

IOSI

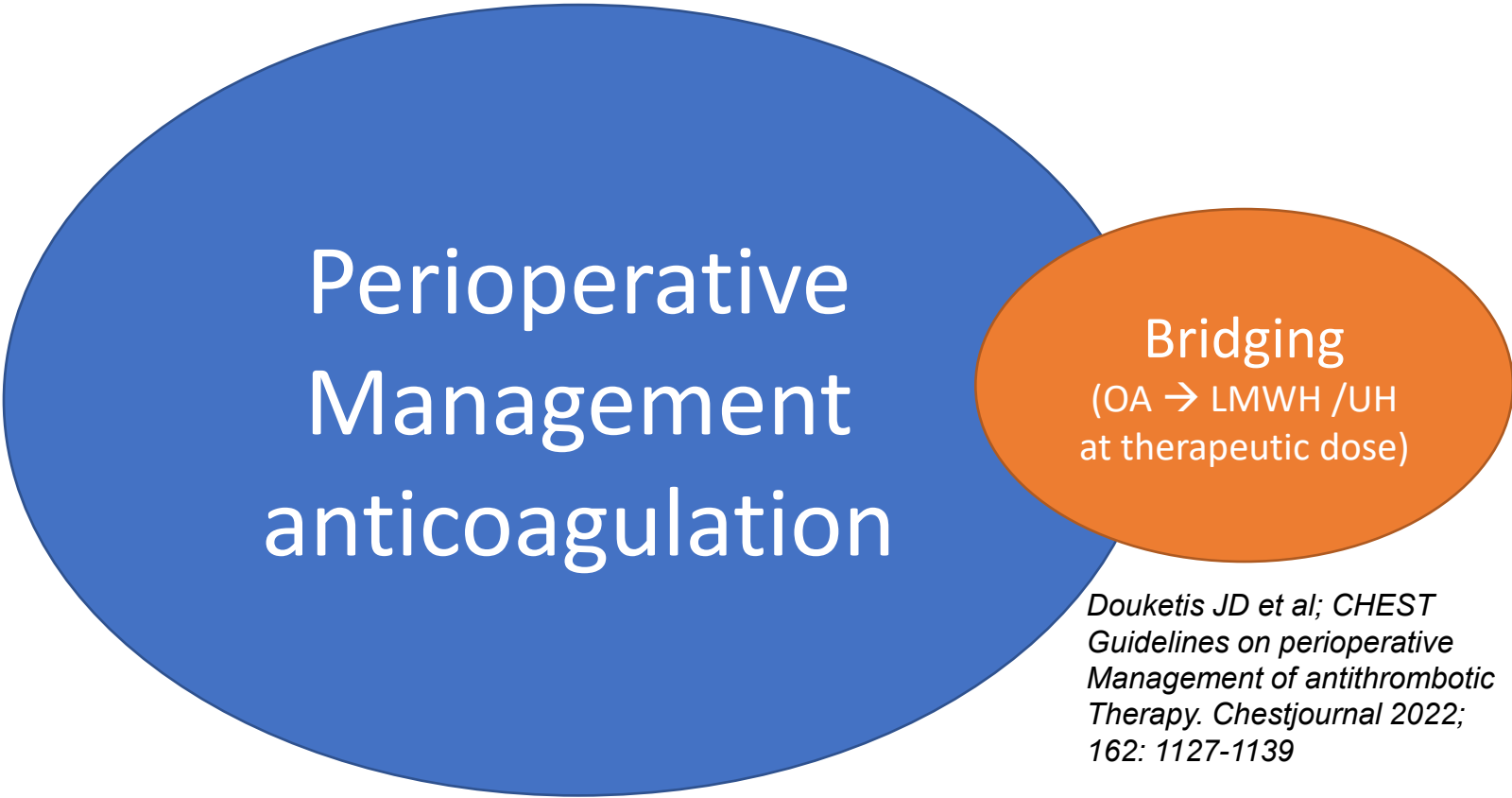
Bridging nell'anticoagulazione

WORKSHOP

23° corso di aggiornamento GMF

15.10.2025

Eugenia Biguzzi



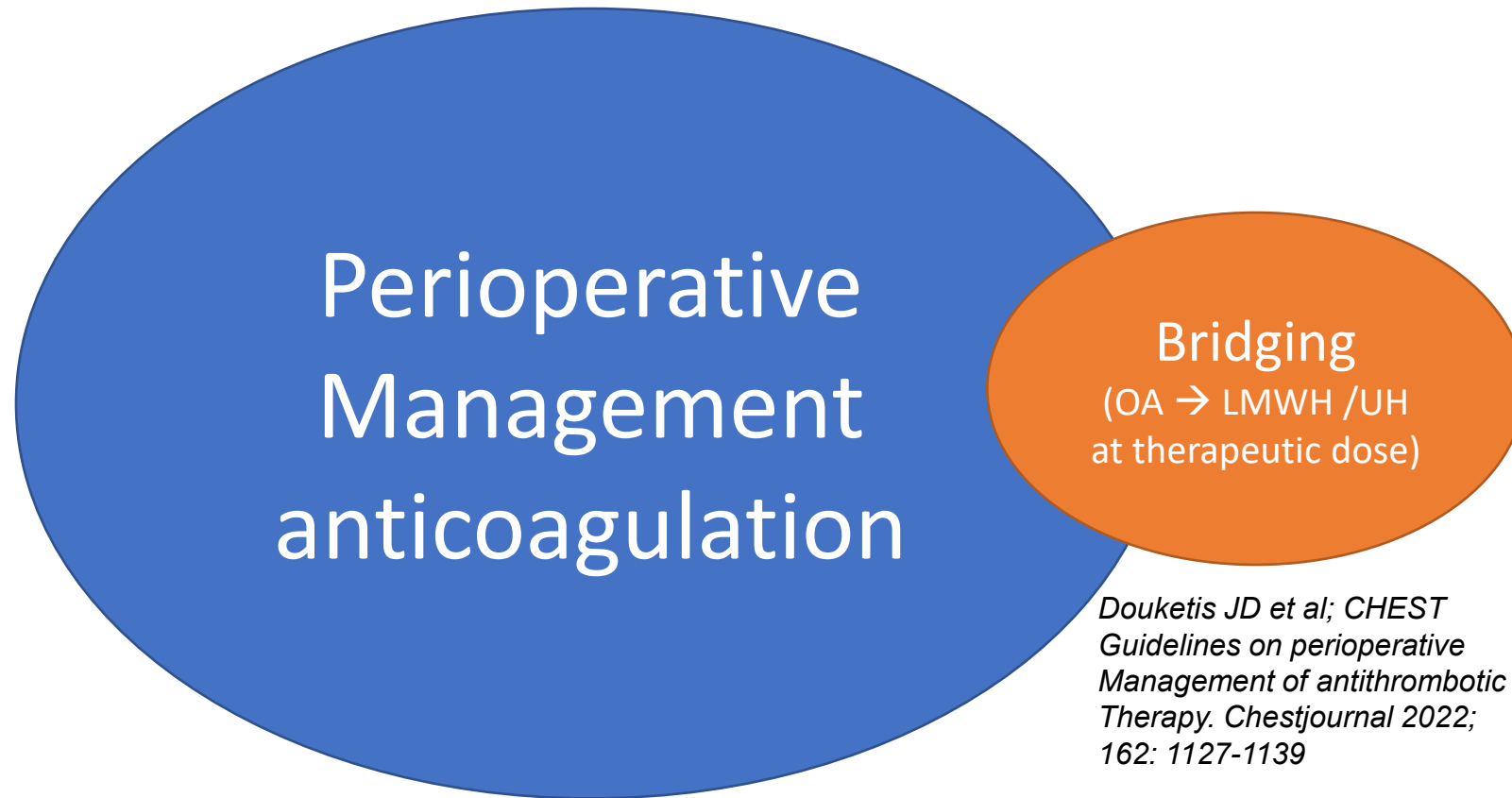
The diagram consists of two overlapping ovals. The larger oval on the left is blue and contains the text 'Perioperative Management anticoagulation'. The smaller oval on the right is orange and contains the text 'Bridging (OA → LMWH /UH at therapeutic dose)'. The two ovals overlap on the right side of the blue oval.

Perioperative Management anticoagulation

Bridging
(OA → LMWH /UH
at therapeutic dose)

*Douketis JD et al; CHEST
Guidelines on perioperative
Management of antithrombotic
Therapy. Chestjournal 2022;
162: 1127-1139*

Who is in charge of the decision?

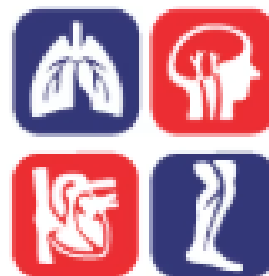


GP → surgeon/specialist → indication to surgery/procedures → date of surgery/procedure

Apps & tools



<https://mappp.ipro.org/>



Thrombosis Canada

Thrombose Canada

https://thrombosiscanada.ca/hcp/practice/clinical_tools?calc=perioperativeAnticoagulantAlgorithm



CoaguSafe - anticoagulanti

Girard Development GmbH

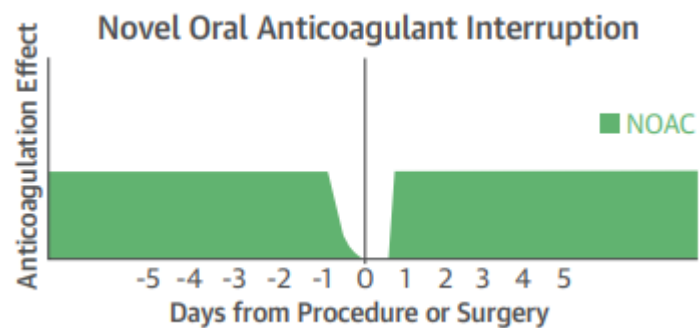
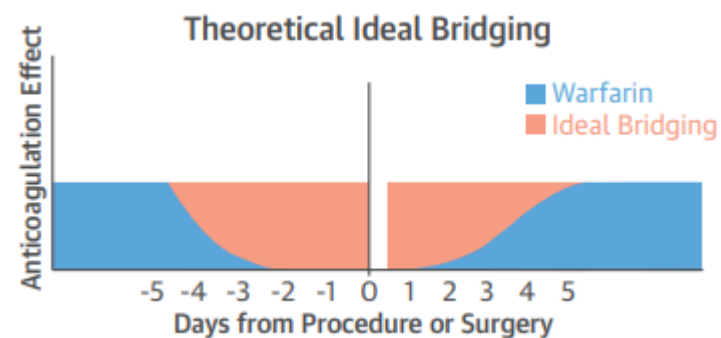
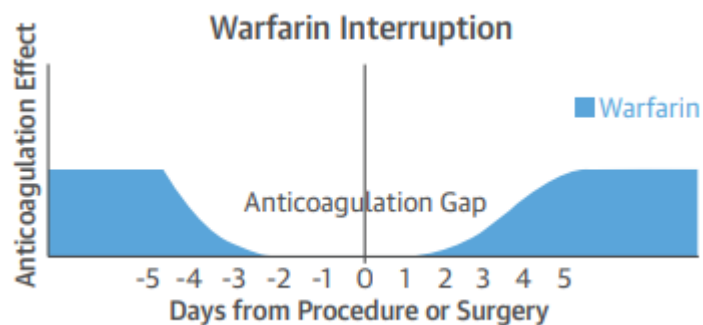
Progettata per iPad

#13 in Medicina

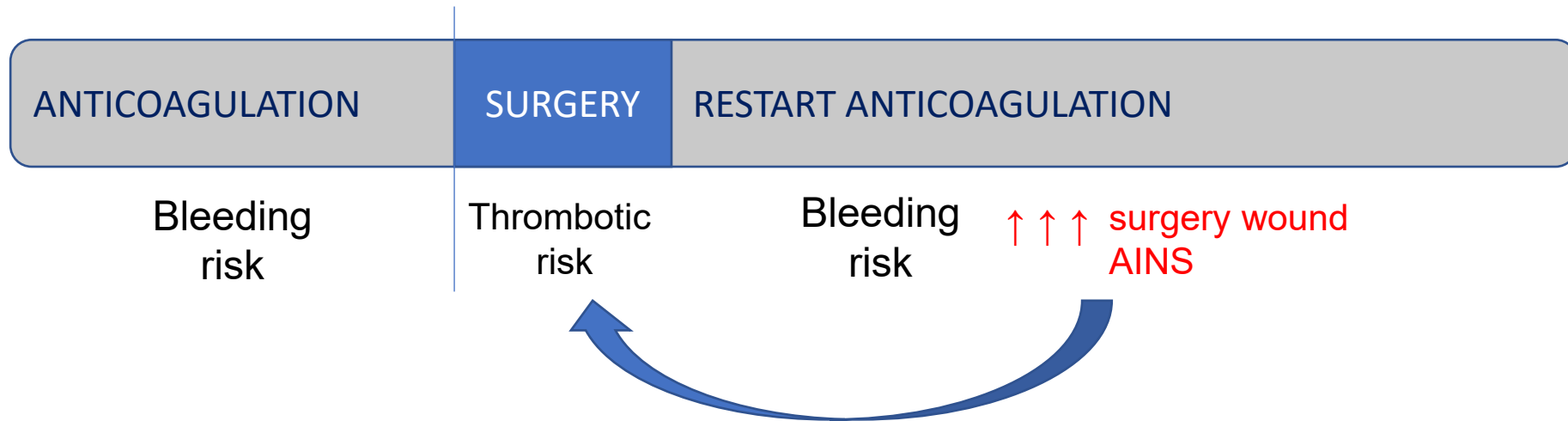
★★★★★ 4,9 • 158 valutazioni

CHF 3.00

The ideal scenario



Scilla and Cariddi: thrombosis and bleeding



Guidelines, websites and app

GUIDELINES

- Douketis JD et al; CHEST **Guidelines on perioperative management of antithrombotic therapy.** Chestjournal 2022;162: 1127-1139
- Douketis JD, Spyropoulos AC. **Perioperative Management of Patients Taking Direct Oral Anticoagulants: A Review.** JAMA. 2024 Sep 10;332(10):825-834
- Thompson A et al. 2024 AHA/ACC/ACS/ASNC/HRS/SCA/SCCT/SCMR/SVM Guideline for **Perioperative Cardiovascular Management for Noncardiac Surgery:** A Report of the American College of Cardiology/American Heart Association Joint Committee on Clinical Practice Guidelines. Circulation. 2024 Nov 5;150(19):e351-e442
- **ORBV_I-ANES-189** Estratto delle raccomandazioni sulla gestione perioperatoria della terapia antiaggregante/anticoagulante per procedure endoscopiche digestive

WEB SITES

<https://mappp.ipro.org/>

https://thrombosiscanada.ca/hcp/practice/clinical_tools?calc=perioperativeAnticoagulantAlgorithm

APP



CoaguSafe - anticoagulanti

Girard Development GmbH

Progettata per iPad

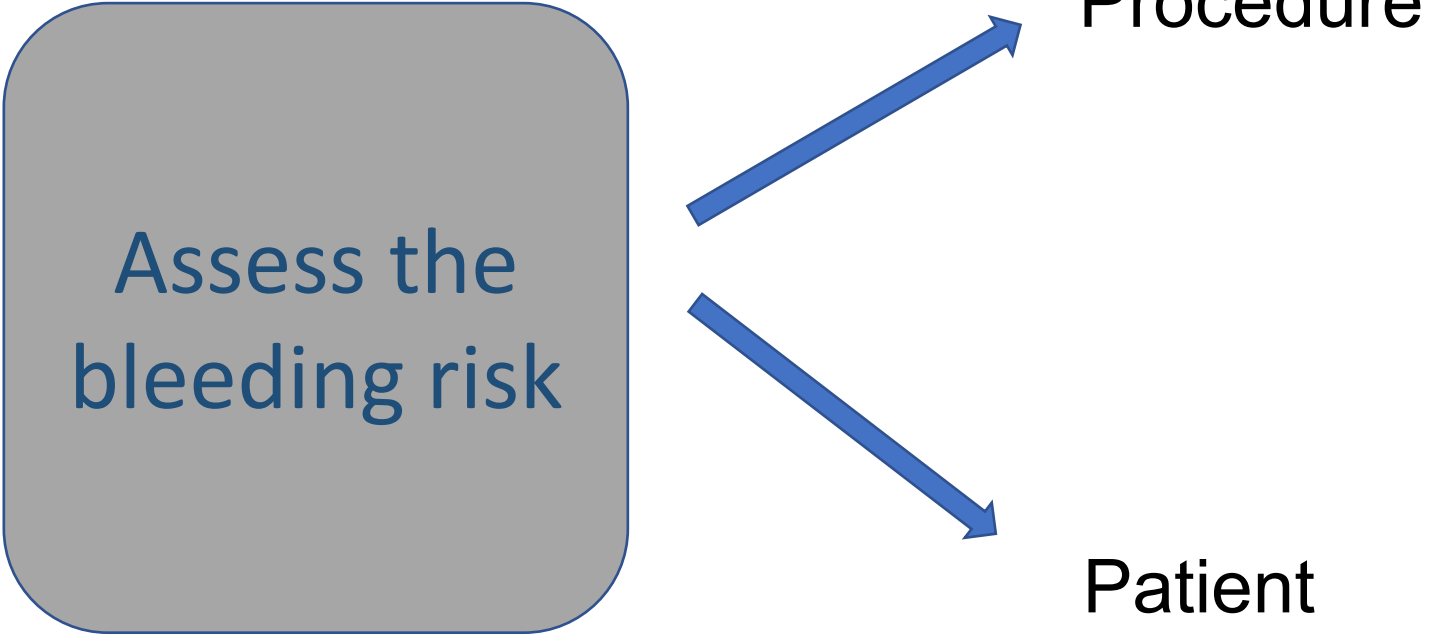
#13 in Medicina

★★★★★ 4,9 • 158 valutazioni

CHF 3.00

**Can the procedure be done
WITHOUT stopping the anticoagulation?**

Assess the
bleeding risk



```
graph LR; A[Assess the bleeding risk] --> B[Procedure]; A --> C[Patient];
```

The diagram consists of a gray rounded rectangle on the left containing the text 'Assess the bleeding risk'. Two blue arrows originate from the right side of this rectangle. The top arrow points diagonally upwards and to the right towards the word 'Procedure'. The bottom arrow points diagonally downwards and to the right towards the word 'Patient'.

Procedure

Patient

Less is more (AVK)

(Bajkin BV et al; Am Ass Oral Maxillof Surg 2009; 67: 990-995)

- 214 patients on oral anticoagulation, simple dental extractions
- randomization
 - 109 patients continue oral anticoagulation INR ≤ 4.0 → bleeding in 7.3%
 - 105 patients bridging with LMWH (mean INR 1.26 ± 0.11) → bleeding in 4.8%



No need for bridging

Less is more (AVK)

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No need for bridging

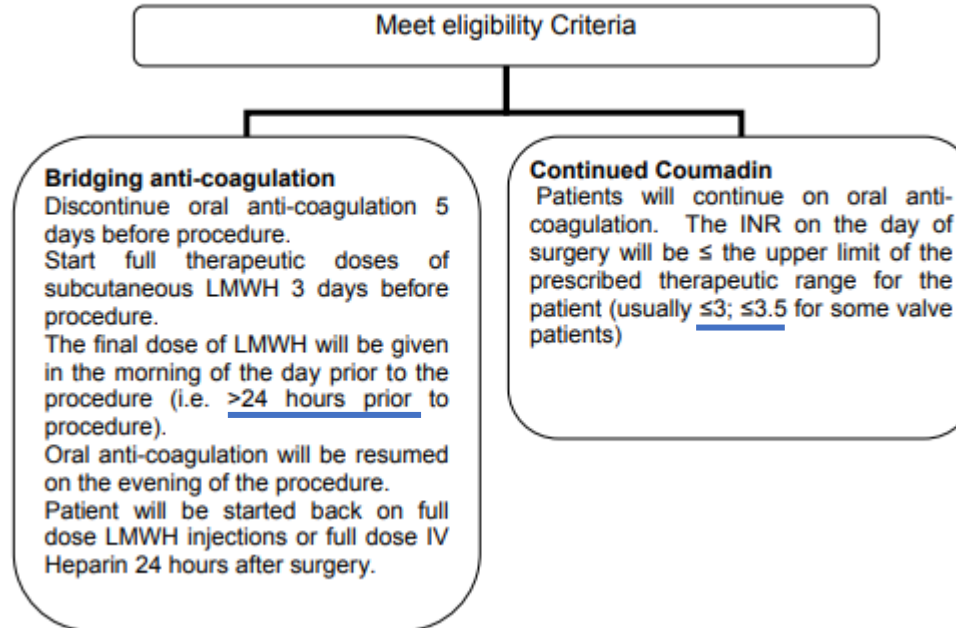
Some suggestions:

- Check INR the day of the procedure
- Accurate surgical hemostasis
- No AINS
- Soft diet + avoid hot beverages/foods for 5-7 days
- Oral rinses with tranexamic acid

Less is more (AVK)

(Birnie et al, **BRUISE** control investigators; NEJM 2013; 368: 2084-2093)

Pacemaker or Defibrillator Surgery without Interruption of Anticoagulation **BRUISE TRIAL (AVK)**



- 681 randomized patients, no blinding (one team implanting the device, one team blind of the randomization assessing possible bleeding complications)
- no restrictions regarding renal function
- Early termination of the trial after the second prespecified interim analysis
- Clinically significant device-pocket hematoma 12/ 343 (3.5%) warfarin vs 54/ 338 (16.0%) bridging
- **Relative risk 0.19 (95% 0.10 - 0.36)**

Less is more (AVK)

(Clark NP et al; Am Heart J; 2018; 195: 108-114)

BRIDGE randomized double-blind placebo controlled trial:

- 1813 patients on warfarin for AF
- Randomization
 - 895 patients stop warfarin, placebo
 - 895 patients stop warfarin, bridging with LMWH (dalteparin 100mg/kg twice daily)
- Major bleeding events n=41
- Risk factors for major bleeding
 - **Bridge therapy** **OR 2.4 (95% CI 1.2-4.8)**
 - History of renal disease OR 2.9 (95% CI 1.4-6.0)
 - Perioperative ASA use OR 3.6 (95% CI 1.1-11.9)
 - Post-procedure INR >3.0 OR 2.1 (95% CI 1.5-5.1)

Less is more (AVK)

(Clark NP et al; Am Heart J; 2018; 195: 108-114)

BRIDGE randomized double-blind placebo controlled trial:

- 1813 patients on warfarin for AF (only 3% with CHADS₂ score of 5-6)
- Randomization
 - 895 patients stop warfarin, placebo
 - 895 patients stop warfarin, bridging with LMWH (dalteparin 100mg/kg twice daily)
- Major bleeding events n=41 (only 11% major surgery/procedure)
- Risk factors for major bleeding
 - **Bridge therapy** **OR 2.4 (95% CI 1.2-4.8)**
 - History of renal disease OR 2.9 (95% CI 1.4-6.0)
 - Perioperative ASA use OR 3.6 (95% CI 1.1-11.9)
 - Post-procedure INR >3.0 OR 2.1 (95% CI 1.5-5.1)

Less is more, also with DOAC

(Douketis JD et al; JAMA Intern Med; 2019; 179: 1469-1478)

PAUSE trial:

3007 patients on DOAC (apixaban/dabigatran/rivaroxaban) for AF, elective procedure

	DOAC	Surgical Procedure-Associated Bleeding Risk	Preoperative DOAC Interruption Schedule					Day of Surgical Procedure (No DOAC)	Postoperative DOAC Resumption Schedule			
			Day -5	Day -4	Day -3	Day -2	Day -1		Day +1	Day +2	Day +3	Day +4
N=1257	Apixaban	High	→							→		
		Low	→									
N=668	Dabigatran etexilate (CrCl ≥50 mL/min)	High	→							→		
		Low	→									
	Dabigatran etexilate (CrCl <50 mL/min) ^a	High	→							→		
		Low	→									
N=1082	Rivaroxaban	High	→							→		
		Low	→									

Less is more, also with DOAC

(Douketis JD et al; JAMA Intern Med; 2019; 179: 1469-1478)

PAUSE trial:

3007 patients on DOAC (apixaban/dabigatran/rivaroxaban) for AF, elective procedure

	Apixaban	Dabigatran	Rivaroxaban
N	1257	668	1082
Major bleeding, n (%)	17 (1.35%)	6 (0.90%)	20 (1.85%)
Procedures with high bleeding risk, n	406	228	373
Major bleeding rate at 30-days, % (95% CI)	2.96 (0-4.7)	0.88 (0-2.6)	2.95 (0.4.76)
Preoperative DOAC levels <50ng/ml, %	90.5%	96.8%	82.6%
Arterial thromboembolism, n (%)	2 (0.16%)	4 (0.60%)	4 (0.37%)

Assess the bleeding risk

PROCEDURES THAT DO NOT REQUIRE TO STOP THE ANTICOAGULATION

Surgical and nonsurgical procedures associated with **minimal** bleeding risk

- | | |
|--|--|
| <ul style="list-style-type: none">• Cardiac device implantation (eg, pacemaker or cardioverter-defibrillator device)• Coronary angiography using radial artery access• Minor dermatologic procedures (eg, excision of basal and squamous cell skin cancers, actinic keratoses, or premalignant or cancerous skin nevi) | <ul style="list-style-type: none">• Phacoemulsification (cataract) procedures• Minor dental procedures (eg, dental extractions, restorations, cleanings, or fillings) |
|--|--|

(Douketis & Spyropoulos, JAMA 2024; 332: 825-834)

- EGD with/without biopsy
- Colonoscopy without biopsy
- Diagnostic ERCP

(ORBV_I-ANES-189, rev 24.02.2025)

Assess the bleeding risk

PROCEDURES THAT DO NOT REQUIRE TO STOP THE ANTICOAGULATION

Surgical and nonsurgical procedures associated with **minimal** bleeding risk

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|--|--|

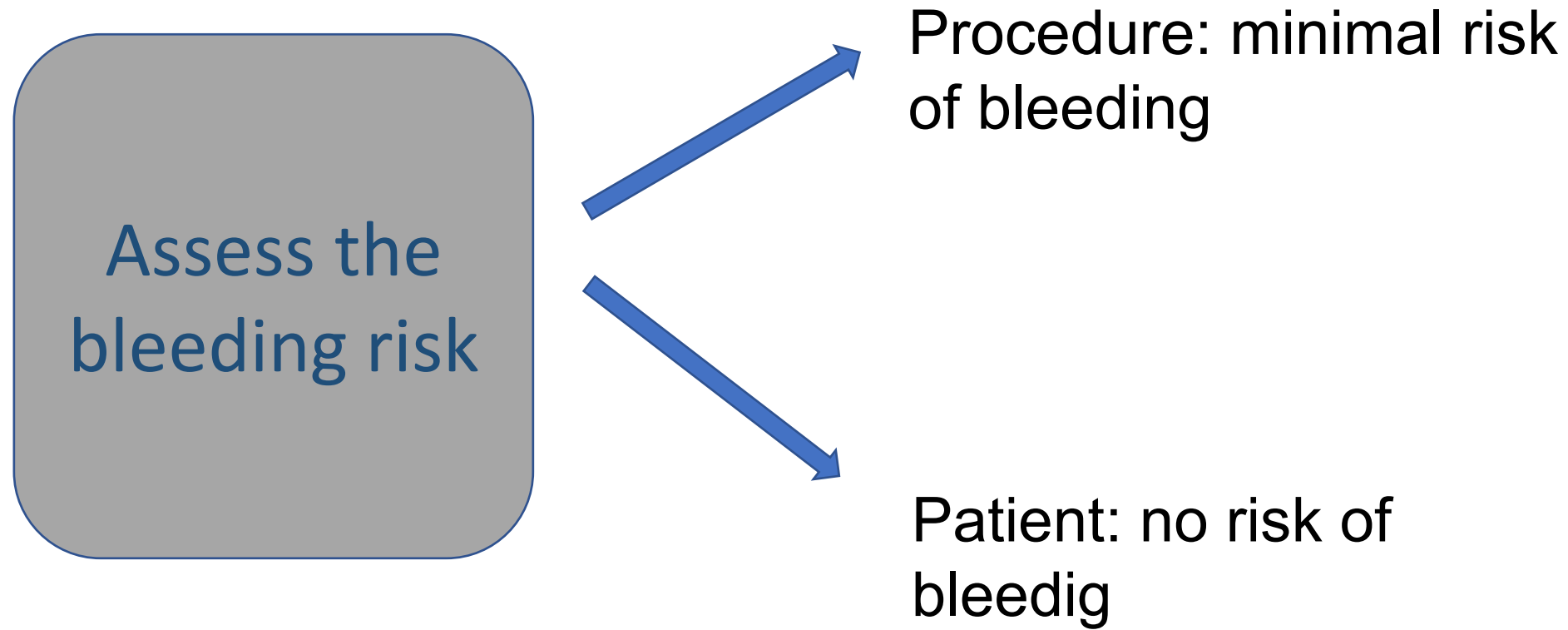
(Douketis & Spyropoulos, JAMA 2024; 332: 825-834)

- EGD with/without biopsy
- Colonoscopy without biopsy
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(ORBV_I-ANES-189, rev 24.02.2025)

PATIENT

- Phytotherapies and over the counter drugs (gingko biloba, ginseng, ginger, omega 3, coenzyme Q10...
→ antiaggregant effect)
- No AINS before and after the procedure
- Uremic patients



Avoid drug peak interval !

- **Rivaroxaban, edoxaban, acenocoumarol, phenprocoumon** → DELAY the morning dose to the evening post-procedure (or avoid dose)
- **Apixaban** → AVOID morning dose

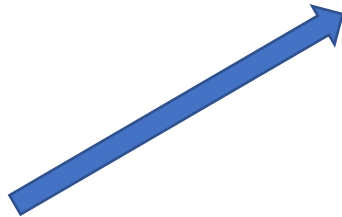
And the other procedures?

Assess the
thrombotic
risk

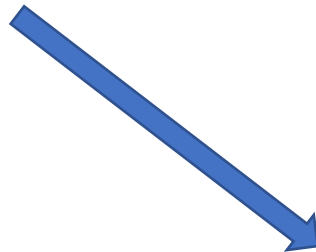


Patient & anticoagulant

Assess the
bleeding risk



Procedure



Patient

Assess the bleeding risk

Procedures associated with low or moderate bleeding risk

- Laparoscopic cholecystectomy
- Abdominal hernia repair
- Abdominal hysterectomy
- Lymph node biopsy
- Foot or hand surgery
- Coronary angiography using femoral artery access
- Gastrointestinal endoscopy with or without biopsy
- Colonoscopy with or without biopsy
- Hemorrhoidal surgery
- Bronchoscopy with or without biopsy

(Douketis & Spyropoulos, JAMA 2024; 332: 825-834)

Procedures associated with high bleeding risk

- Any surgery with spinal or epidural anesthesia
- Major cancer surgery (eg, lung, esophagus, gastric, colon, kidney, or hepatobiliary)
- Major orthopedic surgery (eg, hip or knee replacement)
- Major reconstructive plastic surgery (eg, cancer resection)
- Noncancer major thoracoabdominal surgery (eg, colectomy)
- Transurethral prostate or bladder resection
- Selected biopsies (eg, kidney)
- Selected gastrointestinal procedures (eg, endoscopic retrograde cholangiopancreatography)
- Surgery involving highly vascular organs (eg, kidneys)
- Surgery involving closed space areas (eg, cardiac)
- Deep nerve block procedures

- Colonoscopy with biopsy/polipectomy
- ERCP with sphincterotomy
- Ultrasound endoscopy with biopsy
- GAVE and varices treatment
- PEG

(ORBV_I-ANES-189, rev 24.02.2025)

Assess the bleeding risk

Very detailed list of surgeries /procedures to get an idea

http://jaccjacc.acc.org/Clinical_Document/PMAC_Online_Appendix.pdf



Discuss with the surgeon/specialist !

Assess the thrombotic risk

(Thompson A et al; 2024 AHA/ACC/ASNC/HRS/SCA/SCCT/SCMR/SVM guideline for the perioperative cardiovascular management for noncardiac surgery
Circulation; 2024; 150: e351-e442)

Table 14. Thromboembolic Risk for Common Oral Anticoagulant Indications

Risk Category	Venous Thromboembolism	Atrial Fibrillation	Mechanical Valve	Other Anticoagulation Indications
Low	VTE >12 mo	CHA ₂ DS ₂ -VASc 1-4 (without prior history of stroke)	Bileaflet mechanical AVR without major risk factors for stroke*	
Moderate	VTE ≤3-12 mo Recurrent VTE	CHA ₂ DS ₂ -VASc 5-6	Bileaflet mechanical AVR with major risk factors for stroke*	Nonsevere coagulopathy (heterozygous factor V Leiden or prothrombin gene G20210A mutation) Active cancer
High	Recent VTE (<1 mo or <3 mo)	CHA ₂ DS ₂ -VASc ≥7 (or 5-6 with recent stroke or TIA) AF with rheumatic valvular heart disease	Mechanical mitral valve* Caged ball or tilting-disk valve Mechanical heart valve in any position with recent stroke or TIA (<3 mo)	Recent cardioembolic stroke (<3 mo)† Active cancer associated with high VTE risk LV thrombus (within past 3 mo) Severe thrombophilia‡ Antiphospholipid antibodies

Adapted from Douketis et al.¹⁵ Copyright 2022 Elsevier, with permission from Elsevier.

*Major risk factors for stroke include AF, multiple prior strokes/TIAs (≥3 months), prior perioperative stroke, or prior valve thrombosis.

†Deficiency of protein C, protein S, or antithrombin; homozygous factor V Leiden or prothrombin gene G20210A mutation or double heterozygous for each mutation, multiple thrombophilias.

AF indicated atrial fibrillation; AVR, aortic valve replacement; CHA₂DS₂-VASc, congestive heart failure, hypertension, age ≥75 (doubled), diabetes mellitus, prior stroke or transient ischemic attack or thromboembolism (doubled), vascular disease, age 65 to 74 years, sex category; LV, left ventricular; OAC, oral anticoagulant; TIA, transient ischemic attack; and VTE, venous thromboembolism.

Assess the thrombotic risk

(Thompson A et al; 2024 AHA/ACC(ASNC/HRS/SCA/SCCT/SCMR/SVM guideline for the perioperative cardiovascular management for noncardiac surgery
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Can we delay the surgery/procedure?

Assess the thrombotic risk

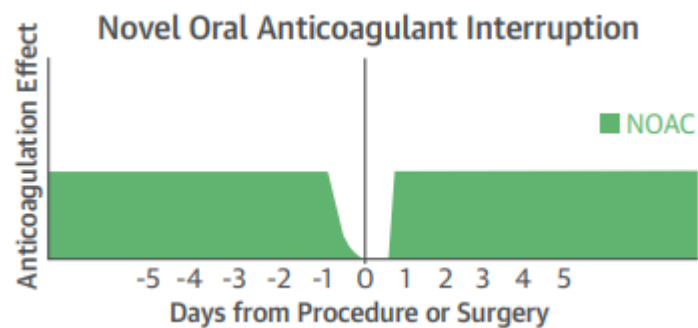
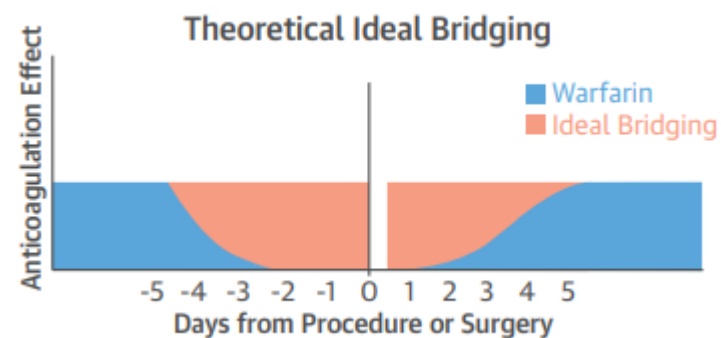
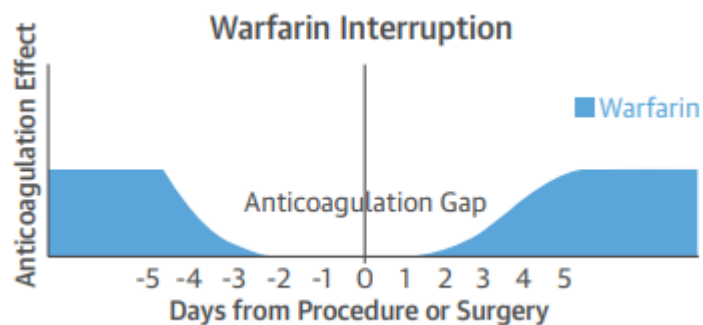
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DOAC

The ideal scenario



Perioperative management: DOAC

	DOAC DOSING REGIMEN	PREOPERATIVE DOAC INTERRUPTION SCHEDULE					SURGERY DAY	POSTOPERATIVE DOAC RESUMPTION SCHEDULE		
		Day -5	Day -4	Day -3	Day -2	Day -1	Day 0	Day +1	Day +2	Day +3
Procedures associated with low or moderate bleeding risk • Laparoscopic cholecystectomy • Abdominal hernia repair • Abdominal hysterectomy • Lymph node biopsy • Foot or hand surgery • Coronary angiography using femoral artery access • Gastrointestinal endoscopy with or without biopsy • Colonoscopy with or without biopsy • Hemorrhoidal surgery • Bronchoscopy with or without biopsy	Rivaroxaban Once a day	●	●	●	●	■	■	●	●	●
	Edoxaban Once a day	●	●	●	●	■	■	●	●	●
	Apixaban Twice a day	● ●	● ●	● ●	● ●	■	■	● ●	● ●	● ●
	Dabigatran Twice a day CrCl ≥50 mL/min	● ●	● ●	● ●	● ●	■	■	● ●	● ●	● ●
	Dabigatran Twice a day CrCl <50 mL/min	● ●	● ●	● ●	■	■	■	● ●	● ●	● ●

(Douketis & Spyropoulos, JAMA 2024; 332: 825-834)

Variable risk definition for EGD and colonoscopy
No evaluation of Creatinine clearance for apixaban

Perioperative management: DOAC

	DOAC DOSING REGIMEN	PREOPERATIVE DOAC INTERRUPTION SCHEDULE					SURGERY DAY Day 0	POSTOPERATIVE DOAC RESUMPTION SCHEDULE		
		Day -5	Day -4	Day -3	Day -2	Day -1		Day +1	Day +2	Day +3
Procedures associated with high bleeding risk - Any surgery with spinal or epidural anesthesia - Major cancer surgery (eg, lung, esophagus, gastric, colon, kidney, or hepatobiliary) - Major orthopedic surgery (eg, hip or knee replacement) - Major reconstructive plastic surgery (eg, cancer resection) - Noncancer major thoracoabdominal surgery (eg, colectomy) - Transurethral prostate or bladder resection - Selected biopsies (eg, kidney) - Selected gastrointestinal procedures (eg, endoscopic retrograde cholangiopancreatography) - Surgery involving highly vascular organs (eg, kidneys) - Surgery involving closed space areas (eg, cardiac) - Deep nerve block procedures	Rivaroxaban Once a day	●	●	●	■	■	■	■	●	●
	Edoxaban Once a day	●	●	●	■	■	■	■	●	●
	Apixaban Twice a day	● ●	● ●	● ●	■	■	■	■	● ●	● ●
	Dabigatran Twice a day CrCl ≥50 mL/min	● ●	● ●	● ●	■	■	■	■	● ●	● ●
	Dabigatran Twice a day CrCl <50 mL/min	●	■	■	■	■	■	■	● ●	● ●

● DOAC dose taken
● DOAC dose taken if hemostasis secured
■ DOAC not taken

(Douketis & Spyropoulos, JAMA 2024; 332: 825-834)

Perioperative management: DOAC

Preoperative DOAC Schedule												
	Procedure Bleeding Risk	Preoperative Interruption						Surgery/ Procedure	Postoperative Resumption			
		Day -6	Day -5	Day -4	Day -3	Day -2	Day -1	Day 0	Day +1	Day +2	Day +3	Day +4
Apixaban, edoxaban, rivaroxaban	High	*	*	*	*	†	†	†	†	†	*	*
	Low/Moderate	*	*	*	*	*	†	†	*	*	*	*
	Minimal	*	*	*	*	*	*	*	*	*	*	*
Apixaban, edoxaban, rivaroxaban with renal impairment (CrCl <30 mL/min)	High	*	*	*	†	†	†	†	†	†	*	*
	Low/Moderate	*	*	*	*	†	†	†	*	*	*	*
	Minimal	*	*	*	*	*	*	*	*	*	*	*
Dabigatran CrCl ≥50 mL/min	High	*	*	*	*	†	†	†	†	†	*	*
	Low/Moderate	*	*	*	*	*	†	†	*	*	*	*
	Minimal	*	*	*	*	*	*	*	*	*	*	*
Dabigatran CrCl <50 mL/min	High	*	*	†	†	†	†	†	†	†	*	*
	Low/Moderate	*	*	*	*	†	†	†	*	*	*	*
	Minimal	*	*	*	*	*	*	*	*	*	*	*

(Thompson A et al; 2024 AHA/ACC/ASNC/HRS/SCA/SCCT/SCMR/SVM guideline for the perioperative cardiovascular management for noncardiac surgery Circulation; 2024: 150: e351-e442)

Perioperative management: DOAC

Schema di sospensione AOD

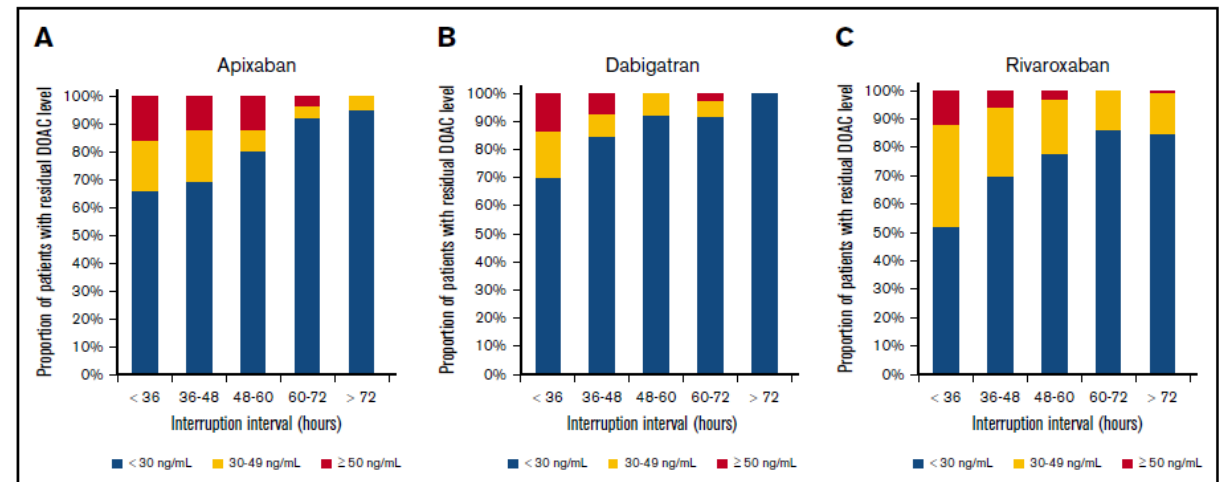
	Dabigatran (Pradaxa®)		Apixaban (Eliquis®) Rdoxaban (Lixiana®) Rivaroxaban (Xarelto®)	
(CrCl calcolata sec. Cockcroft Gault)	Endoscopia NON operativa (Low risk of bleeding)	Endoscopia operativa (High risk of bleeding)	Endoscopia NON operativa (Low risk of bleeding)	Endoscopia operativa (High risk of bleeding)
CrCl ≥ 80 ml/min.	≥24 ore	≥48 ore	≥24 ore	≥48 ore
CrCl 50-79 ml/min.	≥36 ore	≥72 ore	≥24 ore	≥48 ore
CrCl 30-49 ml/min.	≥48 ore	≥96 ore	≥24 ore	≥48 ore
CrCl 15-29 ml/min.	Non indicato l'uso		≥36 ore	≥48 ore
CrCl < 15 ml/min.	Non indicato l'uso			
Non è necessario fare bridging con EBPM o ENF				

(ORBV_I-ANES-189 Estratto delle raccomandazioni sulla gestione perioperatoria della terapia antiaggregante/anticoagulante per procedure endoscopiche digestive)

Predictors of preprocedural DOAC levels before elective surgery

(Shaw JR et al, Blood adv; 2020: 4: 3520-3527)

- **PAUSE study:** 2541 pts on DOAC, undergoing elective surgery
- Low bleeding risk surgery → no DOAC day -1
(dabigatran CrCl <50ml/min → no DOAC day -1-2)
- High bleeding risk surgery → no DOAC day -1-2
(dabigatran CrCl <50ml/min → no DOAC day -1-3)
- **Risk factors for levels >50ng/ml before surgery:**
 - Age ≥75 years
 - Female sex
 - CrCl <50ml/min
- Apixaban and rivaroxaban were more likely than dabigatran to be associated with residual preprocedural levels using the PAUSE protocol.



Do we need to evaluate DOAC levels before surgery?

(Camilleri E et al; DALI study, OC at the ISTH congress 2025)

- **DALI study**
- 2019-2024: 257 pts on DOAC (100 apixaban, 100 rivaroxaban, 57 dabigatran)
- **DOAC stopping according to Creatinine clearance**
- **DOAC levels $\geq 30\text{ng/ml}$ in 7.6% (95% CI 4.9-11.6)**
- 12 major bleeding (all in patients with DOAC levels $< 30\text{ng/ml}$, 4.7% 95%CI 2.7-8.0)
- No association between DOAC levels and surgical blood loss

Do we need to evaluate DOAC levels before surgery?

(Camilleri E et al; DALI study, OC at the ISTH congress 2025)

- **DALI study**
- 2019-2024: 257 pts on DOAC (100 apixaban, 100 rivaroxaban, 57 dabigatran)
- DOAC stopping according to Creatinine clearance
- **DOAC levels $\geq 30\text{ng/ml}$ in 7.6%** (95% CI 4.9-11.6)
- 12 major bleeding (all in patients with DOAC levels $< 30\text{ng/ml}$, 4.7% 95%CI 2.7-8.0)
- No association between DOAC levels and surgical blood loss

Kopp SL et al, ASRAPM guidelines, BMJ 2025: 0:1-29 → neuraxial block

Consider DOAC dosing if shorter discontinuation

Acceptable levels: DOAC $< 30\text{ng/ml}$ or anti-Xa $\leq 0.1 \text{ IU/ml}$

Assess the thrombotic risk

(Thompson A et al; 2024 AHA/ACC(ASNC/HRS/SCA/SCCT/SCMR/SVM guideline for the perioperative cardiovascular management for noncardiac surgery
Circulation; 2024: 150: e351-e442)

Table 14. Thromboembolic Risk for Common Oral Anticoagulant Indications

Risk Category	Venous Thromboembolism	Atrial Fibrillation	Mechanical Valve	Other Anticoagulation Indications
Low	VTE >12 mo	CHA ₂ DS ₂ -VASc 1-4 (without prior history of stroke)	Bileaflet mechanical AVR without major risk factors for stroke*	
Moderate	VTE ≤3-12 mo Recurrent VTE	CHA ₂ DS ₂ -VASc 5-6	Bileaflet mechanical AVR with major risk factors for stroke*	Nonsevere coagulopathy (heterozygous factor V Leiden or prothrombin gene G20210A mutation) Active cancer
High	Recent VTE (<1 mo or <3 mo)	CHA ₂ DS ₂ -VASc ≥7 (or 5-6 with recent stroke or TIA) AF with rheumatic valvular heart disease	Mechanical mitral valve* Caged ball or tilting-disk valve Mechanical heart valve in any position with recent stroke or TIA (<3 mo)	Recent cardioembolic stroke (<3 mo)† Active cancer associated with high VTE risk LV thrombus (within past 3 mo) Severe thrombophilia‡ Antiphospholipid antibodies

AVK

Perioperative management: AVK

VKA Schedule												
	Procedure Bleeding Risk	Preoperative Interruption						Surgery/ Procedure	Postoperative Resumption			
		Day -6	Day -5	Day -4	Day -3	Day -2	Day -1	Day 0	Day +1	Day +2	Day +3	Day +4
Warfarin in low/moderate thrombotic risk	High	*	†	†	†	†	†	†	*	*	*	*
	Low/ Moderate	*	†	†	†	†	†	†	*	*	*	*
	Minimal	*	*	*	*	*	*	*	*	*	*	*
Warfarin in high thrombotic risk	High	*	†	†	‡	‡	‡	†	*	*	*#	*#
	Low/ Moderate	*	†	†	‡	‡	‡	†	*	*#	*#	*#
	Minimal	*	*	*	*	*	*	*	*	*	*	*

(Thompson A et al; 2024 AHA/ACC/ASNC/HRS/SCA/SCCT/SCMR/SVM guideline for the perioperative cardiovascular management for noncardiac surgery. Circulation; 2024; 150: e351-e442)

Bridging with LMWH in very high thrombotic risk

Post-op prophylaxis or full dose LMWH to be discussed

Perioperative management: AVK

VKA Schedule												
	Procedure Bleeding Risk	Preoperative Interruption						Surgery/ Procedure	Postoperative Resumption			
		Day -6	Day -5	Day -4	Day -3	Day -2	Day -1	Day 0	Day +1	Day +2	Day +3	Day +4
Warfarin in low/moderate thrombotic risk	High	*	†	†	†	†	†	†	*	*	*	*
	Low/ Moderate	*	†	†	†	†	†	†	*	*	*	*
	Minimal	*	*	*	*	*	*	*	*	*	*	*
Warfarin in high thrombotic risk	High	*	†	†	‡	‡	‡	†	*	*	*#	*#
	Low/ Moderate	*	†	†	‡	‡	‡	†	*	*#	*#	*#
	Minimal	*	*	*	*	*	*	*	*	*	*	*

(Thompson A et al; 2024 AHA/ACC/ASNC/HRS/SCA/SCCT/SCMR/SVM guideline for the perioperative cardiovascular management of patients undergoing noncardiac surgery. Circulation; 2024; 150: e351-e442)

Bridging with LMWH in high thrombotic risk

Post-op prophylaxis or full dose LMWH to be discussed

Douketis JD et al (2022 Chest guidelines):

- **Bridging** suggested for high thrombotic risk populations with full-dose: subcutaneous LMWH (e.g., enoxaparin, 1 mg/kg bid), with the last dose given the AM of the day prior to the procedure (i.e., D-1) at half the total daily dose.
- **Post-op** low-dose LMWH (e.g., enoxaparin, 40 mg daily or dalteparin 5,000 IU daily) can be used for VTE prophylaxis for first 24-72 hours post-procedure, with full dose LMWH resumed 2-3 days post-procedure.

and Phenprocoumon (Marcoumar) ?

(https://sggssg.ch/fileadmin/user_upload/broschueren/Antikoagulation/Phenprocoucon.pdf)

Discontinuation of vitamin K antagonist anticoagulants (VKA)

Part 1: Phenprocoumon

Discontinuation without bridging (procedure = day 0)

Day -7 to -5	Stop Phenprocoumon; consider INR testing before
Day -2	INR testing; if INR >1.5 administer vitamin K 1-2.5 mg po
Day 0 to +1	Resume phenprocoumon (evening after procedure)

Discontinuation with bridging (procedure = day 0)

Day -7 to -5	Stop Phenprocoumon
Day -4 to -2	INR testing; if INR <2.0 start LMWH (2 sc doses/day);
Day -1	INR testing; if INR >1.5 administer vit. K 1-2.5 mg po; last LMWH dose >24 h before procedure, consider anti-factor Xa testing before procedure in renal insufficiency
Day 0	If adequate hemostasis, LMWH at prophylactic or therapeutic (2 sc doses/day) level according to bleeding risk ≥ 6 h after procedure; resume phenprocoumon, stop LMWH when INR is within target range

The effect of stopping phenprocoumon 5 days preoperatively

(Knol S et al, RPTH 2019; 3:85-88)

- Observational study, 118 patients undergoing an elective procedure
- Phenprocoumon discontinued 5 days before surgery
- **PT-INR ≥ 1.8 in 36%**
- Risk factors: male sex and ASA score 3-4 (5-6 not included)
- **Conclusions:** stop phenprocoumon **7 days before surgery** or customize discontinuation

The effect of stopping phenprocoumon 5 days preoperatively

(van Geest-Daalderop JHH et al, TH 2007; 98:747-755)

Observational study, 111 pts on phenprocoumon undergoing an elective procedure

55 pts, day -2

Vitamin K 2-10mg single dose

Stop phenprocoumon

56 pts, day -2

Vitamin K 2-10mg single dose

The effect of stopping phenprocoumon 5 days preoperatively

(van Geest-Daalderop JHH et al, TH 2007; 98:747-755)

Observational study, 111 pts on phenprocoumon undergoing an elective procedure

55 pts, day -2
Vitamin K 2-10mg single dose
Stop phenprocoumon

PT-INR, mean 1.5 (SD 0.2)

56 pts, day -2
Vitamin K 2-10mg single dose

PT-INR, mean 1.6 (SD 0.3)

Conclusions: administration of vitamin K before an intervention results in an acceptable INR during the intervention, regardless whether phenprocoumon is interrupted or not

Vitamin K dose: 4mg (INR <2.5); 7mg (INR 2.5-3.5); 10mg (INR >3.5)

Perioperative anticoagulant management: guidelines and adherence

(Moesker MJ et al, *BMJ open* 2019, 9: e29879)

- Multicenter retrospective patient record review, 184 elective surgeries in pts on AVK

Preoperative					
No	PAM step	Evaluation criteria		Applicable population	Source
		Compliant	Non-compliant		
1	Timing of patient assessment	► Assessment Performed ≥ 7 days preoperative	► Assessment Performed < 7 days preoperative	Elective	ACCP, 2012
2	VKA interruption interval	► Acenocoumarol interruption = 3 days ► Phenprocoumon interruption = 5 days	► Acenocoumarol interruption $\neq 3$ days ► Phenprocoumon interruption $\neq 5$ days	Elective	CBO, 2008
3	INR testing	► INR is tested on day of surgery	► INR is not tested on day of surgery	Elective	ACCP, 2012
4	Bridging anticoagulation use	Based on thromboembolic risk: ► Low/intermediate risk: no bridging used ► Intermediate/high risk: bridging used	Based on thromboembolic risk: ► Low risk: bridging used ► High risk: no bridging used	Elective	ACCP, 2012
Postoperative					
5	Bridging anticoagulation use	Based on thromboembolic risk: ► Low/intermediate risk: no bridging used ► Intermediate/high risk: bridging used	Based on thromboembolic risk: ► Low risk: bridging used ► High risk: no bridging used	Elective Non-elective	ACCP, 2012
6	Restart time for bridging anticoagulation	► Bridging restart day = 1st day after surgery In case high thromboembolic risk: ► Bridging restart day = day of surgery or 1 st day after surgery	► Bridging restart day $\neq$ 1st day after surgery In case high thromboembolic risk: ► Bridging restart day $\neq$ day of surgery or 1st day after surgery	Elective Non-elective	CBO, 2008
7	Restart time for VKA	► VKA restart day = 1st day after surgery	► VKA restart day $\neq$ 1st day after surgery	Elective Non-elective	CBO, 2008

ACCP, American College of Chest Physicians; CBO, Dutch Quality Institute for Healthcare; INR, international normalised ratio; VKA, vitamin-K antagonist.

Perioperative anticoagulant management: guidelines and adherence

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Preoperative					
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2	VKA interruption interval	► Acenocoumarol interruption = 3 days ► Phenprocoumon interruption = 5 days	► Acenocoumarol interruption ≠ 3 days ► Phenprocoumon interruption ≠ 5 days	Elective	CBO, 2008
3	INR testing	► INR is tested on day of surgery	► INR is not tested on day of surgery	Elective	ACCP, 2012
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Postoperative					
5	Bridging anticoagulation use	Based on thromboembolic risk: ► Low/intermediate risk: no bridging used ► Intermediate/high risk: bridging used	Based on thromboembolic risk: ► Low risk: bridging used ► High risk: no bridging used	Elective Non-elective	ACCP, 2012
6	Restart time for bridging anticoagulation	► Bridging restart day = 1st day after surgery In case high thromboembolic risk: ► Bridging restart day = day of surgery or 1 st day after surgery	► Bridging restart day ≠ 1st day after surgery In case high thromboembolic risk: ► Bridging restart day ≠ day of surgery or 1st day after surgery	Elective Non-elective	CBO, 2008
7	Restart time for VKA	► VKA restart day = 1st day after surgery	► VKA restart day ≠ 1st day after surgery	Elective Non-elective	CBO, 2008

ACCP, American College of Chest Physicians; CBO, Dutch Quality Institute for Healthcare; INR, international normalised ratio; VKA, vitamin-K antagonist.

Valid records	Compliance, %	95% CI
182	81	75-87
119	59	50-68
184	61	54-68
157	80	73-87
181	74	67-81
63	61	48-73
136	42	34-50

Clinical cases

Clinical Case 1, AVK

- Man, 60 years old
- Mechanical mitral valve, since 2020
- On Phenprocoumon with good PT-INR control (time in range >80%)
- Normal weight (75kg), normal creatinine (62 $\mu\text{mol/L}$)

Melanocytic nevus excision (3mm, left forearm)

Clinical Case 1, AVK

- Man, 60 years old
- Mechanical mitral valve, since 2020
- On Phenprocoumon with good PT-INR control (time in range >80%)
- Normal weight (75kg), normal creatinine (62 µmol/L)

Melanocytic nevus excision (3mm, left forearm)

Surgical and nonsurgical procedures associated with **minimal** bleeding risk

- | | |
|--|--|
| <ul style="list-style-type: none">• Cardiac device implantation (eg, pacemaker or cardioverter-defibrillator device)• Coronary angiography using radial artery access• Minor dermatologic procedures (eg, excision of basal and squamous cell skin cancers, actinic keratoses, or premalignant or cancerous skin nevi) | <ul style="list-style-type: none">• Phacoemulsification (cataract) procedures• Minor dental procedures (eg, dental extractions, restorations, cleanings, or fillings) |
|--|--|

(Douketis & Spyropoulos, JAMA 2024; 332: 825-834)

WEB SITES

<https://mappp.ipro.org/>

https://thrombosiscanada.ca/hcp/practice/clinical_tools?calc=perioperativeAnticoagulantAlgorithm

Clinical Case 1, AVK

- Man, 60 years old
- Mechanical mitral valve, since 2020
- On Phenprocoumon with good PT-INR control (time in range >80%)
- Normal weight (75kg), normal creatinine (62 $\mu\text{mol/L}$)

Melanocytic nevus excision (3mm, left forearm)

WEB SITES

<https://mappp.ipro.org/>

Anticoagulant type

Procedure bleeding risk

Thrombotic risk

RESULTS

→ Warfarin (phenprocoumon is not a possible option)

→ minimal (minor dermatologic procedures)

→ moderate (mechanical mitral valve WITHOUT major risk factors for stroke)

→ DO NOT INTERRUPT ANTICOAGULANTS

https://thrombosiscanada.ca/hcp/practice/clinical_tools?calc=perioperativeAnticoagulantAlgorithm

Type of surgery

Procedural bleeding risk

Anticoagulant type

RESULTS

→ elective

→ low (minor non-dental procedure, dermatologic procedures)

→ Warfarin (phenprocoumon is not a possible option)

**→ Continue anticoagulant up to and including day of procedure
and use local hemostatic techniques to control bleeding**

Clinical Case 1, AVK

- Man, 60 years old
- Mechanical mitral valve, since 2020
- On Phenprocoumon with good PT-INR control (time in range >80%)
- Normal weight (75kg), normal creatinine (62 $\mu\text{mol/L}$)

Melanocytic nevus excision (3mm, left forearm)

Tips

- PT-INR (day of procedure, postpone procedure if necessary)
- Konakion corrects INR after at least 4 hours, not useful
- Accurate surgical hemostasis (suture if needed, compressive medication, possible use of local tranexamic acid)

Clinical Case 2, DOAC

- Man, 63 years old
- 11.2023 Pulmonary embolism, no evident transient risk factors, obesity (BMI 33.3)
- Cancer history: prostatic cancer, kidney cancer, urothelial cancer (urological evaluation 01.2024: not active cancer)
- Negative family history of VTE, anticardiolipin IgM positive (276 U/ml, negative rheumatoid factor)
- On Apixaban 5mg BID
- Weight (90kg), increased creatinine (130 $\mu\text{mol/L}$), GFR CKD-EPI 53ml/min, C/G 66ml/min

Colonoscopy (occult neoplasia), 07.2024

Clinical Case 2, DOAC

- Man, 63 years old
- 11.2023 Pulmonary embolism, no evident transient risk factors, obesity (BMI 33.3)
- Cancer history: prostatic cancer, kidney cancer, urothelial cancer (urological evaluation 01.2024: not active cancer)
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- On Apixaban 5mg BID
- Weight (90kg), increased creatinine (130 μ mol/L), GFR CKD-EPI 53ml/min, C/G 66ml/min

Colonscopy (occult neoplasia), 07.2024

Schema di sospensione AOD

	Dabigatran (Pradaxa®)		Apixaban (Eliquis®) Rdoxaban (Lixiana®) Rivaroxaban (Xarelto®)	
(CrCl calcolata sec. Cockcroft Gault)	Endoscopia NON operativa (Low risk of bleeding)	Endoscopia operativa (High risk of bleeding)	Endoscopia NON operativa (Low risk of bleeding)	Endoscopia operativa (High risk of bleeding)
CrCl ≥ 80 ml/min.	≥24 ore	≥48 ore	≥24 ore	≥48 ore
CrCl 50-79 ml/min.	≥36 ore	≥72 ore	≥24 ore	≥48 ore
CrCl 30-49 ml/min.	≥48 ore	≥96 ore	≥24 ore	≥48 ore
CrCl 15-29 ml/min.	Non indicato l'uso		≥36 ore	≥48 ore
CrCl < 15 ml/min.	Non indicato l'uso			
Non è necessario fare bridging con EBPM o ENF				

(ORBV_I-ANES-189 Estratto delle raccomandazioni sulla gestione perioperatoria della terapia antiaggregante/anticoagulante per procedure endoscopiche digestive)

Clinical Case 2, DOAC

- Man, 63 years old
- 11.2023 Pulmonary embolism, no evident transient risk factors, obesity (BMI 33.3)
- Cancer history: prostatic cancer, kidney cancer, urothelial cancer (urological evaluation 01.2024: not active cancer)
- Negative family history of VTE, anticardiolipin IgM positive (276 U/ml, negative rheumatoid factor)
- On Apixaban 5mg BID
- Weight (90kg), increased creatinine (130 μ mol/L), GFR CKD-EPI 53ml/min, C/G 66ml/min

Colonscopy (occult neoplasia), 07.2024

Schema di sospensione AOD

	Dabigatran (Pradaxa®)		Apixaban (Eliquis®) Rdoxaban (Lixiana®) Rivaroxaban (Xarelto®)	
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Non è necessario fare bridging con EBPM o ENF				

(ORBV_I-ANES-189 Estratto delle raccomandazioni sulla gestione perioperatoria della terapiaantiaggregante/anticoagulante per procedure endoscopiche digestive)

No Apixaban day -1 & day -2

When should he restart apixaban? polipectomy no $\rightarrow$ same day
polipectomy yes $\rightarrow$ specialist, wait 48-72h

Clinical Case 3, DOAC

- Man, 53 years old
- 17.10.2024 Emergency department: worsening asthenia, dyspnea on exertion, melena
 - haemorrhagic shock associated with gastric ulcer, HP negative (Hb 57 g/L) → transfusion of 3 units RBC
 - CT scan → bilateral pulmonary embolism
 - Risk factors: obesity (BMI 53.1, occult neoplasia?); thrombophilia (FVL hetero + FII G20210A hetero)
 - **Caval vein filter (unsuccessful retrieval)**
- Comorbidities: metabolic syndrome (arterial hypertension, diabetes, obesity), metabolic liver disease (no information available)
- On Apixaban 5mg BID
- Weight (122 kg), normal creatinine (86 µmol/L), GFR CKD-EPI 92ml/min, C/G not applicable

Eco-endoscopy with biopsy, 09.2025

Clinical Case 3, DOAC

- Man, 53 years old
- 17.10.2024 Emergency department: worsening asthenia, dyspnea on exertion, melena
 - haemorrhagic shock associated with gastric ulcer, HP negative (Hb 57 g/L) → transfusion of 3 units RBC
 - CT scan → bilateral pulmonary embolism
 - Risk factors: obesity (BMI 53.1, occult neoplasia?); thrombophilia (FVL hetero + FII G20210A hetero)
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- Comorbidities: metabolic syndrome (arterial hypertension, diabetes, obesity), metabolic liver disease (no information available)
- On Apixaban 5mg BID
- Weight (122 kg), normal creatinine (86 µmol/L), GFR CKD-EPI 92ml/min, C/G not applicable

Eco-endoscopy with biopsy, 09.2025

Schema di sospensione AOD

	Dabigatran (Pradaxa®)		Apixaban (Eliquis®) Rdoxaban (Lixiana®) Rivaroxaban (Xarelto®)	
(CrCl calcolata sec. Cockcroft Gault)	Endoscopia NON operativa (Low risk of bleeding)	Endoscopia operativa (High risk of bleeding)	Endoscopia NON operativa (Low risk of bleeding)	Endoscopia operativa (High risk of bleeding)
CrCl ≥ 80 ml/min.	≥24 ore	≥48 ore	≥24 ore	≥48 ore
CrCl 50-79 ml/min.	≥36 ore	≥72 ore	≥24 ore	≥48 ore
CrCl 30-49 ml/min.	≥48 ore	≥96 ore	≥24 ore	≥48 ore
CrCl 15-29 ml/min.	Non indicato l'uso		≥36 ore	≥48 ore
CrCl < 15 ml/min.	Non indicato l'uso			
Non è necessario fare bridging con EBPM o ENF				

(ORBV_I-ANES-189 Estratto delle raccomandazioni sulla gestione perioperatoria della terapiaantiaggregante/anticoagulante per procedure endoscopiche digestive)

Clinical Case 4, DOAC

- Woman, 80 years old
- Non valvular Atrial Fibrillation
- On rivaroxaban 20mg/day
- Normal weight (52.5kg), creatinine (85 $\mu\text{mol/L}$, CrCl C/G 39ml/min)
- Comorbidities: arthrosis, arterial hypertension

Tooth avulsion (2 elements)

Surgical and nonsurgical procedures associated with **minimal** bleeding risk

- | | |
|--|--|
| <ul style="list-style-type: none">• Cardiac device implantation (eg, pacemaker or cardioverter-defibrillator device)• Coronary angiography using radial artery access• Minor dermatologic procedures (eg, excision of basal and squamous cell skin cancers, actinic keratoses, or premalignant or cancerous skin nevi) | <ul style="list-style-type: none">• Phacoemulsification (cataract) procedures• Minor dental procedures (eg, dental extractions, restorations, cleanings, or fillings) |
|--|--|

(Douketis & Spyropoulos, JAMA 2024; 332: 825-834)

Clinical Case 4, DOAC

- Woman, 80 years old
- Non valvular Atrial Fibrillation
- On rivaroxaban 20mg/day
- Normal weight (52.5kg), creatinine (85 µmol/L, CrCl C/G 39ml/min)
- Comorbidities: arthrosis, arterial hypertension

Tooth avulsion (2 elements)

Surgical and nonsurgical procedures associated with **minimal** bleeding risk

- | | |
|--|--|
| <ul style="list-style-type: none">• Cardiac device implantation (eg, pacemaker or cardioverter-defibrillator device)• Coronary angiography using radial artery access• Minor dermatologic procedures (eg, excision of basal and squamous cell skin cancers, actinic keratoses, or premalignant or cancerous skin nevi) | <ul style="list-style-type: none">• Phacoemulsification (cataract) procedures• Minor dental procedures (eg, dental extractions, restorations, cleanings, or fillings) |
|--|--|

Tips

(Douketis & Spyropoulos, JAMA 2024; 332: 825-834)

- Warm (not hot!) diet
- Possible use of tranexamic acid rinses
- Think about possible switch to apixaban, reduced dose (2.5mg BID) for age ≥80y + weight <60kg (2.5mg BID)

Clinical Case 5, DOAC

- Woman, 37 years old
- Currently on treatment for breast cancer, BRCA2 mutated, triple negative for hormonal receptors (diagnosis 03.2025)
 - 4 cycles of chemotherapy (neoadjuvant), port-a-cath
 - Oncology program: bilateral mastectomy + ovariectomy (09.2025)
- Comorbidity:
 - Ulcerative rectocolitis (diagnosis 2006, non-responder to immunosuppression, surgery 2012)
 - Obesity (BMI 40.5 kg/m², 101kg x 158cm)
- 09.09.2025 Incidental finding (CT-PET staging): thrombus in right atrium (CVC related)
 - CT scan for dyspnea: segmental pulmonary embolism
 - Lower limbs duplex: negative for deep vein thrombosis (12.09.2025)
 - On anticoagulant (apixaban 5mg BID), creatinine 58 µmol/L, GFR CKD-EPI 116ml/min, C/G 187 ml/min

Bilateral mastectomy + ovariectomy 10.2025

Clinical Case 5, DOAC

- Woman, 37 years old
- Currently on treatment for **breast cancer**, BRCA2 mutated, triple negative for hormonal receptors (diagnosis 03.2025)
 - 4 cycles of chemotherapy (neoadjuvant), **port-a-cath**
 - Oncology program: bilateral mastectomy + ovariectomy (09.2025)
- Comorbidity:
 - **Ulcerative rectocolitis** (diagnosis 2006, non-responder to immunosuppression, surgery 2012)
 - **Obesity** (BMI 40.5 kg/m2, 101kg x 158cm)
- 09.09.2025 Incidental finding (CT-PET staging): thrombus in right atrium (CVC related)
 - CT scan for dispnea: segmental pulmonary embolism
 - Lower limbs duplex: negative for deep vein thrombosis (12.09.2025)
 - On anticoagulant (apixaban 5mg BID), creatinine 58 µmol/L, GFR CKD-EPI 116ml/min, C/G 187 ml/min

Bilateral mastectomy + ovariectomy 10.2025

No Apixaban day -1 & day -2 (GFR CKD-EPI crea (2021) 112 ml/min, C/G 172ml/min

Clinical evaluation pre-op → correct anemia if any deficiency is present

Bridging **pre-op** → enoxaparin 100mg BID (1 dose, day -1 in the morning)
post-op → same day enoxaparin 40mg, daily clinical evaluation to increase dose
 (40mg BID day+1 and +2, 60mg BID day +3 and +4, 80mg BID from day 5 or restart
 apixaban 5mg BID)

Clinical Case 6, AVK

- Woman, 48 years old
- On anticoagulant treatment for ischemic strokes associated with antiphospholipid antibodies (triple positivity) (diagnosis **12.2020**, start **Phenoprocoumon**)
- Uveitis (2020), **renal insufficiency**, livedo reticularis → **Sneddon syndrome (diagnosis 2023)**
- On treatment for **breast cancer**, BRCA2 mutated, triple positive for hormonal receptors (diagnosis 2020)
 - adjuvant therapy (trastuzumab, pertuzumab, paclitaxel)

Left mastectomy (skin reducing) + SLNB + expander 21.04.2021
Right reductive mastectomy

Clinical Case 6, AVK

- Woman, 48 years old
- On anticoagulant treatment for ischemic strokes associated with antiphospholipid antibodies (triple positivity) (diagnosis **12.2020**, start **Phenprocoumon**)
- Uveitis (2020), **renal insufficiency**, livedo reticularis → **Sneddon syndrome (diagnosis 2023)**
- On treatment for **breast cancer**, BRCA2 mutated, triple positive for hormonal receptors (diagnosis 2020)
 - adjuvant therapy (trastuzumab, pertuzumab, paclitaxel)

Left mastectomy (skin reducing) + SLNB + expander 21.04.2021
Right reductive mastectomy

Pre-admission evaluation (20.04.2021)

Body weight 78kg (BMI 27.5), creatinine 141 µmol/L (GFR CKD-EPI 38ml/min, C/G 53ml/min)

Hb 105g/L, PLT 417 G/L, GB 10.5 G/L

Stop Phenprocoumon: last dose 13.04.2021; PT-INR 1.3, **APTT 76 sec**

Bridging (high thrombotic risk for APS) enoxaparin 60mg BID (last dose 20.04.2021, morning)

No AINS in the post-op; ongoing escitalopram 20mg/day

Clinical Case 6, AVK



Recommendation

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

*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 [\(show/hide\)](#)

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 [†]	None
-2	No Warfarin	LMWH at therapeutic or intermediate dose [†]	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)
0 or +1	Provided that adequate hemostasis is achieved, 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

[†] 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.

Warfarin	High Bleeding	High TE	Results
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Clinical Case 6, AVK



Thrombosis Canada

Thrombose Canada

Perioperative Anticoagulant Management Algorithm

Summary

Bleeding Risk: **Moderate** (2-day risk of major bleed 0-2%) **Breast surgery**
Anticoagulant: **Warfarin** **Phenprocoumon not available**
Creatinine Clearance: **53.1 mL/min**
Indication For Antithrombotic: **Mechanical valve** **Antiphospholipid sd not available**
Thromboembolic Risk: **Moderate**

Preoperative Recommendations

No bridging may be acceptable for selected patients but bridging anticoagulation should be considered. Stop warfarin 5-6 days before surgery. Provide therapeutic doses of LMWH (e.g., dalteparin 100 IU/kg SC BID, enoxaparin 1.5 mg/kg SC daily) on preoperative days 3 and 2, and on the day before surgery administer half of the therapeutic dose (e.g., dalteparin 100 IU/kg morning dose only). The last dose of LMWH should be given 24h before surgery. No anticoagulant to be administered the evening before or on the morning of surgery.

Postoperative Recommendations

Postoperative bridging anticoagulation should be considered. Resume warfarin as soon as patient drinking fluids and if further operative intervention is not anticipated. Resume therapeutic dose for LMWH 48-72 hrs after surgery. Consider prophylactic dosing of LMWH (e.g., enoxaparin 40 mg SC daily) until therapeutic dose of LMWH is initiated. Discontinue LMWH once INR in therapeutic range

[Click here for a sample of a perioperative bridging protocol.](#)

Clinical Case 6, AVK



Thrombosis Canada

Thrombose Canada

Sample Bridging Protocol

x

Day	Warfarin	LMWH
-6	✓	×
-5	×	×
-4	×	×
-3	×	✓
-2	×	✓
-1	×	✓*
Surgery	×	×
+1	✓	✓**
+2	✓	✓**
+3	✓	✓***

* Use half daily dose administered 24h prior to procedure

** If high bleeding risk, hold, or use prophylactic dose LMWH

*** Continue LMWH until INR in therapeutic range

Clinical Case 6, AVK

- Woman, 48 years old
- On anticoagulant treatment for ischemic strokes associated with antiphospholipid antibodies (triple positivity) (diagnosis **12.2020**, start **Phenprocoumon**)
- Uveitis (2020), renal insufficiency, livedo reticularis → **Sneddon syndrome (diagnosis 2023)**
- On treatment for **breast cancer**, BRCA2 mutated, triple positive for hormonal receptors (diagnosis 2020)
 - adjuvant therapy (trastuzumab, pertuzumab, paclitaxel)

Left mastectomy (skin reducing) + SLNB + expander 21.04.2021
Right reductive mastectomy

Pre-admission evaluation (20.04.2021)

Body weight 78kg (BMI 27.5), creatinine 141 µmol/L (GFR CKD-EPI 38ml/min, C/G 53ml/min)

Hb 105g/L, PLT 417 G/L, GB 10.5 G/L

Stop Phenprocoumon: last dose 13.04.2021; PT-INR 1.3, **APTT 76 sec**

Bridging enoxaparin 60mg BID (last dose 20.04.2021, morning)

No AINS in the post-op; ongoing escitalopram 20mg/day

Day +0: surgery, no complications, program: enoxaparin 60mg BID from day +1

Day +1: Redon (left side) 110ml hematic, redon (right side) 10ml hematic; Hb 78g/L, local pain, local bruise

→ **hematological consultation** (UH 10.000U/die, APTT not informative, anti-Xa evaluation)

Clinical Case 6, AVK

- Woman, 48 years old
- On anticoagulant treatment for ischemic strokes associated with antiphospholipid antibodies (triple positivity) (diagnosis **12.2020**, start **Phenprocoumon**)
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Day +1: Redon (left side) 110ml hematic, redon (right side) 10ml hematic; Hb 78g/L, local pain, local bruise → hematological consultation (UH 10.000U/die, APTT not informative, anti-Xa evaluation)

From day +3 → Enoxaparin 60mg BID, discharge on day +4; Phenprocoumon from day +5

Day +16 → Hematoma drainage (500ml, serum-hematic, mainly hematic) (PT-INR 1.7, Hb 64g/L), RBC transfusion + INR reversal (Beriplex)

Day +19 → **surgery** hematoma evacuation + expander change → UH 10.000 UI, day 0-2, enoxaparin 60mg BID until day +8, switch to Phenprocoumon by GP

Clinical Case 7, AVK

- Woman, 90 years old
- 1996 mechanical aortic valve (**SJM Master: bileaflet**) → Phenprocoumon
- **07.05.2024** Post-traumatic left femur fracture → surgery, hip prosthesis (planned **10.05.2024**)
- PT-INR 5.4, creatinine 130 µmol/L, GFR epi 34 ml/min; weight 63kg, Hb 125 g/L
- Comorbidities: renal insufficiency

Hip prosthesis

Hematological consultation (08.05.2024, PT-INR 6.8, Hb 112 g/L)

Evaluation: moderate thrombotic risk, high bleeding risk

- Konakion 10mg e.v.
- When PT-INR <2 → bridging with prophylactic UH 10.000/day (stop 3 hours before surgery)
- Post-op: prophylactic UH and switch to Phenprocoumon when possible (surgical evaluation)
- Daily evaluation of PT-INR

Clinical Case 7, AVK

- Woman, 90 years old
- 1996 mechanical aortic valve (**SJM Master: bileaflet**) → Phenprocoumon
- **07.05.2024** Post-traumatic left femur fracture → surgery, hip prothesis (planned **10.05.2024**)
- PT-INR 5.4, creatinine 130 µmol/L, GFR epi 34 ml/min; weight 63kg, Hb 125 g/L
- Comorbidities: renal insufficiency

Hip prothesis

Hematological consultation (08.05.2024, PT-INR 6.8, Hb 112 g/L)

Evaluation: moderate thrombotic risk, high bleeding risk

- Konakion 10mg e.v.
- When PT-INR <2 → bridging with prophylactic UH 10.000/day (stop 3 hours before surgery)
- Post-op: prophylactic UH and switch to Phenprocoumon when possible (surgical evaluation)
- Daily evaluation of PT-INR → **WHY ?**

Clinical Case 7, AVK

- Woman, 90 years old
- 1996 mechanical aortic valve (**SJM Master: bileaflet**) → Phenprocoumon
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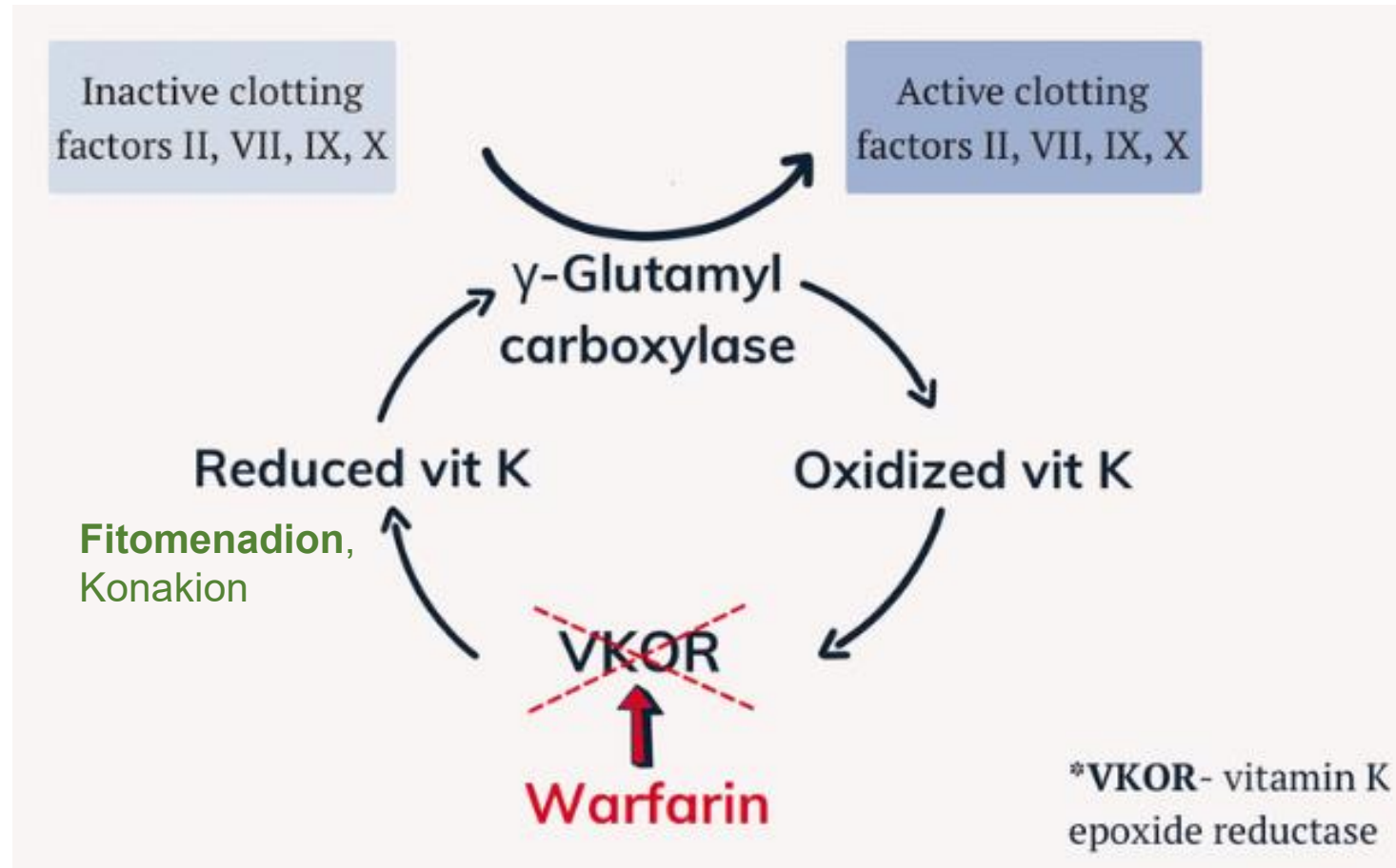
Hip prothesis

Hematological consultation (08.05.2024, PT-INR 6.8, Hb 112 g/L)

Evaluation: moderate thrombotic risk, high bleeding risk

- Konakion 10mg e.v.
- When PT-INR <2 → bridging with prophylactic UH 10.000/day (stop 3 hours before surgery)
- Post-op: prophylactic UH and switch to Phenprocoumon when possible (surgical evaluation)
- Daily evaluation of PT-INR because of possible rebound

Clinical Case 7, AVK



(Mostbauer A et al, J Int Med Res 2022;
50:3000605221103959)

Clinical Case 7, AVK

	07.05	08.05	09.05	10.05
PT-INR	4.3-5.4	6.8	1.3	2.0
APTT, sec	-	-	29	33
Hb, g/L	125		112	
Phenprocoumon		-		
UH			→	
Konakion		10mg		5mg
Consulta ematologia		x		x
Surgery				0
RBC transfusion				

Clinical Case 7, AVK

	07.05	08.05	09.05	10.05	11.05	12.05	13.05	14.05	15.05
PT-INR	4.3-5.4	6.8	1.3	2.0	1.2	1.3	-	-	-
APTT, sec	-	-	29	33	25	29	109	52	54
Hb, g/L	125		112		87	68			
Phenprocoumon		-						1,5	2
UH			→		→	→	→	→	→
Konakion		10mg		5mg					
Consulta ematologia		x		x					
Surgery				0	+1	+2	+3	+4	+5
RBC transfusion						1 unit			

Clinical Case 7, AVK

	07.05	08.05	09.05	10.05	11.05	12.05	13.05	14.05	15.05	16.05	17.05	18.05	19.05	20.05
PT-INR	4.3-5.4	6.8	1.3	2.0	1.2	1.3	-	-	-	5.6	8.2-2.5	2.3		6.8
APTT, sec	-	-	29	33	25	29	109	52	54	71	52	-		
Hb, g/L	125		112		87	68								
Phenprocoumon		-						1,5	2	stop	stop	1	1	
UH			→		→	→	→	→	→	stop	stop			
Konakion		10mg		5mg						2mg	5mg			4mg
Consulta ematologia		x		x						x	x			
Surgery				0	+1	+2	+3	+4	+5	+6	+7	+8	+9	+10
RBC transfusion						1 unit								

Clinical Case 7, AVK

	07.05	08.05	09.05	10.05	11.05	12.05	13.05	14.05	15.05	16.05	17.05	18.05	19.05	20.05	21.05
PT-INR	4.3-5.4	6.8	1.3	2.0	1.2	1.3	-	-	-	5.6	8.2-2.5	2.3		6.8	3.1
APTT, sec	-	-	29	33	25	29	109	52	54	71	52	-			
Hb, g/L	125		112		87	68									
Phenprocoumon		-						1,5	2	stop	stop	1	1		0.25
UH			→		→	→	→	→	→	stop	stop				
Konakion		10mg		5mg						2mg	5mg			4mg	
Consulta ematologia		x		x						x	x				
Surgery				0	+1	+2	+3	+4	+5	+6	+7	+8	+9	+10	+11
RBC transfusion						1 unit									

Phenprocoumon maintenance dose 0.25 tablet/day (not 1 tablet/day) !!!

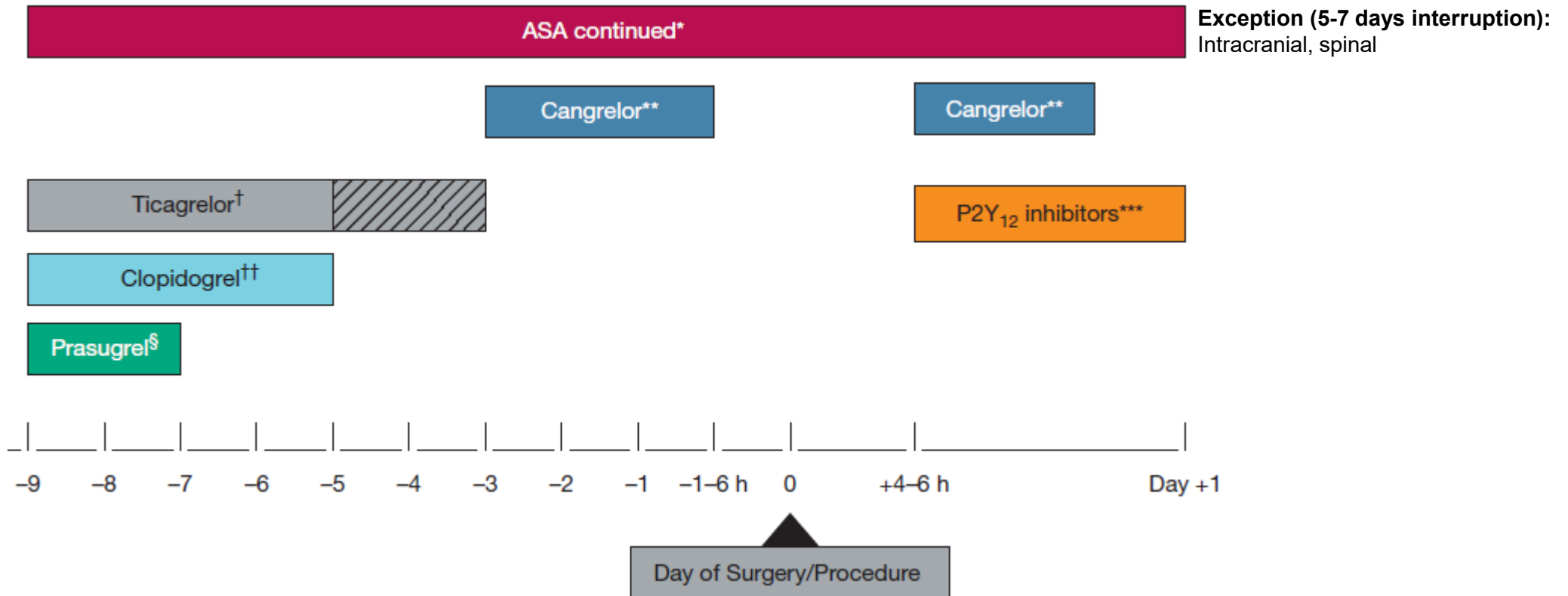
Take home messages

- **Less is more** → reserve bridging therapy ONLY for selected cases
- **Old information are important**
 - type of mechanical valve
 - previous medical history
- **Recent info are equally important**
 - weight, renal function, bleeding events
- **Discuss with surgeons/specialist type of manoeuvre**

Thank you

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Antiplatelet drugs



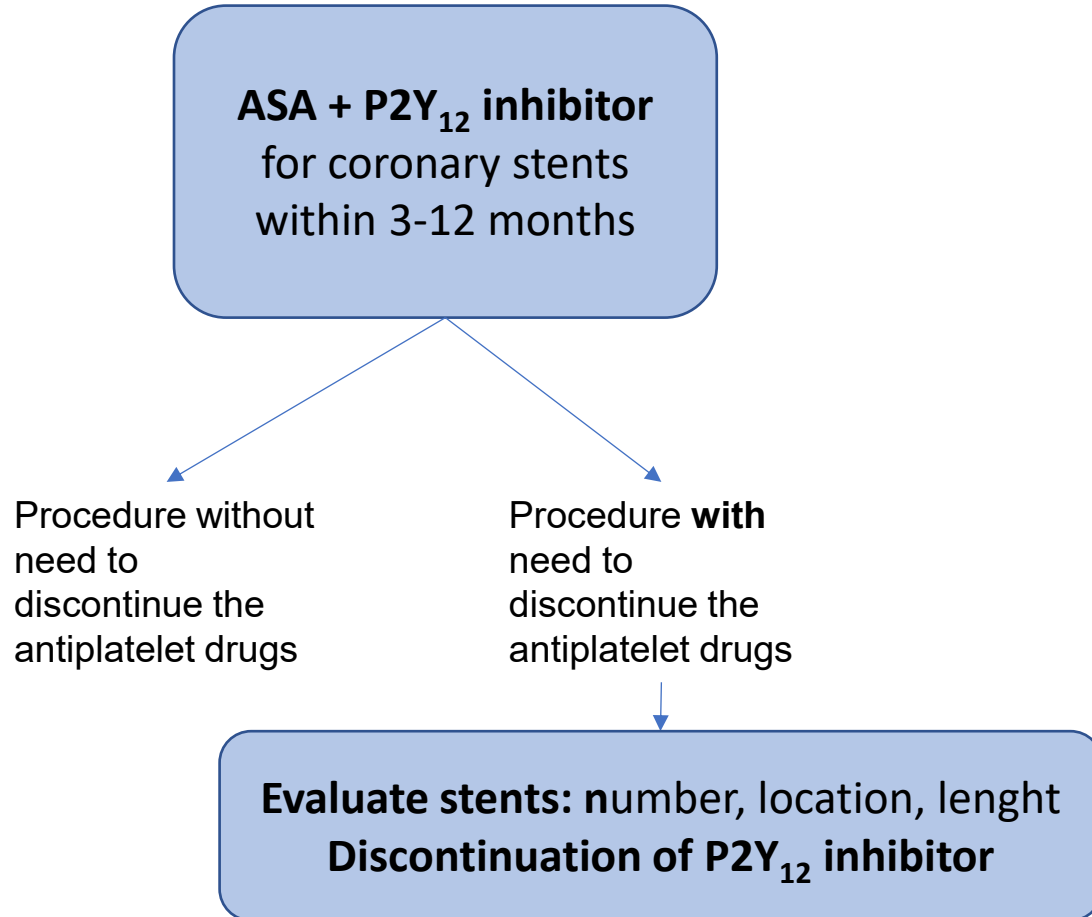
*Based on surgery/procedure bleed risk assessment.

**Routine use not suggested. If used, initiate within 72 hours from P2Y₁₂ inhibitor discontinuation at dose of 0.75 mg/kg/min; resume within 6 hours post-procedure for minimum of 48 hours and maximum of 7 days total. Very low quality data for antiplatelet bridging with glycoprotein IIb/IIIa inhibitors (e.g., eptifibatide, tirofiban).

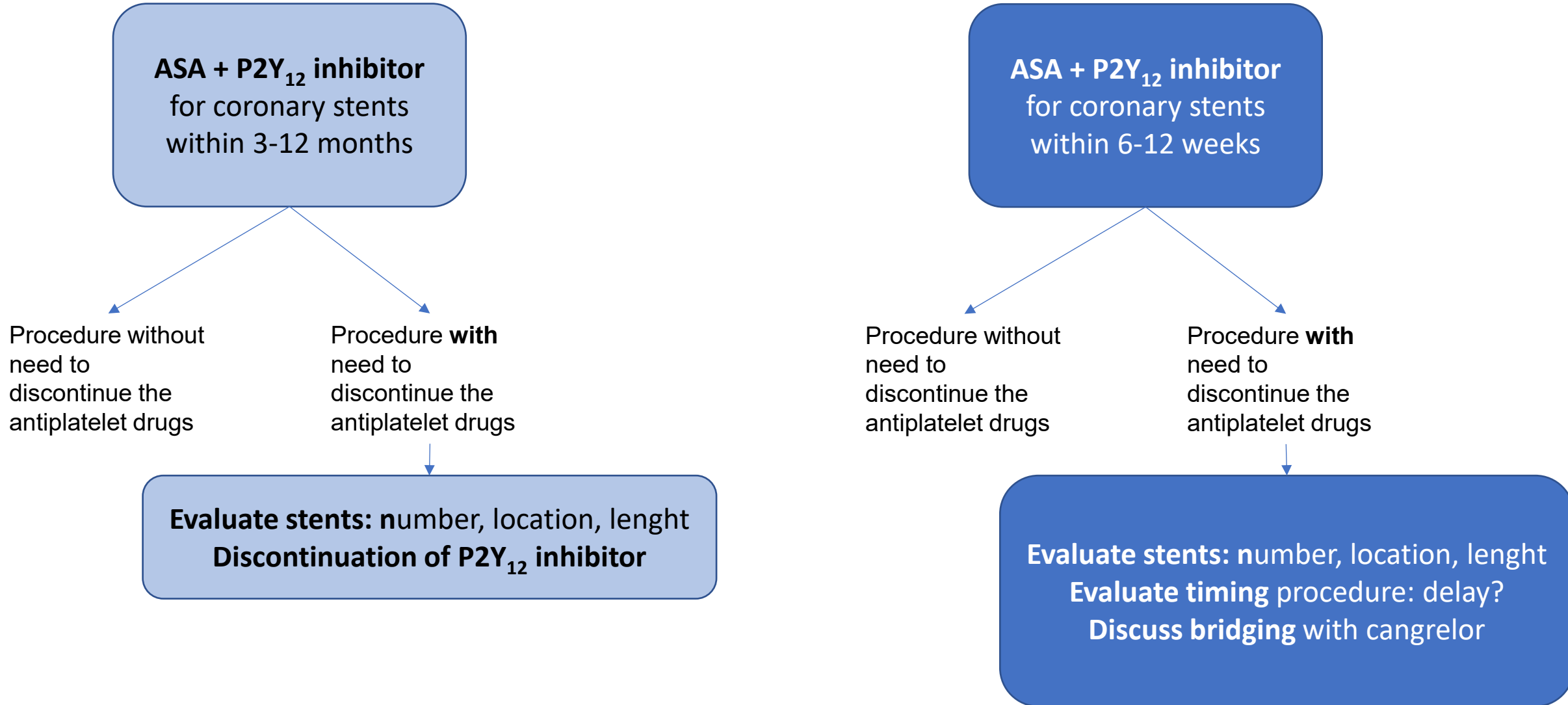
***P2Y₁₂ inhibitors can be resumed within 24 hours post-procedure at a maintenance dose.

†For ticagrelor, 3-5 day interruption
 ††For clopidogrel, 5 day interruption
 §For prasugrel, 7-10 day interruption.

Antiplatelet drugs



Antiplatelet drugs



Clinical Case 8, DAPD

- Man, 75 years old
- **04.04.2024 Myocardial infarction** → stent (2.5x18mm common trunk of the left coronary → ASA + clopidogrel
- 07.04.2024 diagnosis of Hepatocellular Carcinoma (metabolic liver disease) → Selective Internal Radioactive therapy (urgent)

Selective Internal Radioactive therapy (03.06.2024)

Discussion with cardiologist → **indication to bridging with cangrelor** (P2Y12 inhibitor, intravenous administration, short half-life)

Day -5 → stop clopidogrel

Day -3 → start cangrelor 0.75 mcg/kg/min (continuous infusion)

Day 0 → stop cangrelor 1 hour before procedure; restart after 6 hours (clinical evaluation of no bleeding)

Day +1 (24h after procedure) → stop cangrelor, restart clopidogrel