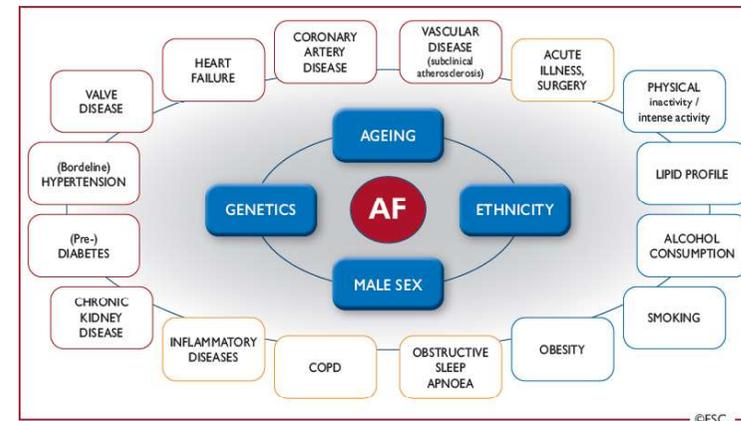


VHF und PCI: OAK, NOAK, Duale oder Triple Therapie: Was für welchen Patienten?

PD Dr. med. Gabor Sütsch

Figure 3 Summary of risk factors for incident AF



www.escardio.org/guidelines

2020 ESC Guidelines for the diagnosis and management of atrial fibrillation (European Heart Journal 2020-doi/10.1093/eurheartj/ehaa612)

Review article

Patients with atrial fibrillation and coronary artery disease – Double trouble

Ewelina Michniewicz^a, Elżbieta Młodawska^a, Paulina Lopatowska^a, Anna Tomaszuk-Kazberuk^a, Jolanta Malyszko^b

Coronary artery disease (CAD) is the **most common cardiovascular disease** while **atrial fibrillation (AF) is the most common cardiac arrhythmia**. Both diseases share associated risk factors – hypertension, diabetes mellitus, sleep apnea, obesity and smoking. Moreover, inflammation plays a causative role in both diseases. **The prevalence of CAD in patients with AF is from 17% to 46.5%** while the **prevalence of AF among patients with CAD is low and it is estimated from 0.2% to 5%**. AF is a well-established factor of poor short- and long-term prognosis in patients with acute myocardial infarction (AMI) and is associated with a marked increase in overall mortality.

The arrhythmia is common after cardiac surgeries and occurs in about 20 to 40% of patients after coronary artery bypass graft (CABG) surgery. It is predicted that between 5 and 15% of AF patients will require stenting at some point in their lives and will receive triple therapy with aspirin, clopidogrel or ticagrelor and oral anticoagulation (OAC).

| Clinical Presentation | AF-related OUTCOMES | | |
|---|---------------------------------------|---|---|
| | AF-Related Outcome | Frequency in AF | Mechanism(s) |
| Asymptomatic or Silent (!) | Death | 1.5-3.5 fold increase | Excess mortality related to: • HF, comorbidities • Stroke |
| Symptomatic Palpitations, dyspnoea, fatigue, Chest tightness/pain, poor effort tolerance, dizziness, syncope, disordered sleep, etc. | Stroke | 20-30% of all ischaemic strokes, 10% of cryptogenic strokes | • Cardioembolic, or • Related to comorbid vascular atheroma |
| | LV dysfunction / Heart failure | In 20-30% of AF patients | • Excessive ventricular rate • Irregular ventricular contractions • A primary underlying cause of AF |
| Haemodynamically unstable • Syncope • Symptomatic hypotension • Acute HF, pulmonary oedema • Ongoing myocardial infarction • Cardiogenic shock | Cognitive decline / Vascular dementia | HR 1.4 / 1.6 (irrespective of stroke history) | • Brain white matter lesions, inflammation, • Hypoperfusion, • Micro-embolism |
| | Depression | Depression in 16-20% (even suicidal ideation) | • Severe symptoms and decreased QoL • Drug side effects |
| Haemodynamically stable | Impaired quality of life | >60% of patients | • Related to AF burden, comorbidities, psychological functioning and medication • Disturbed personality type |
| | Hospitalizations | 10-40% annual hospitalization rate | • AF management, related to HF, MI or AF-related symptoms • Treatment-associated complications |

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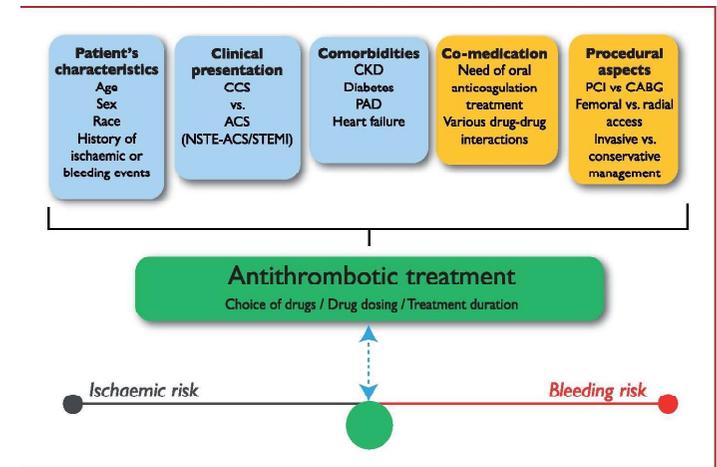
2020 ESC Guidelines for the diagnosis and management of atrial fibrillation (European Heart Journal 2020-doi/10.1093/eurheartj/ehaa612)

Figure 4 Clinical presentation of AF and AF-related outcomes

Patients with atrial fibrillation and coronary artery disease

- The combination of AF and CAD is not only a common clinical scenario, it is also a complex setting to combine anticoagulation and antiplatelet therapy.
- AF patients with relevant CAD have at least a CHA₂DS₂-VASc score of 1 (and mostly higher due to the presence of other cardiovascular risk factors) and hence an indication for OAC.
- The convention is that a period of DAPT (i.e. aspirin and a P2Y₁₂ inhibitor) is necessary to prevent stent thrombosis or recurrent events after an ACS and/or stenting for CAD—but that this is not sufficient for stroke prevention.
- Conversely, NOACs are essential for stroke prevention but on their own insufficient for preventing new coronary events in the immediate phase after ACS or stenting. The choice of antithrombotic drug combinations therefore represents a demanding clinical task: **too little and risk a coronary event and/or stroke, too much and risk a bleeding event.**

Determinants of antithrombotic treatment in coronary artery disease.



Eur Heart J, Volume 42, Issue 14, 7 April 2021, Pages 1289–1367

Typical situation

- A 68-years old patient undergoes elective coronary artery stenting on January 1st 2021. He is discharged on aspirin and clopidogrel. Uneventful interval. LV-EF is normal.
- Arterial hypertension, prediabetes, hypercholesterolemia
- On July 1st, 2021, he is diagnosed with new onset AF.
- It was determined to administer long-term OAC (CHA₂DS₂-VASc-Score >3)
- Strategy?
- Aspirin was stopped, and the patient is treated with OAC plus clopidogrel for one year
- At that time antithrombotic therapy will be re-evaluated.

Table 9 factors for bleeding with OAC and antiplatelet therapy

| Non-modifiable | Potentially modifiable | Modifiable | Biomarkers |
|---|--|---|--|
| Age >65 years Previous major bleeding Severe renal impairment (on dialysis or renal transplant) Severe hepatic dysfunction (cirrhosis) Malignancy Genetic factors (e.g., CYP 2C9 polymorphisms) Previous stroke, small-vessel disease, etc. Diabetes mellitus Cognitive impairment/dementia | Extreme frailty ± excessive risk of falls ^a Anaemia Reduced platelet count or function Renal impairment with CrCl <60 mL/min VKA management strategy ^b | Hypertension/elevate SBP Concomitant antiplatelet/NSAID Excessive alcohol intake Non-adherence to OAC Hazardous hobbies / occupations Bridging therapy with heparin INR control (target 2.0–3.0), target TTR >70% ^c Appropriate choice of OAC and correct dosing ^d | GDF-15 Cystatin C / CKD-EPI cTnT-hs Von Willebrand factor (+ other coagulation markers) |

^aWalking aids, appropriate footwear; home review to remove trip hazards; neurological assessment where appropriate. ^bIncreased INR monitoring, dedicated OAC clinics, self-monitoring/self-management, educational/behavioural interventions. ^cFor patients receiving VKA treatment. ^dDose adaptation based on patient's age, body weight, and serum creatinine level.

2021 European Heart Rhythm Association Practical Guide on the Use of Non-Vitamin K Antagonist Oral Anticoagulants in Patients with Atrial Fibrillation

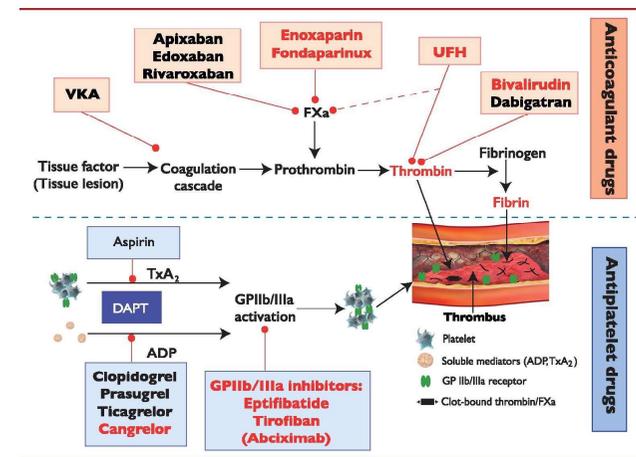
Jan Steffel, Ronan Collins, Matthias Antz, Pieter Cornu, Lien Desteghe, Karl Georg Haeusler, Jonas Oldgren, Holger Reinecke, Vanessa Roldan-Schilling, Nigel Rowell, Peter Sinnaeve, Thomas Vanassche, Tatjana Potpara, A. John Camm, and Hein Heidbüchel

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Pharmacological targets for antithrombotic treatments



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NOACs vs. VKA in dual vs. triple therapy (1)

- Four dedicated prospective RCTs have addressed the issue of using a NOAC or VKA in a variety of combinations with antiplatelet agents to reduce bleeding events after PCI and/or an ACS in patients with AF.
- In essence, these trials focused on bleeding as the primary endpoint, with coronary events and stroke as important secondary outcomes.
- On aggregate, these studies showed that dual therapy with a NOAC plus a P2Y12 inhibitor reduced the risk of bleeding compared to triple therapy with VKA, aspirin and a P2Y12 inhibitor (mostly clopidogrel).
- This bleeding risk reduction appeared to be driven by both receiving a NOAC instead of VKA as well as by omitting aspirin, and this benefit was also observed in medically managed ACS/PCI patients with AF

NOAC dosing in AF patients post-ACS/PCI

| | Standard dose | Comments/dose reduction |
|----------------------------|--------------------------|---|
| Apixaban ²⁴⁴ | 5 mg BID | Dose reduction as for SPAF |
| Dabigatran ²⁴⁷ | 150 mg BID or 110 mg BID | 110mg as for SPAF ⁴⁰³ |
| Edoxaban ²⁴⁵ | 60 mg QD | Dose reduction as for SPAF |
| Rivaroxaban ²⁴⁶ | 15 mg QD | Dose reduction to 10 mg QD if CrCl 30–49 mL/min |

In addition to single/dual antiplatelet therapy, where applicable.
BID, twice daily; CrCl, creatinine clearance; QD, once daily; SPAF, stroke prevention in atrial fibrillation.

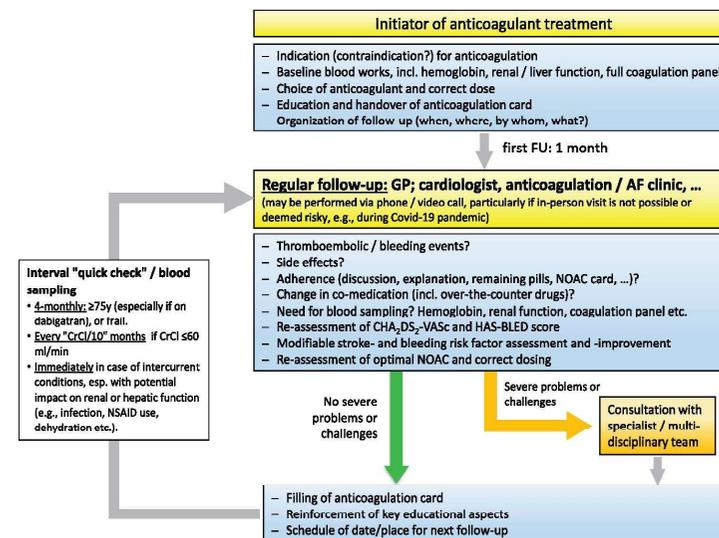
Choice of anticoagulant therapy and initiation

Indication for anticoagulation and choice between VKA and NOAC

- After the indication for OAC is established, **NOACs are preferred** over VKAs in all NOAC-eligible AF patients
- When starting a NOAC, **knowledge of current kidney and liver function** is required as all NOACs are eliminated to some extent via the kidneys, and renal function affects NOAC dosing. Importantly, kidney function should be assessed using the Cockcroft–Gault formula as it was used in the four pivotal phase III trial. Indeed, use of other formulas including ‘Modification of Diet in Renal Disease’ (MDRD) and ‘Chronic Kidney Disease—Epidemiology Collaboration’ (CKD-EPI) may overestimate kidney function particularly in older patients and in those with low body weights.
- A **baseline haematological profile** should be obtained for reference during future follow-up.
- Bleeding risk**, as estimated using the HAS-BLED score, is **not in itself a reason to deny OAC** to AF patients at risk of stroke or reduce the dose of the NOAC. Instead, particularly patients at high bleeding risk (e.g. HAS-BLED ≥ 3) should have their modifiable bleeding risk factors identified and addressed, and should be scheduled for an earlier and more frequent clinical follow-up.
- Similarly, **frailty, cognitive decline and risk of falling** should not generally be a reason not to anticoagulate patients. Care needs to be taken to minimize the risk of falling and to ensure optimal compliance and adherence. This topic is dealt with in detail in the ‘NOACs in advanced age and frailty’ section.

Europace, Volume 23, Issue 10, October 2021, Pages 1612–1676

Initiation and structured follow-up of patients on NOACs

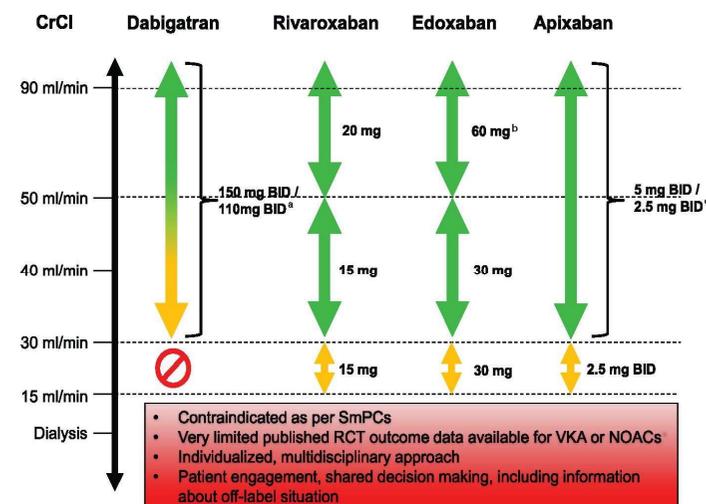


Europace, Volume 23, Issue 10, October 2021, Pages 1612–1676

Absorption and metabolism of the different NOACs

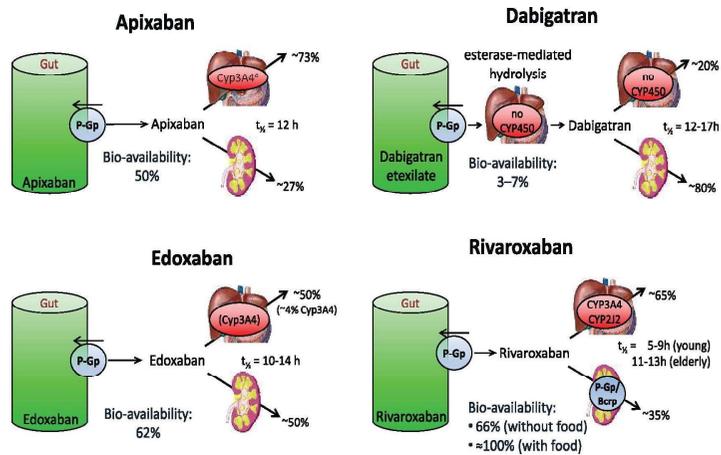
| | Dabigatran ^{106,376} | Apixaban ⁵¹⁷ | Edoxaban ⁵¹⁸ | Rivaroxaban ^{519,520} |
|--|---------------------------------------|---|--|---|
| Bioavailability | 3–7% | 50% | 62% | 15 mg/20 mg: 66% without food, 100% with food |
| Prodrug | Yes | No | No | No |
| Clearance non-renal/renal of absorbed dose | 20%/80% | 73%/27% | 50%/50% | 65%/35% |
| Plasma protein binding | 35% | 87% | 55% | 95% |
| Dialysability | 50–60% (In part dialysable) | 14% (Not dialysable) | NA (Not dialysable) | NA (Not dialysable) |
| Metabolism | Glucuronic acid conjugation | CYP3A4 (25%), CYP1A2, CYP2J2, CYP2C8, CYP2C9, CYP2C19 | CYP3A4 (<4% of elimination) | CYP2A4 (18%) ⁵¹⁹ , CYP2J2 |
| Absorption with food | No effect | No effect | 6–22% more; minimal effect on exposure | +39% more (see above) |
| Absorption with H2B/PPI | –12% to 30% (not clinically relevant) | No effect | No effect | No effect |
| Time to peak levels (h) | 3 | 3 | 2–4 | 2–4 |
| Elimination half-life (h) | 12–17 | 12 | 10–14 | 5–9 (young) 11–13 h (elderly) |

Use of NOACs according to renal function



Europace, Volume 23, Issue 10, October 2021, Pages 1612–1676

Absorption and metabolism of the different NOACs



Dosing Errors

Even in settings with optimal patient education dosing errors are common in daily practice, and patients need to be informed on what to do in such cases.

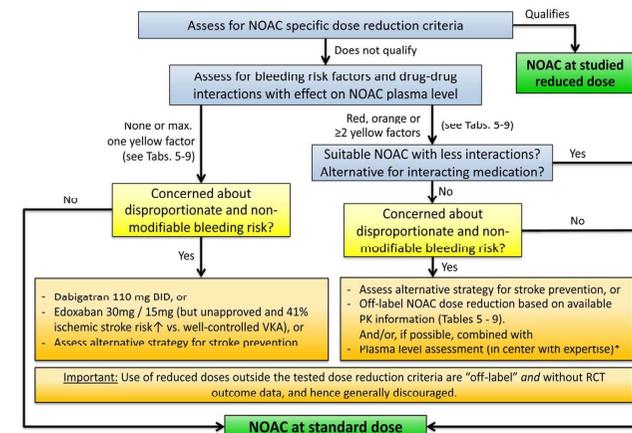
Missed dose

- A **forgotten dose** may be taken until half of the dosing interval has passed. Hence, for NOACs with a **twice daily (BID) dosing regimen** (i.e., intake every 12 h), a forgotten full dose can be taken up **until 6 h after the scheduled intake**. For NOACs with a **once daily (QD) dosing regimen**, a forgotten dose can be taken up **until 12 h after the scheduled intake**. After these time points, the dose should be skipped, and the next scheduled dose should be taken.

Double dose

- For NOACs with a **BID dosing regimen**, the next planned dose (i.e. after 12 h) **may be skipped**, with the regular BID dosing regimen restarted 24 h after the double dose intake.
- For NOACs with a **QD dosing regimen**, the patient should continue **the normal dosing regimen**, i.e. without skipping the next daily dose.

NOAC selection based on drug–drug interactions and/or risk of bleeding.



Dosing Errors

Uncertainty about dose intake

- For NOACs with a **BID dosing regimen**, it is generally advisable to **not take another tablet/capsule**, but to continue with the regular dose regimen, i.e. starting with the next dose at the 12 h interval.
- For NOACs with a **QD dosing regimen**, when **thromboembolic risk is high** ($CHA_2DS_2-VASc \geq 3$), it may generally be advisable to **take another tablet 6–8 h after the original (uncertain) intake** and then continue the planned dose regimen. In case the thromboembolic risk is low ($CHA_2DS_2-VASc \leq 2$) we advise to **wait until the next scheduled dose**.

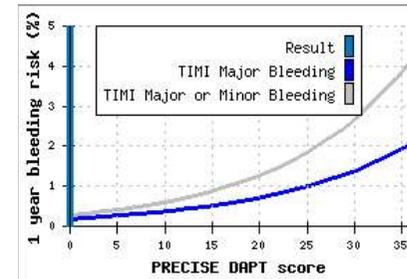
Choice of P2Y12 inhibitor

- In the 2020 ESC AF guidelines, the use of ticagrelor or prasugrel as part of a triple therapy regimen is discouraged.
- Ticagrelor increases bleeding risk in patients on dual therapy when compared to clopidogrel (RE-DUAL PCI trial).
- Although only few patients have been included with a P2Y12-inhibitor other than clopidogrel into the above-mentioned RCTs, the benefit in terms of reduced bleeding risk with NOAC-based dual therapy compared to VKA-based triple therapy appears to be maintained regardless of the type of P2Y12 inhibitor
- In post-ACS patients at high coronary thrombotic risk and low bleeding risk in whom otherwise a VKA- or NOAC-based triple therapy would be warranted, dual therapy with a NOAC plus ticagrelor could be considered instead. Further data, including dedicated RCTs, are warranted in this area. Indeed, up to 40% of patients on clopidogrel may reach insufficient platelet inhibition.



PREdicting bleeding Complications In patients undergoing Stent implantation and subsEquent Dual Anti Platelet Therapy

- Haemoglobin Values collected in close proximity to the index procedure (es. 12 g/dL)
- Age (years)
- White blood cells Values collected in close proximity to the index procedure
- Creatinine Clearance (ml/min) Values collected in close proximity to the index procedure (es. 75 ml/min)
- Prior Bleeding History of spontaneous bleeding requiring medical attention



Europace, Volume 23, Issue 10, October 2021, Pages 1612–1676

| | PRECISE-DAPT-Score | DAPT-Score |
|------------------|--|---|
| Score-Berechnung | <p>Score-Nomogramm</p> <p>0 2 4 6 8 10 12 14 16 18 20 22 24 26 28 30</p> <p>Punkte</p> <p>Hämoglobin (g/dl)</p> <p>≥12,0 11,5 11,0 10,5 ≤10,0</p> <p>Leukozyten (x 10⁹ Zellen/μl)</p> <p>≤5 8 10 12 14 16 18 ≥20</p> <p>Alter (Jahre)</p> <p>≥100 80 60 40 20 0</p> <p>Kreatinin-Clearance (ml/min)</p> <p>Nein Ja</p> <p>Frühere Blutung</p> | <p>Alter</p> <p>≥ 75 -2 Punkte</p> <p>65 bis < 75 -1 Punkt</p> <p>< 65 0 Punkte</p> <p>Rauchen +1 Punkt</p> <p>Diabetes mellitus +1 Punkt</p> <p>ACS bei Präsentation +1 Punkt</p> <p>Vorausgegangen ACS oder PCI +1 Punkt</p> <p>Paclitaxelfreisetzender Stent +1 Punkt</p> <p>Stentdiameter < 3 mm +1 Punkt</p> <p>Herzinsuffizienz oder LVEF < 30% +2 Punkte</p> <p>Stent in Venenbypass +2 Punkte</p> |
| Score-Bereich | 0 bis 100 Punkte | -2 bis 10 Punkte |
| Cut-off-Werte | Score ≥ 25 → kurze DAPT Score < 25 → Standard-/lange DAPT | Score ≥ 2 → lange DAPT Score < 2 → Standard-DAPT |
| Kalkulator | www.precisedaptscore.com | www.daptstudy.org |

Abbildung 1: Risikoscores, validiert für die Dauer der dualen antithrombozytären Therapie (DAPT; ACS: akutes Koronarsyndrom, PCI: perkutane Koronarintervention, LVEF: linksventrikuläre Ejektionsfraktion): Für die Berechnung des PRECISE-DAPT-Scores wird das Score-Nomogramm verwendet. Jeder Wert der 5 Variablen wird markiert, von der Markierung wird eine vertikale Linie zum Nomogramm gezogen und der entsprechende Wert bestimmt. Die Summe aller 5 Variablen ergibt den PRECISE-DAPT-Score (modifiziert nach Valgimigli et al. [2]).

Risk criteria for extended treatment with a second antithrombotic agent

| High thrombotic risk (Class IIa) | Moderate thrombotic risk (Class IIb) |
|--|---|
| Complex CAD and at least 1 criterion | Non-complex CAD and at least 1 criterion |
| Risk enhancers | |
| Diabetes mellitus requiring medication | Diabetes mellitus requiring medication |
| History of recurrent MI | History of recurrent MI |
| Any multivessel CAD | Polyvascular disease (CAD plus PAD) |
| Polyvascular disease (CAD plus PAD) | CKD with eGFR 15–59 mL/min/1.73 m ² |
| Premature (<45 years) or accelerated (new lesion within a 2-year time frame) CAD | |
| Concomitant systemic inflammatory disease (e.g. human immunodeficiency virus, systemic lupus erythematosus, chronic arthritis) | |
| CKD with eGFR 15–59 mL/min/1.73 m ² | |
| Technical aspects | |
| At least 3 stents implanted | |
| At least 3 lesions treated | |
| Total stent length >60 mm | |
| History of complex revascularization (left main, bifurcation stenting with ≥2 stents implanted, chronic total occlusion, stenting of last patent vessel) | |
| History of stent thrombosis on antiplatelet treatment | |

Suggested strategies to reduce bleeding risk related to percutaneous coronary intervention

- Anticoagulant doses adjusted to body weight and renal function, especially in women and older patients

- Radial artery approach as default vascular access

- Proton pump inhibitors in patients on DAPT at higher-than-average risk of gastrointestinal bleeds (i.e. history of gastrointestinal ulcer/haemorrhage, anticoagulant therapy, chronic non-steroidal anti-inflammatory drugs/corticosteroid use, or two or more of:

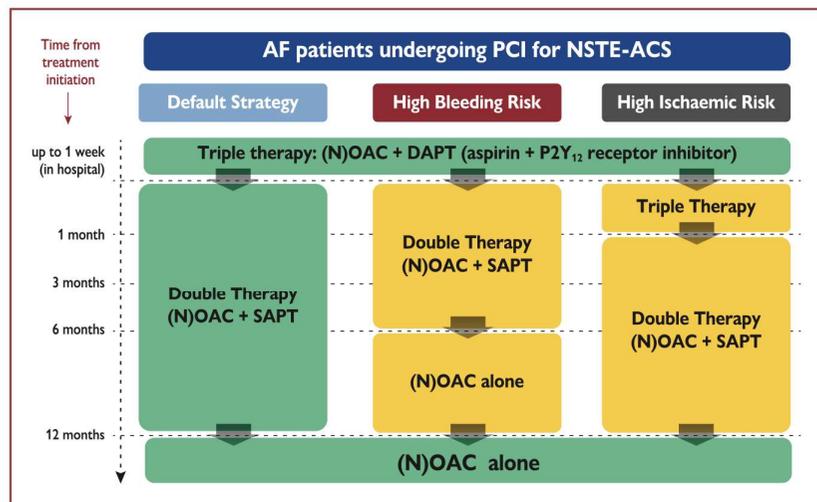
- Age ≥ 65 years
- Dyspepsia
- Gastro-oesophageal reflux disease
- *Helicobacter pylori* infection
- Chronic alcohol use

- In patients on OAC

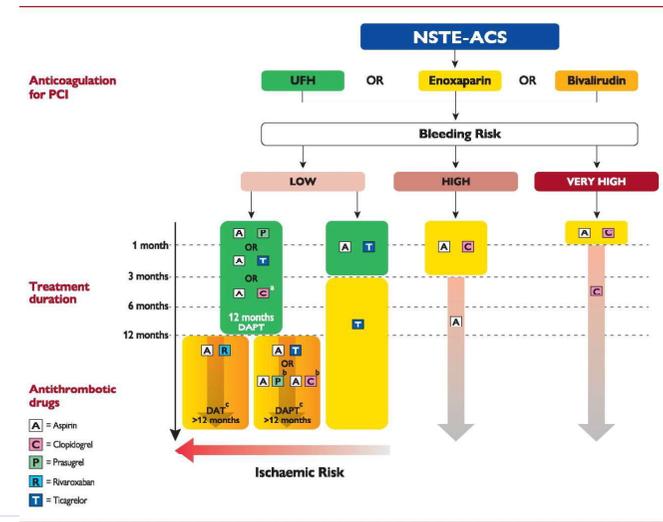
- PCI performed without interruption of VKAs or NOACs
 - In patients on VKAs, do not administer UFH if INR >2.5
 - In patients on NOACs, regardless of the timing of the last administration of NOACs, add low-dose parenteral anticoagulation (e.g. enoxaparin 0.5 mg/kg i.v. or UFH 60 IU/kg)

- Aspirin is indicated but avoid pre-treatment with P2Y₁₂ receptor inhibitors

Algorithm for antithrombotic therapy in non-ST-segment elevation acute coronary syndrome patients with atrial fibrillation undergoing percutaneous coronary intervention or medical management.



Algorithm for antithrombotic therapy in non-ST-segment elevation acute coronary syndrome patients without atrial fibrillation undergoing percutaneous coronary intervention.



Eur Heart J, Volume 42, Issue 14, 7 April 2021, Pages 1289–1367

Duration of triple therapy after ACS/PCI

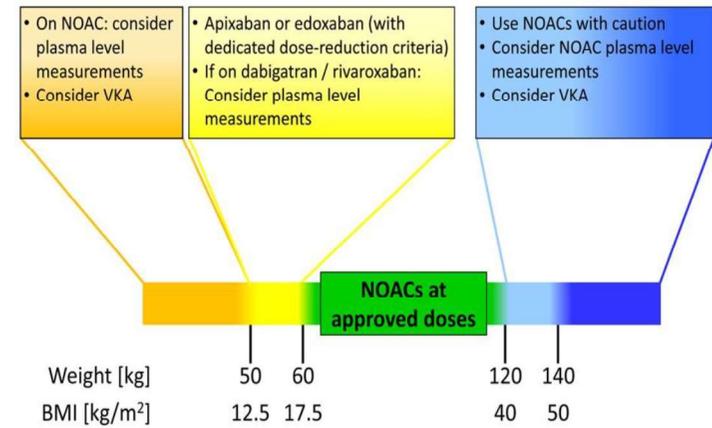
- According to the current 2020 ESC guidelines for AF as well as for non-ST-elevation acute coronary syndrome (NSTEMI-ACS), a short course of triple therapy is recommended for up to 1 week in all patients with AF undergoing PCI
- In medically managed NSTEMI-ACS patients, combination of a NOAC with only a single antiplatelet agent (preferably clopidogrel) is recommended from the event onwards
- However, the time frame of inclusion for the four aforementioned NOAC RCTs ranged from several hours after PCI up to >10 days. As such, a selection bias towards lower-risk patients cannot be excluded; furthermore, a variable course of triple therapy may have been given to a substantial number of patients subsequently randomized to NOAC-based dual therapy
- Finally, although bleeding events were consistently reduced across the four NOAC trials by NOAC-based dual therapy this did not translate into a reduction in all-cause mortality (as compared to VKA-based triple therapy). Therefore, a low threshold for prolonging triple therapy with DAPT and a NOAC up to 30 days may be advisable in patients with a high atherothrombotic risk, including those after a complex PCI or with a history of stent thrombosis. In contrast, continuation of triple therapy beyond 30 days rarely seems warranted.

NOAC use in frail patients

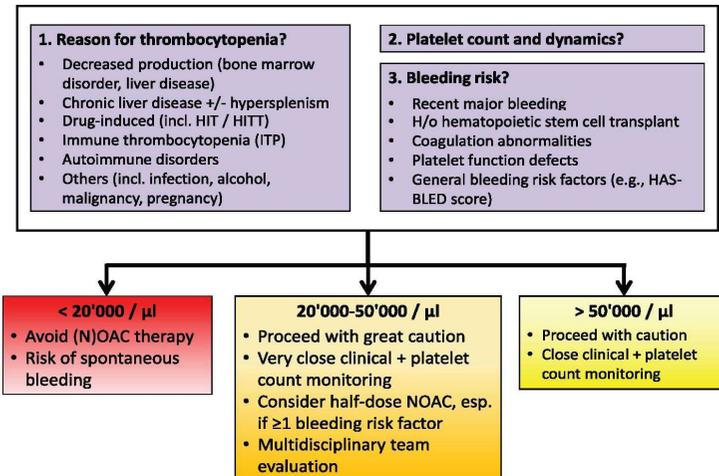
| | |
|----------------------------|--|
| Very Fit | People who are robust, active, energetic and motivated. These people commonly exercise regularly. They are among the fittest for their age. |
| Well | People who have no active disease symptoms but are less fit than category 1. Often, they exercise or are very active occasionally, e.g. seasonally. |
| Managing Well | People whose medical problems are well controlled but are not regularly active beyond routine walking. |
| Vulnerable | While not dependent on others for daily help, often symptoms limit activities. A common complaint is being "slowed up", and/or being tired during the day. |
| Mildly Frail | These people often have more evident slowing and need help in high order with ADLs (finances, transportation, heavy housework, medications). Typically, mild frailty progressively impairs shopping and walking outside alone, meal preparation and housework. |
| Moderately Frail | People need help with all outside activities and with keeping house. Inside, they often have problems with stairs and need help with bathing and might need minimal assistance (cuing, standby) with dressing. |
| Severely Frail | Completely dependent for personal care, from whatever cause (physical or cognitive). Even so, they seem stable and not at high risk of dying (within ~ 6 months). |
| Very Severely Frail | Completely dependent, approaching the end of life. Typically, they could not recover even from a minor illness. |
| Terminally ill | Approaching the end of life. This category applies to people with a life expectancy <6 months, who are not otherwise evidently frail. |

The 'Canadian Study of Health and Aging' (CHSA) Clinical Frailty Scale, based on comprehensive geriatric assessment including structured interview

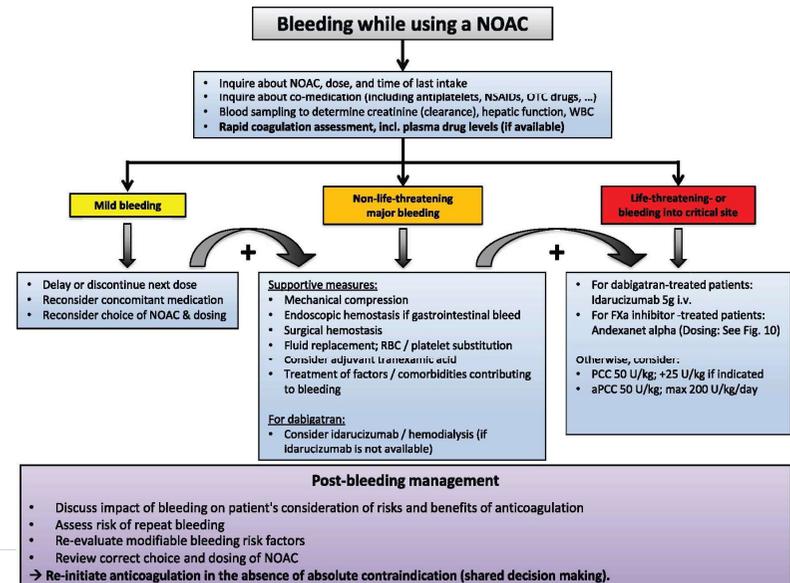
NOACs in under- and overweight patients



NOACs in patients with thrombocytopenia



Management of bleeding in patients taking NOACs



Treatment of patients with chronic coronary syndrome

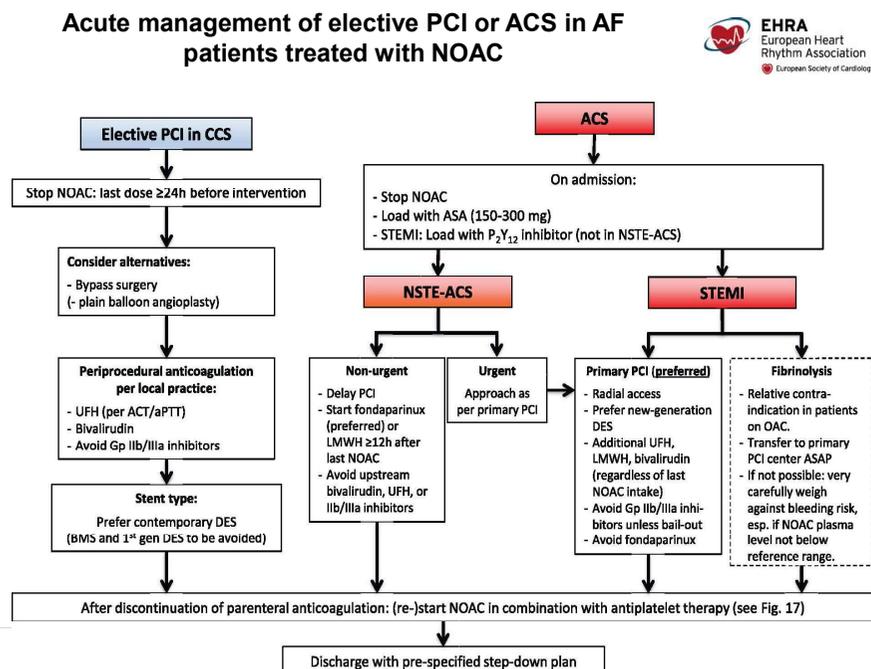
- The Japanese multi-centre, open-label 'Atrial Fibrillation and Ischemic Events with Rivaroxaban in Patients with Stable Coronary Artery Disease' (AFIRE) trial demonstrated that **continuing rivaroxaban 15 mg QD monotherapy** beyond 1 year after a revascularization procedure in AF patients **not only decreased the risk of ISTH bleeding** (primary safety outcome) but also demonstrated non-inferiority for the **primary composite endpoint of cardiovascular events** (stroke, systemic embolism, MI, unstable angina requiring revascularization) or death from any cause compared with the combination of rivaroxaban and antiplatelet therapy.
- Indeed, the trial was stopped prematurely due to an increased mortality in the combination therapy arm. Although it is formally unclear if these results translate to other NOACs, other doses, and other populations, these data suggest that most AF patients with chronic CAD should be transitioned to NOAC monotherapy without an antiplatelet agent as recommended in current ESC AF guidelines

Europace, Volume 23, Issue 10, October 2021, Pages 1612–1676

Scenario 1: coronary interventions in atrial fibrillation patients on non-vitamin K antagonist oral anticoagulants (1)

- Performing a PCI (scheduled or not) under NOAC is different than under VKA for several reasons, and various aspects and uncertainties need to be taken into consideration, including:
 - timepoint of the last dose, adherence, and renal function;
 - variability in renal function in an acute setting;
 - singular factor II or Xa blockade vs. multifactor antagonism;
 - uncertainty about the extent of anticoagulation in the absence of established tests, and hence
 - uncertainty about stacking of additional periprocedural anticoagulants, etc.
- Temporary discontinuation of the short-acting NOACs may allow for safe initiation of antiplatelet therapy and standard local anticoagulation practices peri-procedurally.
- In contrast, NOACs should be continued in non-invasively managed ACS patients.

Europace, Volume 23, Issue 10, October 2021, Pages 1612–1676



Europace, Volume 23, Issue 10, October 2021, Pages 1612–1676

Scenario 1: coronary interventions in atrial fibrillation patients on non-vitamin K antagonist oral anticoagulants (2)

- New-generation drug-eluting stents are preferred to shorten exposure to dual or triple therapy after the procedure but also to avoid the need for repeat interventions.
- Sole balloon angioplasty or bypass surgery should always be considered as an alternative in patients in need for chronic anticoagulation since they can reduce the need for long-term dual or triple therapy.
- There is no longer a reason to opt for a bare metal stent as a strategy to reduce DAPT duration.

Europace, Volume 23, Issue 10, October 2021, Pages 1612–1676

Scenario 2: management of the patient with a recent acute coronary syndrome (<1 year) who develops new-onset atrial fibrillation

- ACS guidelines recommended DAPT for up to 1 year after the acute event in patients without indication for OAC, and high-risk patients might require an even longer DAPT duration.
- In high bleeding-risk ACS patients, however, current ESC guidelines allow for shorter DAPT durations (3–6 months).
- If AF develops during the first year after an ACS and there is an indication for anticoagulation, a NOAC should be started and the need for continuing DAPT should be carefully weighed against the increased bleeding risk. Beyond 1 month after the event, aspirin can be stopped in the majority of such patients as discussed above.

Europace, Volume 23, Issue 10, October 2021, Pages 1612–1676

Treatment of left ventricular thrombus after myocardial infarction in patients with atrial fibrillation

- In the absence of randomized studies, it **remains uncertain** whether a NOAC is effective in the treatment of left ventricular thrombi complicating a large infarction. One observational study suggests that NOACs were associated with a **higher incidence** of thromboembolic events compared to VKA in (mostly non-AF) patients with a left ventricular thrombus, while others showed a similar rate of thrombus resolution.
- Although residual confounding can never be excluded in these settings, **VKA should be viewed as standard of care** for the treatment of patients with LV thrombus until more data are available.
- **Only in very special situations** (e.g. no VKA monitoring possible, no stable INR despite maximal efforts, etc.) NOACs may be evaluated after clear communication and consent from the patient about the lack of data and the off-label situation.

Europace, Volume 23, Issue 10, October 2021, Pages 1612–1676

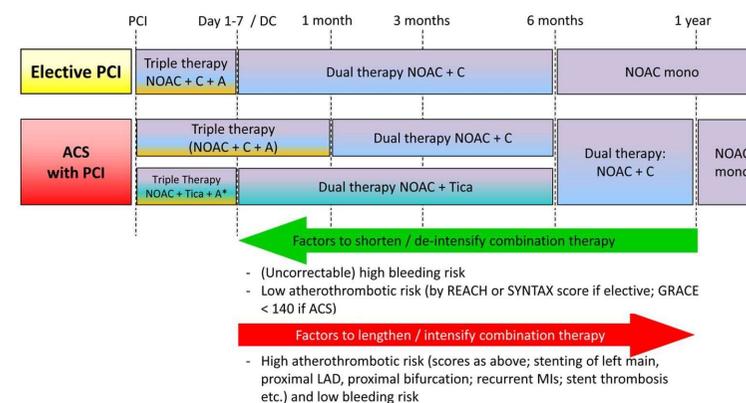
Scenario 3: a chronic coronary syndrome patient (acute coronary syndrome ≥1 year ago) develops atrial fibrillation

- Patients with a CCS **developing AF** should receive anticoagulation, depending on their CHA₂DS₂-VASc score (which per definition will be ≥1).
- A **NOAC without any antiplatelet agent** appears to be the preferred strategy for these patients as discussed above, based on the results of the four landmark NOAC trials (which included up to 15–20% of patients with a prior MI) and the ‘Atrial Fibrillation and Ischaemic Events with Rivaroxaban in Patients with Stable Coronary Artery Disease’ (AFIRE) trial.
- An additional antiplatelet agent should only be considered in individual patients with a **very high ischaemic and low bleeding risk**.

Europace, Volume 23, Issue 10, October 2021, Pages 1612–1676

Summary

Anticoagulation therapy after elective PCI or ACS in patients with AF



In all patients:

- Avoid use of BMS / first generation DES
- Use PPI if on triple / dual therapy
- Minimize bleeding risk by assessing and treating modifiable bleeding risk factors (e.g., hypertension, etc.)
- Close follow-up; check for signs of (occult) bleeding

Europace, Volume 23, Issue 10, October 2021, Pages 1612–1676

Thank you for your attention
