Bridging with LMWH not suggested for DOACs Warfarin users: Interrupt warfarin. Bridging with LMWH not necessary †	DOAC users: Interrupt DOAC, Bridging with LMWH not suggested for DOACS Warfarin users: Interrupt warfarin. Bridging with LMWH not necessary	Do not interrupt anticoagulants**



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Spyropoulos AC et al J of Thromb Haemost 2016;14(5):875 85

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#### Validated Periprocedural and Bridging Protocol

Northwell Health®

Table 4 Validated	periprocedural	warfarin and low	molecular weight h	eparin (LMWH)	bridging protocol
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Day	Warfarin dose	Bridging with LMWH	INR monitoring
- 7 to - 10	Maintenance dose	Assess for perioperative bridging anticoagulation; classify patients as undergoing high or low bleeding risk procedures	Check baseline laboratory findings (hemoglobin, platdet count, serum creatinine, INR)
- 6to - 5	Begin to hold warfarin on day = 5 or day = 6	No LMWH	None
- 4	No wafarin	No LMWH	None
- 3	No wafarin	Sart LMWH at a therapeutic or intermediate dose*	None
- 2	No wafarin	LMWH at a therapeutic or intermediate dose*	None
- 1	No wafarin	Last preprocedural dose of LMWH administered no less than 24 h hefore the start of surgery at half the total daily dose	Assess INR before the procedure; proceed with surgery if the INR is < 1.5. If the INR is > 1.5 and < 1.8, consider low-dose or al vitamin K reversal (1-2.5 mg)
0 or + 1	Resume the maintenance dose of warfarin on the evening of or morning after the procedure	None	None
+1	Maintenance dese	Low bleding risk: restart LMWH at the previous dese High bleding risk: no LMWH administration	According eto linician judgement
+ 2 or + 3	Maintenance dese	Low bleding risk: LMWH administration continued High bleding risk: restart LMWH at the previous dose	According to clinician judgement
+4	Maintenance dose	Low bleding risk: INR testing (discontinue LMWH if the INR is > 1.9) High bleding risk: INR testing (discontinue LMWH if the INR is > 1.9)	INR
+7 to + 10	Maintenance dose	-	INR

INR, International Normalized Ratio. Both twice-daily LMWH regimens (i.e. enotaparin 1 mg kg-1 subcutaneous, daltepearin 100 IU kg-1) and once-daily LMWH regimens (i.e. enotaparin 1.5 mg kg-1 subcutaneous, dalteparin 200 IU kg-1 subcutaneous) have been used. Intermediate-dose LMWH has been less studied in this setting.



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Spyropoulos AC et al J of Thromb Haemost 2016;14(5):875 85

				Resumption	of therapy
Drug	Renal function	Low bleeding risk surgery	High bleeding risk surgery*	Low bleeding risk surgery	High bleeding risk surgery
Dabigatran	CrCl > 50 mL min <sup>-1</sup> CrCl 30– 50 mL min <sup>-1</sup>	Last dose: 2 days before procedure Last dose: 3 days before procedure	Last dose: 3 days before procedure Last dose: 4–5 days before procedure	Resume ~ 24 h after procedure	Resume 2–3 days after procedure (48– 72 h postoperatively)†
Rivaroxaban	CrCl > 50 mL min <sup>-1</sup> CrCl 30– 50 mL min <sup>-1</sup> CrCl 15– 29.9 mL min <sup>-1</sup> ‡	Last dose: 2 days before procedure Last dose: 2 days before procedure Last dose: indivualized on the basis of patient and procedural factors for bleeding and thrombosis	Last dose: 3 days before procedure Last dose: 3 days before procedure Last dose: indivualized on the basis of patient and procedural factors for bleeding and thrombosis	Resume ~ 24 h after procedure	Resume 2-3 days after procedure (48- 72 h postoperatively)†
Apixaban	CrCl > 50 mL min <sup>-1</sup> CrCl 30– 50 mL min <sup>-1</sup> CrCl 15– 29.9 mL min <sup>-1</sup>	Last dose: 2 days before procedure Last dose: 2 days before procedure Last dose: indivualized on the basis of patient and procedural factors for bleeding and thrombosis	Last dose: 3 days before procedure Last dose: 3 days before procedure Last dose: indivualized on the basis of patient and procedural factors for bleeding and thrombosis	Resume ~ 24 h after procedure	Resume 2–3 days after procedure (48– 72 h postoperatively)†
Edoxaban	CrCl > 50 mL min <sup>-1</sup>	Last dose: 2 days before procedure	Last dose: 3 days before procedure	Resume ~ 24 h after procedure	Resume 2–3 days after procedure (48– 72 h postoperatively)†

Table 6 Suggested periprocedural direct oral anticoagulant therapy interruptions (adapted from [4])

CrCl, creatinine clearance. \*Includes any procedure/surgery requiring neuraxial anesthesia. †For patients at high risk for thromboembolism and with a high bleeding risk after surgery, consider administering a reduced dose of dabigatran (75 mg twice daily), rivaroxaban (10 mg once daily) or apixaban (2.5 mg twice daily) on the evening after surgery and on the following day (first postoperative day) after surgery. ‡Value for patients receiving rivaroxaban 15 mg once daily.

### **2015 ASRA Guidelines for DOACs**

 TABLE 4. Recommended Intervals Between Discontinuation of the New Anticoagulants and Interventional Pain Procedure

 and Between the Procedure and Resumption of the New Anticoagulants

Drug	Half-life	Recommended of Drug and Interven	Interval Between tional Pain Proce	Discontinuation edure* (5 Half-lives)†‡	Recommended Interval Between Procedure and Resumption of Drug
Dabigatran	12–17 h		4–5 d		24 h
	28 h (renal disease)		6 d (renal disease	)	
Rivaroxaban	9–13 h		3 d		24 h
Apixaban	$15.2 \pm 8.5$ h		3–5 d‡		24 h

\*The procedures include medium- and high-risk interventional pairs procedures. For low-risk procedures, a shared decision making should be followed, a 2 half-life interval may be considered.

<sup>†</sup>Because of the lack of published studies and in view of the added risks involved in patients with spine abnormalities, we took the upper limit of the halflife of each drug in calculating the 5 half-lives.

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<sup>‡</sup>The potency and the wide variability in the pharmacokinetics of these drugs make us recommend a longer interval.



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Narouze S et al Reg Anesth Pain Med 2015;40: 182 212)

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### How to Apply the MAPPP App



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### Applicability of the MAPPP App



- Performs patient anticoagulation assessment 7+ days prior to procedures
- Categorizes procedure-related bleeding risk and underlying thrombosis risk for each patient
- Provides final recommendation for anticoagulant interruption and bridging related to bleeding and thromboembolic risk

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 Each recommendation is coupled to specific guidance for DOAC users, warfarin users and/or antiplatelet users



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To download the app or view the web-based version, please visit: http://mappp.ipro.org/

# MAPPP!

Welcome to IPRO's Management of Anticoagulation in the Peri-Procedural Period app.

Try out the app yourself!







- Once clicking on accept and continue (disclaimer screens), you'll be presented with a screen displaying various antithrombotic options
- Select the antithrombotic agent relevant to your patient

🗢 TATA 👓	11:29 AM	@ 🕴 99% 🚥	orthwel
	Antithrombotics	Help	ealth®
Warfarin	(Coumadin)	>	
Antiplate	let Agent	>	
Dabigatr	an (Pradaxa)	>	
Rivaroxa	ban (Xarelto)	>	
Apixabar	n (Eliquis)	>	
Edoxaba	n (Savaysa)	>	



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- The next screen then prompts you to categorize the specific procedure bleeding risk as High, Low or Minimal
- If the procedure bleeding risk is known simply click on the appropriate choice
- If the procedure bleeding risk is unknown, click on the "Click here for more information on the above choices" which will allow you to view definitions of each level of bleeding risk





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- The "Click here for more information on the above choices" selection reveals the full definition guidance for High, Low and Minimal Bleeding Risk Procedures.
- Procedure bleeding risk can also be selected from this page by clicking on the appropriate choice

Bleeding Risk Northwell Health<sup>®</sup> **High Bleeding Risk Procedures** (2 day risk of major bleed  $\geq$  2%) Major surgery with extensive tissue injury Cancer surgery Major orthopedic surgery Reconstructive plastic surgery Select Urologic or Gastrointestinal surgery Transurethral prostate resection, bladder resection or tumor ablation Nephrectomy, kidney biopsy Colonic polyp resection Bowel resection Percutaneous endoscopic gastrotomy (PEG) placement, endoscopic retrograde cholangiopancreatography (ERCP) Select **Other High Risk Procedures**  Cardiac, intracranial, or spinal surgery Surgery in highly vascular organs (kidneys, liver, spleen) Multiple tooth extractions Any major operation (procedure duration >45) minutes) Select **IPRO** Serving (last updated: 05-25-2016 V1.1.1) 45 New York IPRO Improving Healthcare for the Common Good ®

5:17 PM

●●●○ AT&T 🤶



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 This screenshot depicts the Low Bleeding Risk and Minimal Bleeding Risk Procedure categories

●●●○○ AT&T 🤶 5:41 PM Northwell **Bleeding Risk** Health<sup>®</sup> Low Bleeding Risk Procedures (2 day risk of major bleed <2%) Minor dental procedures (simple dental extractions, restorations, prosthetics, endodontics) Cutaneous/lymph node biopsies Shoulder/foot/hand surgery Coronary angiography Gastrointestinal endoscopy +/- biopsy Colonoscopy +/- biopsy Abdominal hysterectomy Laparoscopic cholecystectomy Abdominal hernia repair Hemorrhoidal surgery Bronchoscopy +/- biopsy Epidural injections with INR <1.2</li> Pacemaker battery change Pacemaker or cardioverter-defibrillator device implantation' Arthroscopy Select **Minimal Bleeding Risk Procedures**  Minor dermatologic procedures (excision of basal and squamous cell skin cancers, actinic keratoses, and premalignant or cancerous skin nevi) Cataract procedures Dental cleanings, fillings Select (last updated: 05-25-2016 V1.1.1) IPRO Improving Healthcare for the Common Good ® 46



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- The next screen then prompts you to categorize the specific thromboembolic risk as High, Moderate/Medium or Low
- If the thromboembolic risk is known simply click on the appropriate choice
- If the thromboembolic risk is unknown, click on the "Click here for more information on the above choices" which will allow you to view definitions of each level of thromboembolic risk

●●○ AT&T ᅙ	5:51 PM	© ∦ 71% <b>■</b> ⊃	Northw
<	Thromboembolic Risk	Help	Health <sup>®</sup>
High Ris	k	>	
Moderat	e Risk	>	
Low Risł	<	>	
Click here for mo	ore information on the above cl	noices.	



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- The "Click here for more information on the above choices" selection reveals the full definition guidance for High, Moderate/Medium and Low Thromboembolic Risk
- Thromboembolic risk can also be selected from this page by clicking on the appropriate choice

High Thromboembolic Risk Procedures >10%/vr. risk of arterial thromboembolism [ATE] or >10%/month risk of venous thromboembolism [VTE] Antiphospholipid antibodies Caged ball or tilting disc valve in mitral/aortic position • Stroke or transient ischemic attack (TIA) within valve or 6 with AF Severe thrombophilia antithrombin Multiple thrombophilias VTE Recurrent VTE Non-severe thrombophilia Active cancer

●●●○○ AT&T 🤶

last 6 months in patients with a mechanical Atrial fibrilliation (AF) with CHADS2 score of 5 Stroke or TIA within past 3 months in patients Rheumatic valvular heart disease VTE within past 3 months Deficiency of protein C, protein S or Any mechanical mitral valve Select

5:53 PM

Thromboembolic Risk

@ 71% 🔳

X

Medium Thromboembolic Risk Procedures 4-10%/yr. risk of ATE or 4-10%/month risk of

- Bileaflet aortic valve replacement (AVR) WITH major risk factors for stroke
- AF with CHADS2 score of 3 or 4
- VTE within past 3-12 months

Select

Low Thromboembolic Risk Procedures <4%/yr. risk of ATE or <4%/month risk of VTE

- Bileaflet AVR WITHOUT major risk factors for stroke
- AF with CHADS2 score of 0–2 (and no prior stroke or TIA)
- VTE more than 12 months ago

Select

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- Once a Bleeding Risk and Thromboembolic Risk is selected for each patient, the MAPPP app will automatically select the appropriate recommendation
- The final "Results" section will provide a Recommendation with References (upper right corner) and option to select another patient (upper left corner)
- At any point in time, you can double check your input data for Antithrombotic agent selection, Bleeding Risk and Thromboembolic Risk by viewing the information bar at bottom of the screen. Backward navigation can occur by clicking this bar or swiping the screen

#### Recommendation

●●●○ AT&T 🤶

New Patient

Interrupt warfarin with LMWH bridging suggested based on clinician judgment and most current evidence

6:04 PM

Results

88% 🕑 🕸

References

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Atrial fibrillation: Bridging NOT recommended based on Level 1 evidence, but evidence in few high risk CHADS2 patients (score 5 and 6); MHV and VTE: Retrospective studies suggest bridging increases bleeding risk without reducing thrombosis.

#### Warfarin Interruption and Bridging Suggestions (show/hide)





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#### Case 1



A 58-year-old female with a bileaflet AVR without major risk factors for stroke is scheduled for a laparoscopic cholecystectomy. She is on warfarin 4mg daily with stable INR within therapeutic range.

Using the MAPPP app, what recommendations would you make regarding the patient's peri-procedural anticoagulation?



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### **Case 1- Antithrombotic Agent Selection**

#### Step 1:

 Since the patient is currently taking warfarin, select warfarin (Coumadin) as the proper Antithrombotic

🔹 T&TA 😳	11:29 AM	@ 🕸 99% 🚥
	Antithrombotics	Help
Warfarin (	Coumadin)	>
Antiplatel	et Agent	>
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Rivaroxat	oan (Xarelto)	>
Apixaban	(Eliquis)	>
Edoxabar	n (Savaysa)	>
	fast updated: 01-15-2016 V1.0)	

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#### Step 2

- You now have access to the Bleeding Risk screen and will be prompted to select a procedure-specific Bleeding Risk
- Click on the "Click here for more information on the above choices"
- Note that the drug selection confirmation appears in the bottom information navigation bar

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••000 AI&I 🗢	5:12 PM	e• ⊼ 80% <b>—</b>
<	Bleeding Risk	Help
High Blee	eding	>
Low Blee	eding	>
Minimal I	Bleeding	>
Click here for mo	ore information on the abov	e choices.
Wastaria		
Warfarin		
Warfarin	(last updated: 05-25-2016 V1.1	1)

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#### **Case 1- Bleeding Risk Evaluation**

Since patient is scheduled for a laparoscopic cholecystectomy, the Low Bleeding Risk category should be selected



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#### Bleeding Risk X

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 Bronchoscopy +/- biopsy Epidural injections with INR <1.2</li> Pacemaker battery change Pacemaker or cardioverter-defibrillator device implantation' Arthroscopy

5:41 PM

Select



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## Case 1- Thromboembolic Risk Evaluation Health

#### Step 3

- You now have access to the Thromboembolic Risk screen and will be prompted to select the Thromboembolic Risk
- Click on the "Click here for more information on the above choices"
- Note that the drug selection and the Bleeding Risk confirmation appears in the bottom information navigation bar

●○○○ AT&T LT	E	2:13 PM	@ 1 %	100% 🗰
<	Thrombo	oembolic R	lisk	Help
High I	Risk			>
Mode	rate Risk			>
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Click here fo	r more informa	tion on the ab	ove choices.	
Warfarin	Low Bleedi	ng		





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#### Case 1- Thromboembolic Risk Evaluation

 Since patient presents with a bileaflet AVR without major risk factors for stroke, the Low Thromboembolic Risk category should be selected

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### **Case 1- Recommendation**

- Based on patient's anticoagulant, procedure bleeding risk and thromboembolic risk, the MAPPP-generated result is shown
- The Warfarin Interruption guidance appears below the recommendation

6:43 PM •••• AT&T 🗢 New Patient Results References Recommendation Interrupt warfarin. Bridging with LMWH not necessary. Warfarin Interruption (show/hide) Bridging with International Low Molecular Warfarin Normalized Day Dose Weight Heparin Ratio (INR) (LMWH) Monitoring Assess for perioperative bridging Check baseline anticoagulation; labs -7 Maintenance classify (hemoglobin, to dose patients as platelet count, -10 undergoing serum high or low creatinine, INR) bleeding risk procedures Begin to -6 hold No LMWH None or warfarin dav -5 and and G Low TE Results Warfarin Low Bleeding

(last updated: 05-25-2016 V1.1.1)

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#### **Case 1- Recommendation**



The Warfarin Interruption guide provides a detailed chart guiding anticoagulation bridging or interruption protocols on days leading up to procedures:

Day	Warfarin Dose	Bridging with Low Molecular Weight Heparin (LMWH)	International Normalized Ratio (INR) Monitoring
-7 to -10	Maintenance dose	Assess for perioperative bridging anticoagulation; classify patients as undergoing high or low bleeding risk procedures	Check baseline labs (hemoglobin, platelet count, serum creatinine, INR)
-6 or -5	Begin to hold warfarin day -5 or day -6	No LMWH	None
-4	No Warfarin	No LMWH	None
-3	No Warfarin	Start LMWH at therapeutic or intermediate dose <sup>†</sup>	None
-2	No Warfarin	LMWH at therapeutic or intermediate dose <sup>†</sup>	None
-1	No Warfarin	Last preprocedural dose of LMWH administered no less than 24h before start of surgery at half the total daily dose	Assess INR before the procedure; proceed with surgery if INR <1.5; If INR > 1.5 and <1.8, consider low-dose oral vitamin K reversal (1-2.5 mg)



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#### **Case 1- Recommendation**



The recommendation will additionally extend guidance to include anticoagulation regimens for days following a patient's procedure:

0 or +1	Resume maintenance dose of warfarin on evening of or morning after procedure	None	None
+ 1	Maintenance dose	Restart LMWH at previous dose	Per clinician judgment
+2 or +3	Maintenance dose	LMWH administration continued	Per clinician judgment
+4	Maintenance dose	INR testing (discontinue LMWH if INR > 1.9)	INR
+7 to +10	Maintenance dose		INR

† Either twice daily LMWH regimens (i.e. enoxaparin 1mg/kg subcutaneous, dalteparin 100 IU/kg subcutaneous) or once-daily LMWH regimens have been used (i.e. enoxaparin 1.5 mg/kg subcutaneous, dalteparin 200 IU/kg subcutaneous). Intermediate-dose LMWH has been less studied in this setting.

Decisions to interrupt, bridge, and resume anticoagulants MUST be clearly communicated among providers and to patient.							
	Warfarin	Low Bleeding	Low TE	Results			

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#### Case 2



A 76-year-old female with a history of HF, Atrial Fibrillation and HTN is scheduled to undergo a total hip replacement. She is currently on warfarin therapy for a recent DVT (2 months ago).

- $CHADS_2 = 3$
- CrCl = 42 ml/min

Using the MAPPP app, what recommendations would you make regarding the patient's peri-procedural anticoagulation?

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### **Case 2- Antithrombotic Agent Selection**

#### Step 1:

 Since the patient is currently taking warfarin, select warfarin (Coumadin) as the proper Antithrombotic

🔹 T&TA 😳	11:29 AM	* ۶ 99%
	Antithrombotics	Help
Warfarin (0	Coumadin)	>
Antiplatele	et Agent	>
Dabigatra	n (Pradaxa)	>
Rivaroxab	an (Xarelto)	>
Apixaban	(Eliquis)	>
Edoxaban	(Savaysa)	>
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### **Case 2- Bleeding Risk Evaluation**



 Since patient is undergoing major orthopedic surgery, the High Bleeding Risk category should be selected



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#### Case 2- Thromboembolic Risk Evaluation

#### Step 3:

 Due to patient's recent DVT (2 months ago), the High Thromboembolic Risk category should be selected





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### **Case 2 - Recommendation**

- Based on patient's anticoagulant, procedure bleeding risk and thromboembolic risk, the MAPPP-generated result is shown
- The Warfarin Interruption and Bridging Suggestions appear below the recommendation

	●●●○ AT&T 🔶	7:03 PM	@∦53%■⊃
	New Patient	Results	References
	Recommenda	ition	
	Interrupt warfarin with LMWH bridging suggested based on clinician judgment and most current evidence		
d r	Atrial fibrillation: Bridging NOT recommended based on Level 1 evidence, but evidence in few high risk CHADS2 patients (score 5 and 6); MHV and VTE: Retrospective studies suggest bridging increases bleeding risk without reducing thrombosis.		ommended dence in few and 6); MHV ggest bridging ucing
Warfarin Interruption and Bridging Suggestions (show/hide)			ing
		Bridging with	International

Warfarin

Dose

Maintenance

Day

-7

Warfarin



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High Bleeding

Low Molecular

Weight Heparin

(LMWH)

Assess for perioperative bridging

anticoagulation;

classifv

High TE

Normalized

Ratio (INR)

Monitoring

Check baseline

labs

(hemoglobin,

Results



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#### **Case 2- Recommendation**



The recommendation will provide a detailed chart guiding anticoagulation bridging or interruption protocols on days leading up to procedures:

Day	Warfarin Dose	Bridging with Low Molecular Weight Heparin (LMWH)	International Normalized Ratio (INR) Monitoring
7 to -10	Maintenance dose	Assess for perioperative bridging anticoagulation; classify patients as undergoing high or low bleeding risk procedures	Check baseline labs (hemoglobin, platelet count, serum creatinine, INR)
6 or -5	Begin to hold warfarin day -5 or day -6	No LMWH	None
-4	No Warfarin	No LMWH	None
-3	No Warfarin	Start LMWH at therapeutic or intermediate dose <sup>†</sup>	None
-2	No Warfarin	LMWH at therapeutic or intermediate dose <sup>†</sup>	None
-1	No Warfarin	Last preprocedural dose of LMWH administered no less than 24h before start of surgery at half the total daily dose	Assess INR before the procedure; proceed with surgery if INR <1.5; If INR > 1.5 and <1.8, consider low-dose oral vitamin K reversal (1-2.5 mg)



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#### **Case 2- Recommendation**



The recommendation will additionally extend guidance to include anticoagulation regimens for days following a patient's procedure:

0 or +1	Resume maintenance dose of warfarin on evening of or morning after procedure	None	None
+ 1	Maintenance dose	No LMWH administration	Per clinician judgment
+2 or +3	Maintenance dose	Restart LMWH at previous dose	Per clinician judgment
+4	Maintenance dose	INR testing (discontinue LMWH if INR > 1.9)	INR
+7 to +10	Maintenance dose		INR



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#### Case 3



A 64-year-old male with a history of Atrial Fibrillation, HTN, and Type 2 Diabetes is scheduled to undergo a coronary angiography in 2 weeks. He is on Eliquis (apixaban) 5mg BID.

- $CHADS_2 = 2$
- CrCl = 84 ml/min

Using the MAPPP app, what recommendations would you make regarding the patient's peri-procedural anticoagulation?

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### **Case 3 - Antithrombotic Agent Selection**

#### Step 1:

 Since the patient is currently taking Eliquis, select Eliquis (apixaban) as the proper Antithrombotic

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		Antithrombotics	Help
	Warfarin (C	Coumadin)	>
	Antiplatele	t Agent	>
	Dabigatrar	n (Pradaxa)	>
	Rivaroxaba	an (Xarelto)	>
-	Apixaban (	(Eliquis)	>
	Edoxaban	(Savaysa)	>

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### **Case 3 - Bleeding Risk Evaluation**



 Since patient is undergoing coronary angiography, the Low Bleeding Risk category should be selected





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#### Case 3 - Thromboembolic Risk Evaluation

#### Step 3:

 Due to patient's CHADS<sub>2</sub> score of 2 and lack of significant past medical history (prior stroke/TIA), the Low Thromboembolic Risk category should be selected



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### **Case 3 - Recommendation**

- Based on patient's anticoagulant, procedure bleeding risk and thromboembolic risk, the MAPPP-generated result is shown
- The Apixaban Interruption Suggestions appear below the recommendation

●●●○○ AT&	T 🗢 7:19 PM	@∦ 52% ■□	
New Par	tient Results	References	
Recom	mendation		
Interrupt DOAC. Bridging with LMWH not suggested for DOACs.			
Apixaban Interruption Suggestions (show/hide)			
CrCl	Last Dose	Resumption of Therapy	
≥50 ml/min	2 days before procedure		
30- 49.9 ml/min	2 days before procedure	Resume within 24	
15- 29.9 ml/min	Individualized based on patient and procedural factors for bleed and thrombosis	procedure	



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Apixaban Low Bleeding

Low TE Results

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# **Questions/Discussion**

Please complete the program evaluation you will be directed to when you close the webinar.



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#### **Contact Information**



Anne Myrka, RPh, MAT Director, Drug Safety (518) 426-3300 ext 191 Anne.Myrka@area-I.hcqis.org

Teresa Lubowski, PharmD Pharmacist (518) 426-3300 ext 125 Teresa.Lubowski@area-I.hcqis.org IPRO CORPORATE HEADQUARTERS 1979 Marcus Avenue Lake Success, NY 11042-1002 IPRO REGIONAL OFFICE 20 Corporate Woods Boulevard Albany, NY 12211-2370 www.ipro.org



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## References



Management of Anticoagulation in the Peri-Procedural Period: A Tool for Clinicians. IPRO; The Medicare Quality Improvement Organization for New York State. Revised 2016.

This material was prepared by the Atlantic Quality Innovation Network/IPRO, the Medicare Quality Innovation Network Quality Improvement Organization for New York State, South Carolina, and the District of Columbia, under contract with the Centers for Medicare & Medicaid Services (CMS), an agency of the U.S. Department of Health and Human Services. The contents do not necessarily reflect CMS policy. 11SOW-AQINNY-TskC.3-16-19



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