SPECIAL ISSUE ON ENHANCED RECOVERY AFTER SURGERY

Preoperative medication management turnkey order set for nonemergent adult cardiac surgery

Check for updates

Amanda Rea, DNP, CRNP,^a Rawn Salenger, MD,^{a,b} Michael C. Grant, MD, MSE,^c Jennifer Yeh, PharmD,^a Barbara Damas, PharmD,^d Cheryl Crisalfi, MSN, RN,^d Rakesh Arora, MD, PhD,^e Alexander J. Gregory, MD,^f Vicki Morton-Bailey, DNP, CRNP,^g and Daniel T. Engelman, MD,^d on behalf of the ERAS Cardiac Working Group*

ABSTRACT

Objective: The management of preoperative medications is an essential component of perioperative care for the cardiac surgical patient. This turnkey order set is part of a series created by the Enhanced Recovery After Surgery Cardiac Society, first presented at the Annual Meeting of The American Association for Thoracic Surgery in 2023. Numerous guidelines and expert consensus documents have been published to provide guidance in preoperative medication management. Our objective is to integrate these documents into an evidence-based order set that will facilitate standardized implementation of best practices for preoperative medication management for nonemergent adult cardiac surgery.

Methods: Subject matter experts were consulted to translate existing guidelines and peer reviewed literature into a sample turnkey order set for the preoperative management of patients' medications. Orders derived from consistent Class I, IIA, or equivalent recommendations across referenced guidelines and consensus manuscripts appear in the order set in **bold** type. Selected orders that were inconsistently Class I or IIA, Class IIB, or supported by published evidence, were also included in *italicized* type.

Results: Holding antiplatelet and anticoagulant medications before nonemergent cardiac surgical procedures may reduce the risk of bleeding. Sodium-glucose co-transporter-2 inhibitors and glucagon-like peptide-1 agonists should also be discontinued to prevent acidosis and aspiration, respectively. Specific guidance for frequently used medications are complied within the manuscript, less frequently used medications are listed seperately.

Conclusions: Despite strong recommendations from major guidelines and consensus manuscripts, variation exists in preoperative medication orders, with limited availability of succinct implementation tools. This turnkey order set may facilitate standardized comprehensive preoperative medication management before nonemergent cardiac surgery. (JTCVS Open 2024;22:1-13)

Key aspects of preoperative medication management across perioperative phases of care.

CENTRAL MESSAGE

Integrating guideline recommendations and consensus statements into a turnkey order set may facilitate the implementation of preoperative medication management of nonemergent cardiac surgery patient.

PERSPECTIVE

Multiple professional societies have released evidence-based recommendations for preoperative medication management. Variation in practice still exists. The Enhanced Recovery After Surgery (ERAS) Cardiac Society has developed a "turnkey" order set to assist clinicians in adhering to best practices for preoperative medication management of the nonemergent adult cardiac surgery.

New York, NY; ⁱDepartment of Surgery, Baylor College of Medicine, Houston, Tex; ⁱSanger Heart & Vascular Institute, Charlotte, NC; ^kWakeMed Health, Raleigh, NC; ⁱTriStar Centennial Medical Center, Nashville, Tenn.

Received for publication March 5, 2024; revisions received May 25, 2024; accepted for publication June 10, 2024; available ahead of print July 29, 2024.

Address for reprints: Amanda Rea, DNP, CRNP, Division of Cardiac Surgery, University of Maryland St Joseph Medical Center, Suite 302, 7601 Osler Dr, Towson, MD 21204 (E-mail: amanda.rea@umm.edu).

2666-2736

Copyright © 2024 The Author(s). Published by Elsevier Inc. on behalf of The American Association for Thoracic Surgery. This is an open access article under the CC BY-NC-ND license (http://creativecommons.org/licenses/by-nc-nd/4.0/). https://doi.org/10.1016/j.xjon.2024.06.009

EBASE Condities Society

 Turnkey Order Set for Preopensities Medication Management of the Nonemergent Cardia: Surgery Planta

 Image: Stranger Planta

 I

From the ^aDivision of Cardiac Surgery, University of Maryland St Joseph Medical Center, Towson, Md; ^bDepartment of Surgery, University of Maryland School of Medicine, Baltimore Md; ^cDepartment of Anesthesiology and Critical Care Medicine, Johns Hopkins University, School of Medicine, Baltimore Md; ^dHeart & Vascular Program, Baystate Health, University of Massachusetts Chan Medical School-Baystate, Springfield, Mass; ^cDepartment of Surgery, Division of Cardiac Surgery, Harrington Heart and Vascular, Institute, University Hospitals, Case Western Reserve University, Cleveland, Ohio; ^fDepartment of Anesthesiology, Cumming School of Medicine & Libin Cardiovascular Institute, University of Calgary, Calgary, Alberta, Canada; and ^gProvidence Anesthesiology Associates, Charlotte, NC.

^{*} ERAS Cardiac Working Group: Busra Cangut, MD,^h Subhasis Chatterjee, MD,ⁱ Kevin W. Lobdell, MD,^j Gina McConnell BSN, RN,^k Shannon Crotwell BSN, RN,^j and Seenu Reddy MD, MBA¹. ^hIcahn School of Medicine at Mount Sinai,

Abbreviatio	ons and Acronyms
ACC	= American College of Cardiology
AHA	= American Heart Association
AmSECT	$\Gamma =$ American Society of ExtraCorporeal
	Technology
ASA	= American Society of Anesthesiologists
EACTS	= European Association for Cardio-
	Thoracic Surgery
ERAS	= Enhanced Recovery After Surgery
GLP-1	= glucagon-like peptide agonist
SABM	= Society for the Advancement of Patient
	Blood Management
SCA	= Society of Cardiovascular
	Anesthesiologists
SCAI	= Society for Cardiac Angiography and
	Interventions
SGLT2i	= sodium-glucose co-transporter-2
	inhibitor
STS	= Society of Thoracic Surgeons
TKO	= "turnkey" order set

The management of preoperative medications requires a provider to weigh the risk to benefits of drug therapy versus the potential for increased complications before the surgical procedure. For example, holding anticoagulation to reduce the risk of bleeding must be weighed against the risk of increased thrombotic complications.¹ Similar risk-benefit decisions must be considered for all medications during the perioperative period. There are no current comprehensive resources incorporating recommendations for most medication classes before cardiac surgery. This work used published guidelines and consensus documents to develop a practical preoperative medication order set for all nonemergent adult cardiac surgery patients. This "turnkey" order set (TKO) is part of a continued series created by the Enhanced Recovery After Surgery (ERAS) Cardiac Society to aid in the comprehensive implementation of evidence based standardized care.

METHODS

Leading multidisciplinary subject matter experts were consulted to analyze and translate existing guidelines and peer-reviewed literature into a sample TKO for preoperative medication management of the nonemergent cardiac surgery patient. Orders from consistent Class I, IIA, or comparable recommendations across referenced guidelines and consensus manuscripts appear in the TKO in bold type. Selected orders inconsistent with Class I or IIA, Class IIB, or supported by evidence published in peer-reviewed journals, are also included in italicized type. Table 1²⁻⁵ provides a synopsis of existing Class I and IIA (or equivalent) recommendations from applicable guidelines and consensus statements. Table 2 translates the recommendations, guidelines, consensus statements, and peer-reviewed literature into a TKO, with less frequently used medication noted in Appendix E1. Notably, the intent was not to recapitulate the evidence base justifying existing recommendations, as this task was part of the original guidelines and consensus statements efforts. Decisions regarding order inclusion were made on the basis of estimated benefit, risk, cost, implementation complexity, and generalizability. These orders should be considered on the basis of local institutional priorities, resources, practices, and expertise. When variation was found in literature, we used expert consensus and strength of literature to determine order inclusion. This TKO does not include consideration for patients with renal and/or hepatic dysfunction; prescribers should use additional precautions and consult a pharmacist when treating these patients. Whenever prescribing or holding a medication, consideration should include risk versus benefit, need for bridging therapies, medication to medication interactions, allergies, and potential adverse reactions. Medication holding day definition example can be viewed in Figure 1.

Comparisons Across Existing Guidelines

Strong-, moderate-, and low-risk recommendations from multiple sources were used to create the TKO. The guidelines chosen for comparison were published by The European Association for Cardiothoracic Surgery (EACTS),² The American College of Cardiology/American Heart Association/Society for Cardiac Angiography and Interventions (ACC/AHA/ SCAI),³ The Society of Thoracic Surgeons/Society of Cardiovascular Anesthesiologists/American Society of ExtraCorporeal Technology/Society for the Advancement of Patient Blood Management (STS/SCA/Am-SECT/SABM),⁴ and American Society of Anesthesiologists (ASA)⁵ (Table 1). The EACTS,² ACC/AHA/SCAI,³ and STS/SCA/AmSECT/ SABM⁴ documents are guidelines, and the ASA⁵ publication is an expert consensus document. The EACTS² and STS/SCA/AmSECT/SABM⁴ guidelines focus exclusively on cardiac surgery, whereas the ACC/AHA/ SCAI³ and ASA⁵ publications were intended for all surgical subspecialties.

Antithrombotic Agents

There is literature to support holding preoperative antithrombotic agents to reduce the risk of bleeding during cardiac surgery. EACTS and ACC/ AHA/SCAI both support continuation of aspirin throughout the preoperative period for patients with coronary artery disease.^{2,3} EACTS and STS/SCA/ AMSECT/SABM both state that it is reasonable to stop aspirin 7 days before noncoronary cardiac surgery to reduce the risk of bleeding.^{2,4} EACTS, ACC/ AHA/SCAI, and STS/SCA/AmSECT/SABM all recommend ticagrelor should be held preoperatively for a minimum of 3 days, clopidogrel for 5 days, and prasugrel for 7 days before nonemergent surgery.^{2,3,5-9} The half-life of ticagrelor is about 7 hours, and it inhibits platelet aggregation in a reversible manner. This is in contrast to other antiplatelet medications like clopidogrel, which irreversibly inhibit platelets for the lifetime of the platelet (7-10 days).^{23,24} As the result of its relatively short half-life and reversible action, the antiplatelet effect of ticagrelor diminishes more quickly once the medication is stopped, allowing platelet function to recover in time for surgery, thereby reducing the risk of excessive bleeding. Although hold recommendations for cangrelor range from 1 to 6 hours, providers are now routinely holding the medication for 1 hour before surgery.¹⁰ Providers may choose to be more conservative in high-risk cases, such as a redo sternotomy, multiple valve, and advanced aortic cases.

EACTS and a clinical recommendation from the American College of Chest Physicians recommend holding non-vitamin K antagonist oral anticoagulants for 2 to 4 days. The range depends on the medication and guidelines differ slightly on recommended time to hold before surgery.^{2,8} EACTS recommends abciximab to be held for 24 hours and eptifibatide tirofiban, 4 hours before surgery.² Warfarin is recommended to be held for 5 days before surgery and to check an international normalized ratio.^{2,8} Intravenous heparin is recommended to be held 6 hours and low molecular weight heparin held 1 day before surgery.^{2,8} Patients should be assessed for need of bridging therapy with either intravenous short-acting anticoagulation or antiplatelet medications.

· · · ·				
Recommendation	EACTS ² 2017	ACC/AHA/SCAI ³ 2021	STS/SCA/AmSECT/SABM ⁴ 2021	ASA ⁵ 2023
Antithrombotic management				
Aspirin	\checkmark	\checkmark	\checkmark	
DAPT	\checkmark	\checkmark	\checkmark	
GPIIb/IIIa inhibitors	\checkmark	\checkmark		
Vitamin K antagonists	\checkmark			
NOAC/DOAC	\checkmark			
Renin-angiotensin aldosterone system inhibitors				
ACE/ARB	\checkmark			
Beta-blockers				
Beta-blocker	\checkmark			
Statins				
Statin	\checkmark			
Glucose management				
Oral hypoglycemics and long-acting subcutaneous insulin	\checkmark			
GLP-1 agonists				\checkmark

TABLE 1. Summary of Class I/IIa or equivalent recommendations

A " $\sqrt{}$ " indicates class I/IIA or equivalent recommendations for preoperative medication for nonemergent **adult cardiac surgery**. *EACTS*, European Association for Cardiothoracic Surgery; *ACC/AHA/SCAI*, The American College of Cardiology American Heart Association, Society for Cardiac Angiography and Interventions; *STS/SCA/AmSECT/SABM*, Society of Thoracic Surgeons, Society of Cardiovascular Anesthesiologists, American Society of ExtraCorporeal Technology, Society for the Advancement of Patient Blood Management; *ASA*, American Society of Anesthesiologists; *DAPT*, dual antiplatelet therapy; *GPIIb/IIa*, glycoprotein IIb/IIIa; *NOAC*, non-vitamin K antagonist oral anticoagulant; *DOAC*, direct-acting oral anticoagulant; *ACE-I*, angiotensin-converting-enzyme inhibitor; *ARB*, angiotensin receptor blocker; *GLP-1*, glucagon-like peptide 1.

Endocrine Agents

There has been increasing use of glucagon-like peptide (GLP-1) agonists and sodium-glucose co-transporter-2 inhibitors (SGLT2i), as benefits outside the treatment of diabetes have now been demonstrated.²⁵ There are concerns for increased risk of aspiration as the result of delayed gastric emptying associated with the use of GLP-1 medications. A recent consensus statement from the ASA recommends holding daily dosed GLP-1 agonists the day of surgery and weekly dosed GLP-1 agonists for 7 days before surgery to theoretically reduce the potential for aspiration with induction of anesthesia.^{5,11,12} Other adverse events that have been reported with the use of GLP-1 medications include gastrointestinal disorders, gallbladder-related disorders (mostly cholelithiasis), cardiovascular and psychiatric disorders, and malignant neoplasm.²⁶

SGLT2i are recommended to be held 3 days before surgery, except ertugliflozin, which should be held 4 days before surgery because of its long half-life and increased risk of euglycemic diabetic ketoacidosis.^{7,11-18} All other oral hypoglycemics are recommended to be held the day of surgery. If metformin is co-prescribed with an SGLT2i, it should be held for 3 days before surgery.^{7,11-16}

Short-acting insulin is recommended to be held the day of surgery. For long-acting insulin, 75% of the dose is commonly taken the day before surgery and 50% of the dose on the day of surgery.^{7,11-13,15} The EACTS recommendation does not include specific medication classes and states that antidiabetics and long-acting subcutaneous insulin should be omitted the day before surgery.²

Cardiac Agents

 β -blockers are recommended to be continued until surgery.^{2,3,7,18-21} Angiotensin-converting enzyme inhibitors and angiotensin receptor blockers are commonly held the day before cardiac surgery to reduce the risk of vasoplegia.^{2,7,18} In contrast, it is reasonable to continue vasodilators such as hydralazine and nitroglycerin.^{7,18} Statin therapy is frequently continued before cardiac surgery.^{2,7,21} There are no large studies, guidelines, or recommendations regarding other cardiac agents. Although not addressed by guidelines or consensus manuscripts, diuretics and direct renin inhibitors are generally held the day of surgery because of concern for adverse effects on renal function in the setting of volume depletion and/or hypotension.^{7,18} phosphodiesterase type 5 inhibitors (ie, sildenafil and tadalafil) are frequently held 72 hours before surgery when used for erectile dysfunction as the result of concerns related to vasodilation and hypotension.^{7,18,22} When phosphodiesterase type 5 inhibitors are used for pulmonary hypertension or cardiac optimization, they are regularly continued, as they have been shown to decrease inotropic support and ventilator hours in patients with right ventricular failure and pulmonary hypertension.^{7,18,22,27} Calcium channel blockers and antiarrhythmics are generally continued.^{2,3,7,18}

Future Research

There are insufficient publications to guide the use of cannabis before surgery. The use of delta-9-tetrahydrocannabinol, or THC, should be disclosed to the anesthesiologist.²⁸ Herbal supplements are another area in which the literature is inconsistent, with some resources recommending discontinuation of all supplements and herbal medications for 7 days, whereas others recommend 2 weeks before surgery.^{7,29} Further research and publications are needed. Additional research is needed to study how best to educate patients and assist with consistent safe multidisciplinary preoperative medication guidance. This can include formal multidisciplinary education and introducing materials through multiple platforms, including web-based or personal phonebased applications.³⁰

CONCLUSIONS

Literature exists to guide practitioners in evidence-based best practice for most preoperative medications used by adult cardiac surgical patients. However, the availability of a centralized single reference of up-to-date

TABLE 2. Orders

Medication class	Medication example(s)	Recommendation
Antiplatelet agents		
In the absence of	Aspirin	Hold 7 d before surgery (patients without
platelet function studies		CAD or other strong indication) ^{2-4,6,7}
		Continue for patients with CAD ^{2,3,050}
	Dipyridamole	Hold 2 d before surgery'
	Cilostazol	Hold 5 d before surgery ⁷
	Clopidogrel	Hold 5 d before surgery ^{2-4,6-9}
	Prasugrel	Hold 7 d before surgery ^{2-4,6-8}
	Ticagrelor	Hold 3 d before surgery ^{2-4,7-9}
	Cangrelor	Hold 1-6 h before surgery ^{2,10}
	Eptifibatide	Hold 4 h before surgery ^{2,3,6}
	Tirofiban	
	Abciximab	Hold 24 h before surgery ^{2,3,6}
Anticoagulants		
	Warfarin	Hold 5 d before surgery and check an INR ^{2,8}
	Henarin IV	Hold 6 h before surgery ^{2,8}
		Hold 1 d b for an and 2.8
	LMWH	Hold 1 d before surgery
Anticoagulants-NOAC		
	Apixaban	Hold 2-4 d before surgery; consider renal impairment when holding ^{2,4,8}
	Dabigatran	Hold 2-4 d before surgery; consider renal impairment when holding ^{2,4,8}
	Edoxaban	Hold 2-4 d before surgery; consider renal impairment when holding ^{2,4,8}
	Rivaroxaban	Hold 2-4 d before surgery; consider renal impairment when holding ^{2,4,8}
Endocrine agents		
GLP-1 agonists (daily dosing)	Exenatide	Hold the day of surgery ^{5,11,12}
	Liraglutide	
	Lixisenatide	
	Semaglutide	512
GLP-1 agonists (weekly dosing)	Dulaglutide	Hold 7 d before surgery ^{3,12}
	Exenatide extended release	
Dipentidul pentidase-4 inhibitors	Glyburide	Hold the day of surgery $7,11-15$
Secretogogues	Glipizide	nou ne day of surgery
Thiazolidinediones	Glimepiride	
	Pioglitazone	
	Saxagliptin	
	Linagliptin	
	Vildagliptin	
	Sitagliptin	
	Alogliptin	
Oral nypogiycemics-biguanides	Metformin	<i>co-prescribed with an SGLT-2, then hold</i> <i>3 d before</i> ^{7,11-16}

TABLE 2. Continued

Medication class	Medication example(s)	Recommendation
Sodium-glucose co-transporter-2	Canaglifozin	Hold 3 d before surgery except ^{7,11-13,15-18}
inhibitors (SGLT2i)	Dapagliflozin	*Ertugliflozin should be held 4 d before surgery ^{12,13,18}
	Empagliflozin	
	Ertugliflozin*	
Insulin, long-acting	Detemir	Take 75% of dose day before surgery, take
	Glargine	50% of dose day of surgery ^{11,13,15}
Insulin, short-acting	Lispro	Hold the day of surgery ^{11,13,15}
	Aspart	
	Regular	
Cardiac medications		
ACE-I	Benazapril	Hold the day before surgery ^{2,7,18}
ARB	Lisinopril	
	Enalapril	
	Ramipril	
	Captopril	
	Perindopril	
	Losartan	
	Valsartan	
	Candesartan	
	Irbesartan	
	Sacubitril/valsartan	
	Olmesartan	
	Telmisartan	
	Azilsartan	
	Eprosartan	
Antiarrhythmics	Amiodarone	$Continue^{2,3,7,18}$
	Dronaderone	
	Digoxin	
Beta blocker	Metoprolol	<i>Continue</i> ^{2,3,7,18-21}
	Carvedilol	
Calcium channel blockers	Amlodipine	Continue ^{7,18}
	Diltiazem	
	Felodipine	
	Isradipine	
	Nicardipine	
	Nifedipine	
	Verapamil	
Diuretics	Bumetanide	Hold the day of surgery ^{7,18}
	Furosemide	
	Hydrochlorothiazide	
	Triamterene	
	Torsemide	
	Spironolactone	
Phosphodiesterase	Sildenafil	Hold 72 h before surgery when used for
type 5 inhibitors	Tadalafil	the indication of erectile dysfunction for
	Vardenafil	pulmonary hypertension continue ^{7,18,22}

CAD, Coronary artery disease; *INR*, international normalize ratio; *IV*, intravenous; *LMWH*, low-molecular-weight heparin; *NOAC*, non-vitamin K antagonist oral anticoagulant; *GLP-1*, glucagon-like peptide 1; *SGLT-2*, sodium/glucose cotransporter 2; *ACE-1*, angiotensin-converting-enzyme inhibitor; *ARB*, angiotensin receptor blocker; *MAOI*, mono-amine oxidase inhibitor.

recommendations is lacking. We have combined these resources and guidelines to provide a referenced practical TKO reference for clinicians caring for nonemergent adult cardiac surgery patients.

Conflict of Interest Statement

A.R.: Speaker's Bureau for Edwards Lifesciences; R.S.: consulting/advisory relationships with Zimmer Biomet, AtriCure, La Jolla, Terumo, Encare, and Edwards

		Last	Hold 2 d	ays prior to s	surgery				
					Day of Surgery				
-5	-4	-3	-2	-1	POD 0	POD 1	POD 2	POD 3	POD 4

FIGURE 1. Holding day definition: day dosing does not include surgery day (ex: "hold 2 days before surgery" if Wednesday is surgery day, last dose should be taken Sunday).

Lifesciences. R.C.A.: honoraria from Edwards Lifesciences and HLS Therapeutics. In addition, he has function on an Ad Board for Renibus Therapeutics Inc. A.J.G.: speaker and advisory activities for Edwards Lifesciences. V.M.B.: Speaker's Bureau Edwards Lifesciences. D.T.E.: Device Safety Monitoring Board: Edwards Lifesciences Transcatheter Valves; Trial Steering Committees: Renibus, Alexion, Cardiorenal Systems, Genentech; Medical Advisory Boards: Medela, Arthrex, and AtriCure. S.C.: served on advisory boards for Edwards Lifesciences, La Jolla Pharmaceutical Company, Baxter Healthcare, and Eagle Pharmaceuticals. K.W.L.: consultant for Abiomed, Alexion, Medela, Medtronic, and Renibus. G.M.: consultant for Edwards Lifesciences. S.R.: advisor and speaker for 3M. All other authors reported no conflicts of interest.

The *Journal* policy requires editors and reviewers to disclose conflicts of interest and to decline handling or reviewing manuscripts for which they may have a conflict of interest. The editors and reviewers of this article have no conflicts of interest.

References

- Polania Gutierrez JJ, Rocuts KR. Perioperative anticoagulation management. In: StatPearls. StatPearls Publishing; 2023.
- Sousa-Uva M, Head S, Milojevic M, et al. 2017 EACTS guidelines on perioperative medication in adult cardiac surgery. *Eur J Cardiothorac Surg.* 2018;53: 5-33. https://doi.org/10.1093/ejcts/ezx314
- Lawton JS, Tamis-Holland JE, Bangalore S, et al. 2021 ACC/AHA/SCAI guideline for coronary artery revascularization. JAm Coll Cardiol. 2021;79(2). https:// doi.org/10.1016/j.jacc.2021.09.006
- Tibi P, McClure RS, Huang J, et al. STS/SCA/AmSECT/SABM update to the clinical practice guidelines on patient blood management. J Extra Corpor Technol. 2021;53(2):97-124. https://doi.org/10.1182/ject-2100053
- American Society of Anesthesiologists consensus-based guidance on preoperative management of patients (adults and children) on glucagon-like peptide-1 (GLP-1) receptor agonists. Accessed February 19, 2024. www.asahq.org, https://www.asahq.org/about-asa/newsroom/news-releases/2023/06/americansociety-of-anesthesiologists-consensus-based-guidance-on-preoperative
- Demertzis S, Cassina T, Casso G. Antiplatelet therapy before cardiac surgery. Cardiovasc Med. 2016;19(4):110-116.
- Froedtert. Guideline for perioperative medication management; 2020. Accessed February 19, 2024. https://www.froedtert.com/sites/default/files/upload/docs/ professionals/physicians/preoperative/perioperative-medication-management.pdf
- Douketis JD, Spyropoulos AC, Murad MH, et al. Perioperative management of antithrombotic therapy: an American College of Chest Physicians clinical practice guideline. *Chest.* 2022;162(5):e207-e243.

- Hansson EC, Jideus L, Aberg B, et al. Coronary artery bypass grafting-related bleeding complications in patients treated with ticagrelor or clopidogrel: a nationwide study. *Eur Heart J.* 2016;37(2):189-197.
- Angiolillo DJ, Firstenberg MS, Price MJ, et al. Bridging antiplatelet therapy with cangrelor in patients undergoing cardiac surgery: a randomized controlled trial. *JAMA*. 2012;307(3):265-274.
- Sreedharan R, Khanna S, Shaw A. Perioperative glycemic management in adults presenting for elective cardiac and non-cardiac surgery. *Periop Med*. 2023;12(1):13.
- Pfeifer KJ, Selzer A, Mendex CE, et al. Preoperative management of endocrine, hormonal, and urologic medications: Society for Perioperative Assessment and Quality Improvement (SPAQI) consensus statement. *Mayo Clinic Proc.* 2021; 96(6):1655-1669.
- American Diabetes Association Professional Practice Committee. Diabetes care in the hospital: standards of medical care in diabetes—2022. *Diabetes Care*. 2022;45(Suppl 1):S244-S253.
- Preiser JC, Provenzano B, Mongkolpun W, Halenarova K, Cnop M. Perioperative management of oral glucose-lowering drugs in the patient with type 2 diabetes. *Anesthesiology*. 2020;133:430-438.
- Duggan EW, Carlson K, Umpierrez GE. Perioperative hyperglycemia management. Anesthesiology. 2017;126(3):547-560.
- Australian Diabetes Society. Periprocedural diabetic ketoacidoss (DKA) with SGLT2 inhibitor use; 2020. Accessed February 19, 2024. https://diabetessociety. com.au/documents/ADS_DKA_SGLT2i_Alert_update_2020.pdf
- Thiruvenkatarajan V, Jesudason D, Nanjappa N, Meyer EJ, Van Wijk RM. Perioperative management of glucose-lowering drugs: comment. *Anesthesiology*. 2021;134(2):349-350. https://doi.org/10.1097/ALN.00000000003626
- Sahai SK, Balonov K, Bentov N, et al. Preoperative management of cardiovascular medications: a Society for Perioperative Assessment and Quality Improvement (SPAQI) consensus statement. *Mayo Clin Proc.* 2022;97(9):1734-1751.
- Chan AY, McAlister FA, Norris CM, Johnstone D, Bakal JA, Ross DB. Effect of beta-blocker use on outcomes after discharge in patients who underwent cardiac surgery. J Thorac Cardiovasc Surg. 2010;140:182-187. 187.e1.
- 20. ten Broecke PW, De Hert SG, Mertens E, Adriaensen HF. Effect of preoperative beta-blockade on perioperative mortality in coronary surgery. *Br J Anaesth.* 2003;90(1):27-31.
- Daumerie G, Fleisher LA. Perioperative beta-blocker and statin therapy. *Curr* Opin Anaesthesiol. 2008;21:60-65.
- Pfeifer KJ, Selzer A, Whinney CM, et al. Preoperative management of gastrointestinal and pulmonary medications: Society for Perioperative Assessment and Quality Improvement (SPAQI) consensus statement. *Mayo Clinic Proc.* 2021; 96(12):3158-3177.
- Hansson EC, Malm CJ, Hesse C, et al. Platelet function recovery after ticagrelor withdrawal in patients awaiting urgent coronary surgery. *Eur J Cardiothorac* Surg. 2017;51(4):633-637.
- Adamski P, Skonieczny G, Hajdukiewicz T, Kern A, Kubica J. Reversal of platelet inhibition in patients receiving ticagrelor. *Rev Cardiovasc Med.* 2022;23:300.
- Brown E, Heerspink HJL, Cuthbertson DJ, Wilding JPH. SGLT2 inhibitors and GLP-1 receptor agonists: established and emerging indications. *Lancet.* 2021; 398(10296):262-276.
- **26.** Rubino DM, Greenway FL, Khalid U, et al. Effect of weekly subcutaneous semaglutide vs daily liraglutide on body weight in adults with overweight or obesity without diabetes: the STEP 8 randomized clinical trial. *JAMA*. 2022;327(2): 138-150.

- Villanueva DLE, Agustin RD, Llanes EJ. Pre-operative sildenafil for patients with pulmonary hypertension undergoing mitral valve surgery: a systematic review and meta-analysis. *Cardiol Res.* 2019;10(6):369-377.
- Shah S, Schwenk ES, Sondekoppam RV, et al. ASRA pain medicine consensus guidelines on the management of the perioperative patient on cannabis and cannabinoids. *Reg Anesth Pain Med.* 2023;48(3):97-117. https://doi.org/10.1136/ rapm-2022-104013
- Cummings KC, Keshock M, Ganesh R, et al. Preoperative management of surgical patients using dietary supplements: Society for Perioperative Assessment and Quality Improvement (SPAQI) consensus statement. *Mayo Clinic Proc.* 2021; 96(5):1342-1355.
- Grant MC, Crisafi C, Alvarez A, et al. Perioperative care in cardiac surgery: a joint consensus statement by the Enhanced Recovery After Surgery (ERAS) Cardiac Society, ERAS International Society, and The Society of Thoracic Sur geons (STS). *Ann Thorac Surg.* 2024;117(4):669-689. https://doi.org/10.1016/j. athoracsur.2023.12.006
- Kumar A, Moises A, Aneja A, Mohr F, Jain A, Shen B. Inflammatory bowel disease: Perioperative pharmacological considerations. *Mayo Clinic Proc.* 2011; 86(8):748-757.
- Lim S, Rogers LK, Tessler O, Mundinger GS, Rogers C, Lau FH. Phentermine: a systematic review for plastic and reconstructive surgeons. *Ann Plastic Surg.* 2018;81(4):503-507. https://doi.org/10.1097/SAP.000000000001478
- 33. Goodman SM, Springer B, Guyatt G, et al. 2017 American College of Rheumatology/American Association of Hip and Knee Surgeons guideline for the perioperative management of antirheumatic medication in patients with rheumatic diseases undergoing elective total hip or total knee arthroplasty. *Arthritis Care Res.* 2017;69(8):1111-1124.
- MacKenzie CR. Perioperative medication management in the rheumatic diseases. Curr Rev Musculoskelet Med. 2017;10:404-406.

- 35. Goodman SM, Springer BD, Chen AF, et al. 2022 American College of Rheumatology/American Association of Hip and Knee Surgeons guideline for the perioperative management of antirheumatic medication in patients with rheumatic diseases undergoing elective total hip or total knee arthroplasty. *Arthritis Care Res.* 2022;74:1399-1408.
- 36. Russell LA, Craig C, Flores E, et al. Preoperative management of medications for rheumatologic and HIV diseases: Society for Perioperative Assessment and Quality Improvement (SPAQI) consensus statement. *Mayo Clinic Proc.* 2022; 97(8):1551-1571.
- Oprea AD, Keshock MC, O'Glasser AY, et al. Preoperative management of medications for psychiatric diseases. *Mayo Clinic Proc.* 2022;97(2):397-416.
- O'Rourke MJ, Keshock MC, Boxhorn CE, et al. Preoperative management of opioid and nonopioid analgesics: Society for Perioperative Assessment and Quality Improvement (SPAQI) consensus statement. *Mayo Clinic Proc.* 2021;96(5):1325-1341.
- 39. Doleman B, Leonardi-Bee J, Heinink TP, et al. Pre-emptive and preventive NSAIDs for postoperative pain in adults undergoing all types of surgery. *Cochrane Database Syst Rev.* 2021;6(6):CD012978.
- Hyland SJ, Brickhaus KK, Vincent WR, et al. Perioperative pain management and opioid stewardship: a practical guide. *Healthcare*, 2021;9(3):333.
- Kaye AD, Kline RJ, Thompson ER, et al. Perioperative implications of common and newer psychotropic medications used in clinical practice. *Best Pract Res Clin Anaesthesiol.* 2018;32(2):187-202.
- Libert N, Tourtier J, Védrine L, Chargari C, Riou B. Inhibitors of angiogenesis: new hopes for oncologists, new challenges for anesthesiologists. *Anesthesiology*. 2010;113:704-712.

Key Words: preop medication, turnkey order set, cardiac surgery, perioperative, medication management

APPENDIX E1. MEDICATIONS BY CLASS

Class	Example	Recommendation
Acetylcholinesterase inhibitors	Donazepil Galantamine Rivastigmine Tacrine	<i>Continue</i> ⁷
Alpha blocker	Alfuzosin Doxazosin Phenoxybenzamine Phentolamine Prazosin Silodosin Tamsulosin Terazosin	Continue ^{7,12,18}
Alpha 2 agonists—central-acting	Clonidine Guanfacine	Continue ^{7,18}
Aminosalicylates	Mesalamine Sulfasalazine Azathioprine	Hold 1 d before surgery ^{7,31} Hold the day of surgery ³¹
ACE-I ARB	Benazapril Lisinopril Enalapril Ramipril Captopril Perindopril Losartan Valsartan Candesartan Irbesartan Sacubitril/valsartan Olmesartan Telmisartan Azilsartan Eprosartan	Hold the day before surgery ^{2,7,18}
Anorectics	Phentermine	Hold for 7 days before surgery ³²
Antiarrhythmics	Amiodarone Dronaderone Digoxin	<i>Continue</i> ^{2,10}
Antibiotic prophylaxis	Mupirocin	<i>Continue</i> ²
Anticonvulsants	Carbamazepine Felbamate Gabapentin Levetiracetam Lamotrigine Oxcarbazepine Phenytoin Pregabalin Primidone Tiagabine Topiramate Valproic acid Zonisamide	<i>Continue</i> ⁷

Class	Example	Recommendation
Antigout agents	Allopurinol Colchicine Febuxostat Probenecid	Continue ⁷
Antihistamines	Loratadine Diphenhydramine Cetirizine Chlorpheniramine	<i>Continue</i> ⁷
Antimetabolites Antirheumatics	Methotrexate* Leflunomide* Hydroxychloroquine* Sulfasalazine	Continue ^{7,33-36} *Methotrexate is recommended to be continued preoperatively in patients with normal renal function, but held for 2 wk before surgery in patients with renal impairment. *Leflunomide stopped 2 wk before surgery. *Hydroxychloroquine—insufficient evidence for recommendation of cardiac surgery
Antimigraine agents—triptans	Almotriptan Eletriptan Frovatriptan Naratriptan Rizatriptan Sumatriptan Zolmitriptan	Hold the day before surgery ⁷
Antipsychotics	Haloperidol Risperidone Olanzapine	Continue ^{7,37}
Antispasmodic	Oxybutynin Darifenacin Tolterodine	Continue ^{7,38}
Antivirals Antiretrovirals	Non-nucleoside reverse transcriptase inhibitor Protease inhibitors, Integrase inhibitors (would be too many to list individually)	Continue ^{7,36}
Appetite suppressant	Phentermine	Hold 7 d before surgery ⁷
Benzodiazepines	Alprazolam Chlordiazepoxide Diazepam Clonazepam	Continue ^{7,37}
Beta blocker	Metoprolol Carvedilol	<i>Continue</i> ^{2,3,7,18-21}
Biologic agents (including any applicable biosimilars)	Adalimumab Etanercept Golimumab Infliximab Abatacept Certolizumab Rituximab Tocilizumab Anakinra Secukinumab	Hold timing for end of dosing cycle 2-8 wk pending medication ³³⁻³⁵

Class	Example	Recommendation
Biologic disease-modifying agents	Interferon beta Glatiramer acetate Siponimod Fingolimod Ustekinumab Belimumab Tofacitinib* Upadacitinib* Baricitinib*	General should be stopped at end of dosing cycle but should be discussed with prescribing team. *Should be stopped 7 d before surgery ^{33,35,36}
Bisphosphonates and antiosteoporosis agents	Alendronate Ibandronate Risedronate Zoledronic acid Denosumab Calcitonin-salmon	Hold the day of surgery ^{7,12}
Calcium channel blockers	Amlodipine Diltiazem Felodipine Isradipine Nicardipine Nifedipine Verapamil	<i>Continue</i> ^{7,18}
Cannabis		No guidance at this time, but should be disclosed to the anesthesiologist ²⁸
Central nervous system depressant	Zolpidem	Continue ³⁷
Central nervous system stimulants	Dextroamphetamine Lisdexamfetamine Dexmethylphenidate Methylphenidate	Hold the day of surgery ^{7,37}
Cox-2 inhibitors	Celecoxib	Hold 3 d before surgery ⁷
Direct renin inhibitors	Aliskiren and combination products	Hold the day before surgery ⁷
Diuretics	Bumetanide Furosemide Hydrochlorothiazide Triamterene Torsemide Spironolactone	Hold the day of surgery ^{7,18}
Dopamine agonist	Ropinirole Pramipexole Carbidopa/Levodopa	<i>Continue</i> ^{7,12}
Glucocorticoids	Budesonide Dexamethasone Hydrocortisone Methylprednisolone Prednisolone Prednisone	Continue ^{7,31,36}
Histamine H2 blockers	Cimetidine Famotidine Ranitidine Nizatidine	<i>Continue</i> ^{7,22}

Class	Example	Recommendation
IgG monoclonal antibody	Brodalumab Guselkumab Risankizumab Secukinumab Tildrakizumab Ustekinumab	Agents are recommended to be stopped before surgery and that surgery is scheduled at the end of the dosing cycle ^{33,36}
Immunosuppressive agents	Mycophenolate Mofetil Cyclosporine Tacrolimus Azathioprine Prednisone	Continue ^{7,33-36}
Inhaled pulmonary agents	Budesonide Fluticasone Montelukast Prednisone Tiotropium Albuterol Albuterol/ipratropium Formoterol/budesonide Formoterol/mometasone Ipratropium Levalbuterol Salmeterol Salmeterol/fluticasone	Continue ^{7,18}
MAOI	Isocarboxazid Moclobemide Phenelzine Tranylcypromine	Continue unless unable to provide MAOI-safe anesthetic, then discontinue under psychiatric guidance 2 wk before the procedure ^{7,37}
Mood stabilizers	Lithium* Carbamazepine Lamotrigine Oxcarbazepine Valproic acid	Continue *Hold for 72 h ³⁷
Non-statin lipid-lowering agents	Cholestyramine Colestipol Ezetimibe Fenofibrate Gemfibrozil Niacin	Hold the day before surgery ⁷
Nonsteroidal anti-inflammatory agents	Short-acting Ibuprofen, Diclofenac Etodolac Indomethacin Intermediate-acting Naproxen Meloxicam Oxaprozin	Hold 7 d before surgery ^{7,36,39}

Opioid agonists

Class

Example	Recommendation
Codeine Fentanyl Hydromorphone Methadone Morphine Oxycodone Oxymorphone Tramadol	Continue in perioperative period, with some considerations ^{38,40}
Buprenorphine Naltrexone*	Continue *Intramuscular naltrexone should be held 24-30 d after the last injection; oral naltrexone should be held 3-4 d ^{37,39,41}
Sildenafil	Hold 72 h before surgery when used

Opioid antagonists	Buprenorphine Naltrexone*	Continue *Intramuscular naltrexone should be held 24-30 d after the last injection; oral naltrexone should be held 3-4 d ^{37,39,}
Phosphodiesterase type 5 inhibitors	Sildenafil Tadalafil Vardenafil	Hold 72 h before surgery when used for the indication of erectile dysfunction for pulmonary hypertension continue ^{7,18,22}
Postmenopausal hormone therapy, oral contraceptive(s)	Estrogens (oral)	Hold 6 wk before surgery ⁷
Proton pump inhibitors	Esomeprazole Lansoprazole Omeprazole Pantoprazole Rabeprazole	Continue ^{7,22}
Pulmonary agents	Theophylline	Hold the day before surgery ^{7,22}
Selective estrogen receptor modulators	Tamoxifen Raloxifene	Hold the day of surgery ^{7,12}
Selective serotonin reuptake inhibitors Serotonin and norepinephrine reuptake inhibitor Aminoketone	Fluoxetine Citalopram Escitalopram Duloxetine Desvenlafaxine Duloxetine Levomilnacipran Milnacipran Venlafaxine Bupropion Hydrochloride	Continue ^{7,37}
Skeletal muscle relaxants	Cyclobenzaprine Metaxalone Tizanidine Methocarbamol	<i>Continue</i> ⁷
Statin	Atorvastatin	<i>Continue</i> ^{2,7,21}
Supplements, herbals		Discontinue all supplements and herbals 7 d before surgery ^{7,29} (2 wk)
Thyroid agents	Levothyroxine Methimazole Propylthiouracil	Continue ^{7,12}
Triazolopyridines		Continue ³⁷

Class	Example	Recommendation
Tricyclic antidepressant	Amitriptyline Nortriptyline Clomipramine Doxepin Imipramine	Continue ^{7,37}
Tyrosine kinase inhibitors (oncologic indications only)	Acalabrutinib Dasatinib Imatinib Erlotinib Everolimus Nintedanib	Hold 7 d before surgery in coordination with the patient's prescribing physician ⁴²
Vasodilators	Hydralazine Isosorbide dinitrate Isosorbide Mononitrate Minoxidil	Continue ^{7,18}

ACE-I, Angiotensin-converting enzyme inhibitors; ARB, angiotensin II receptor blockers; MAOI, monoamine oxidase inhibitor.