

# ELIQUIS<sup>®</sup> (apixaban) prescriber checklist

This checklist applies to the ELIQUIS indication for the treatment of DVT and PE, and for the prevention of recurrent DVT and PE in adults. It is not to be used in relation to other ELIQUIS indications or until the prescribing decision has been made to consider the patient for ELIQUIS. This checklist is intended for healthcare professional use only and should not be used by or given to patients.

Patient name:

MRN number:

Date of birth:

Signed by healthcare professional:

Print name:

Date:

Signed by patient / carer:

Print name:

Date:

ELIQUIS (apixaban): an oral, direct factor Xa inhibitor indicated for the treatment of deep vein thrombosis (DVT) and pulmonary embolism (PE), and prevention of recurrent DVT and PE in adults.<sup>1</sup>

DVT = Deep Vein Thrombosis PE = Pulmonary Embolism

**Prescribing information and Adverse Event reporting information can be found at the end of this document. This promotional checklist has been produced by the BMS / Pfizer Alliance.**

# ELIQUIS® prescriber checklist\*

The table below provides an example checklist that may be used to assess a patient's suitability prior to initiating treatment with ELIQUIS. Please note that this is a sample of potential assessment questions that could be used and is not an exhaustive list. Please refer to the SmPC before prescribing.<sup>1</sup>

This entire document is intended for healthcare professionals only and should not be given to patients.

Have you reviewed the following in this patient?	
<b>Diagnosis and risk assessment</b>	<b>Assessed</b>
Patient has VTE (DVT or PE) or has had a recurrent VTE	<input type="checkbox"/>
Patient's risk factors for VTE (e.g. known thrombophilias, over 60 years old, obesity (BMI over 30 kg/m <sup>2</sup> ))	<input type="checkbox"/>
Individual risk of bleeding	<input type="checkbox"/>
<b>Renal function</b>	
Prior to initiating ELIQUIS, renal function testing should be performed. ELIQUIS is NOT recommended in patients with CrCl <15 ml/min or in patients undergoing dialysis. Use with caution in patients with CrCl 15–29 ml/min due to an increased risk of bleeding.	<input type="checkbox"/>
<b>Hepatic function</b>	
ELIQUIS is contraindicated in patients with hepatic disease associated with coagulopathy and clinically relevant bleeding risk. ELIQUIS is <b>NOT</b> recommended in patients with severe hepatic impairment. It should be used with caution in patients with mild or moderate hepatic impairment (Child Pugh A or B). Patients with elevated liver enzymes alanine aminotransferase (ALT) / aspartate aminotransferase (AST) >2 x ULN or total bilirubin ≥ 1.5 x ULN were excluded in clinical trials. Therefore ELIQUIS should be used with caution in this population. Prior to initiating ELIQUIS, liver function testing should be performed.	<input type="checkbox"/>
<b>Special warnings and precautions for use</b>	
Any relevant "special warnings and precautions for use" see SmPC for further information. Patients with active cancer can be at high risk of both venous thromboembolism and bleeding events. When apixaban is considered for DVT or PE treatment in cancer patients, a careful assessment of the benefits against the risks should be made.	<input type="checkbox"/>
Please note ELIQUIS is <b>NOT</b> recommended in haemodynamically unstable PE patients or patients who require thrombolysis or pulmonary embolectomy. ELIQUIS is <b>NOT</b> recommended in patients with prosthetic heart valves as safety and efficacy have not been studied. ELIQUIS is <b>NOT</b> recommended in patients with a history of thrombosis who are diagnosed with antiphospholipid syndrome. Refer to SmPC for full prescribing information.	<input type="checkbox"/>
<b>Contraindications to ELIQUIS</b>	
Concomitant treatment with any other anticoagulant agent <b>except</b> under specific circumstances of switching anticoagulant therapy to or from ELIQUIS or when unfractionated heparin is given at doses necessary to maintain an open central venous or arterial catheter, or when unfractionated heparin is given during catheter ablation for AF	<input type="checkbox"/>
Hypersensitivity to the active substance or any of the excipients (ELIQUIS contains lactose – patients with rare hereditary problems of galactose intolerance, total lactase deficiency, or glucose-galactose malabsorption should <b>NOT</b> take ELIQUIS)	<input type="checkbox"/>
Active clinically significant bleeding	<input type="checkbox"/>
Hepatic disease associated with coagulopathy and clinically relevant bleeding risk	<input type="checkbox"/>
Lesion or condition if considered a significant risk factor for major bleeding (see SmPC for list of examples)	<input type="checkbox"/>
<b>Concomitant medications</b>	
Use of strong inhibitors of both CYP3A4 and P-gp (ELIQUIS is <b>NOT</b> recommended)	<input type="checkbox"/>
Use of strong inducers of both CYP3A4 and P-gp (for the <u>treatment of VTE</u> , ELIQUIS should <b>NOT</b> be used since efficacy may be compromised. For the prevention of recurrent VTE, ELIQUIS should be used with caution).	<input type="checkbox"/>
Use of other anticoagulants (See contraindications section above)	<input type="checkbox"/>
Use with caution with SSRIs / SNRIs, NSAIDs, aspirin, and / or P2Y <sub>12</sub> inhibitors due to increased risk of bleeding	<input type="checkbox"/>
Use of other platelet aggregation inhibitors (such as GPIIb / IIIa receptor antagonists, dipyridamole, dextran or sulfinpyrazone) or thrombolytic agents increase the bleeding risk (ELIQUIS is <b>NOT</b> recommended)	<input type="checkbox"/>
<b>Pregnancy and breastfeeding</b>	
Avoid use of ELIQUIS during pregnancy or when breastfeeding	<input type="checkbox"/>
<b>Dose of ELIQUIS prescribed</b>	
The duration of overall therapy should be individualised after careful assessment of the treatment benefit against the risk for bleeding	
<ul style="list-style-type: none"> <li>● <b>Treatment of VTE:</b> 10 mg BD for the first 7 days followed by 5 mg BD. As per available medical guidelines, short duration of treatment (at least 3 months) should be based on transient risk factors (e.g., recent surgery, trauma, immobilisation) <b>OR</b></li> <li>● <b>Prevention of recurrent VTE:</b> 2.5 mg BD following completion of 6 months of anticoagulant treatment for DVT or PE</li> </ul>	<input type="checkbox"/> <b>OR</b> <input type="checkbox"/>

\* This checklist applies to the ELIQUIS indication for the treatment of DVT and PE, and for the prevention of recurrent DVT and PE in adults. It is not to be used in relation to other ELIQUIS indications.

# Patient discussion checklist\*

The table below provides an example of a checklist that may be used to guide a discussion with patients starting ELIQUIS®. Please note that this is a sample of potential assessment questions that could be used and is not an exhaustive list. Please refer to the SmPC before prescribing.<sup>1</sup>

This entire document is intended for healthcare professionals only and should not be given to patients.

Have you discussed the following points with your patient?		
Diagnosis and VTE risk	Informing healthcare professionals	
<ul style="list-style-type: none"> <li>■ Diagnosis of VTE</li> <li>■ The patient's individual risk factors for VTE</li> </ul>	<input type="checkbox"/> As with all anticoagulants, patients should inform the following HCPs that they are taking ELIQUIS: <ul style="list-style-type: none"> <li>■ Doctor/Nurse – inform before any additional medication or surgery/procedure is required</li> <li>■ Dentist – needs to be informed because there is an increased risk of bleeding if dental treatment is needed</li> <li>■ Pharmacist – so they can advise about medicines suitable to take with anticoagulants</li> <li>■ All HCPs – ensure patient has a patient alert card and emphasise the importance of carrying at all times and producing when seeing a healthcare professional</li> </ul>	<input type="checkbox"/>
<ul style="list-style-type: none"> <li>■ The role of anticoagulants in the treatment and/or prevention of recurrent VTE</li> </ul>		
Dose of ELIQUIS prescribed		
<ul style="list-style-type: none"> <li>■ Advise patient on the dose of ELIQUIS they have been prescribed</li> </ul>	<input type="checkbox"/>	
How to take ELIQUIS		
<ul style="list-style-type: none"> <li>■ ELIQUIS should be swallowed with water, with or without food and taken twice a day as directed</li> </ul>	<input type="checkbox"/>	
How long to take ELIQUIS for	Surgery and invasive procedures	
<ul style="list-style-type: none"> <li>■ Continue until told by doctor to stop</li> <li>■ Discuss ways of remembering to take twice daily</li> <li>■ Discuss the importance of treatment compliance</li> <li>■ Arrange appropriate follow-up appointments</li> <li>■ Ensure patient knows how to order repeat prescription</li> </ul>	<ul style="list-style-type: none"> <li>■ Patient should inform surgeon that they are taking ELIQUIS as it is recommended that ELIQUIS is discontinued before and during surgery (See SmPC for further information)</li> </ul>	<input type="checkbox"/>
	Drug interactions	
	<ul style="list-style-type: none"> <li>■ Check if the patient is taking any drugs other than those recorded in their notes, e.g. OTC preparations, herbal remedies, other medicines</li> <li>■ Advise patients to consult a pharmacist (or other healthcare professional) before taking any other medication (including OTC preparations, herbal remedies, other medicines)</li> </ul>	<input type="checkbox"/>
When to take ELIQUIS	Side effects	
<ul style="list-style-type: none"> <li>■ At roughly the same times each day and 12 hours apart</li> </ul>	<ul style="list-style-type: none"> <li>■ Most relate to bruising and bleeding. Discuss signs and symptoms of bleeding</li> <li>■ Discuss actions in case of minor and major bleeding episodes. In case of bleeding event that does not stop on its own or is serious, seek medical attention immediately</li> <li>■ Advise patient to read the patient information leaflet found inside their ELIQUIS medicine pack and inform their doctor, nurse or pharmacist about any side effects they experience, even if they are not listed in the patient information leaflet</li> </ul>	<input type="checkbox"/>
What to do if a dose is missed?		
<ul style="list-style-type: none"> <li>■ If a dose is missed, take ELIQUIS immediately and then continue with twice-daily intake as before</li> </ul>	<input type="checkbox"/>	
What to do if an extra dose is taken accidentally?	ELIQUIS patient support	
<ul style="list-style-type: none"> <li>■ Contact pharmacist, doctor or nurse straight away</li> </ul>	<ul style="list-style-type: none"> <li>■ Provide patients with ELIQUIS patient information and advise to also read the Patient Information Leaflet contained within their pack of ELIQUIS and complete the patient alert card</li> </ul>	<input type="checkbox"/>
Anticoagulation monitoring	ELIQUIS patient alert card	
<ul style="list-style-type: none"> <li>■ Advise patients who are accustomed to INR monitoring with warfarin, that no INR monitoring is required with ELIQUIS</li> </ul>	<ul style="list-style-type: none"> <li>■ Emphasise the importance of carrying this at all times and producing when seeing a healthcare professional</li> </ul>	<input type="checkbox"/>
Food interactions	Contact numbers	
<ul style="list-style-type: none"> <li>■ Advise patients that ELIQUIS should be swallowed with water, with or without food, and there are no known food interactions</li> </ul>	<ul style="list-style-type: none"> <li>■ Check that the patient has the appropriate contact numbers to contact the relevant healthcare professional(s) should they have any questions/concerns</li> </ul>	<input type="checkbox"/>

\* This checklist applies to the ELIQUIS indication for the treatment of DVT and PE, and for the prevention of recurrent DVT and PE in adults. It is not to be used in relation to other ELIQUIS indications.

BD = Twice Daily CrCL = Creatinine Clearance DVT = Deep Vein Thrombosis INR = International Normalised Ratio NSAIDs = Non Steroidal Anti-Inflammatory Drugs OTC = Over The Counter PE = Pulmonary Embolism P-gp = P-glycoprotein SmPC = Summary of Product Characteristics SNRI = Serotonin Norepinephrine Reuptake Inhibitor SSRI = Selective Serotonin Reuptake Inhibitor ULN = Upper Limit of Normal VTE = Venous Thromboembolic Events

## References:

1. ELIQUIS® (apixaban) Summary of Product Characteristics. Available at <http://www.medicines.ie>

# ELIQUIS® (apixaban) PRESCRIBING INFORMATION

## Ireland

Consult Summary of Product Characteristics (SmPC) before prescribing

**PRESENTATION:** Film-coated tablets; 5 mg and 2.5 mg apixaban.

**INDICATION (SPC section 4.1):** Prevention of stroke and systemic embolism in adults with non-valvular atrial fibrillation (NVAf) with one or more risk factors, such as prior stroke or transient ischaemic attack (TIA), age  $\geq 75$  years, hypertension, diabetes mellitus or symptomatic heart failure (NYHA Class  $\geq$  II). Treatment of deep vein thrombosis (DVT) and pulmonary embolism (PE), and prevention of recurrent DVT and PE in adults (see Special warnings and precautions for information on haemodynamically unstable PE patients). Prevention of venous thromboembolic events (VTE) in adults who have undergone elective hip or knee replacement surgery (2.5 mg only).

**DOSAGE AND ADMINISTRATION (SPC section 4.2):** Oral. Taken with water, with or without food. Prevention of stroke and systemic embolism in patients with NVAf: The recommended dose is 5 mg twice a day. In patients who meet at least two of the following criteria: serum creatinine  $\geq 1.5$  mg/dL (133 micromole/L), age  $\geq 80$  years, or body weight  $\leq 60$  kg the recommended dose is Eliquis, 2.5 mg twice daily. Patients with severe renal impairment (creatinine clearance 15-29 mL/min) should receive Eliquis 2.5 mg twice daily. Therapy should be continued long term. Treatment of DVT, treatment of PE and prevention of recurrent DVT and PE (VTEt): The recommended dose for the treatment of acute DVT and treatment of PE is 10 mg twice daily for the first 7 days followed by 5 mg twice daily. As per available medical guidelines, short duration of treatment (at least 3 months) should be based on transient risk factors (e.g. recent surgery, trauma, immobilisation). The recommended dose for the prevention of recurrent DVT and PE is 2.5 mg twice daily. When prevention of recurrent DVT and PE is indicated, the 2.5 mg twice daily dose should be initiated following completion of 6 months of treatment with Eliquis 5 mg twice daily or with another anticoagulant. The duration of overall therapy should be individualised after careful assessment of the treatment benefit against the risk for bleeding. Prevention of VTE (VTEp): elective hip or knee replacement surgery: The recommended dose is 2.5 mg twice a day. The initial dose should be taken 12 to 24 hours after surgery. Hip replacement surgery, the recommended duration of treatment is 32 to 38 days. Knee replacement surgery, the recommended duration of treatment is 10 to 14 days. Missed Dose for All Indications: If a dose is missed, Eliquis should be taken immediately and then continue with twice daily dose as before. Switching: Switching treatment from parenteral anticoagulants to Eliquis (and vice versa) can be done at the next scheduled dose. These medicinal products should not be administered simultaneously. Switching treatment from VKA therapy to Eliquis: Warfarin or other VKA therapy should be discontinued and Eliquis started when the international normalized ratio (INR) is  $< 2$ . Switching treatment from Eliquis to VKA therapy: Administration of Eliquis should be continued for at least 2 days after beginning VKA therapy. After 2 days of co-administration of Eliquis with VKA therapy, an INR should be obtained prior to next scheduled dose of Eliquis. Co-administration of Eliquis and VKA therapy should be continued until the INR is  $\geq 2$ . Renal Impairment - mild or moderate renal impairment: For the prevention of VTE in elective hip or knee replacement surgery (VTEp), for the treatment of DVT, treatment of PE and prevention of recurrent DVT and PE (VTEt), no dose adjustment is necessary. For the prevention of stroke and systemic embolism in patients with NVAf and serum creatinine  $\geq 1.5$  mg/dL (133 micromole/L) associated with age  $\geq 80$  years or body weight  $\leq 60$  kg, a dose reduction is necessary. In the absence of other criteria for dose reduction (age, body weight), no dose adjustment is necessary. Severe renal impairment (creatinine clearance 15-29 mL/min): For the prevention of VTE in elective hip or knee replacement surgery (VTEp), for the treatment of DVT, treatment of PE and prevention of recurrent DVT and PE (VTEt), Eliquis is to be used with caution. For the prevention of stroke and systemic embolism in patients with NVAf, patients should receive the lower dose of Eliquis 2.5mg twice daily. In patients with creatinine clearance  $< 15$  mL/min, or in patients undergoing dialysis, there is no clinical experience therefore Eliquis is not recommended. See SmPC for further details. Hepatic impairment: Contraindicated in patients with hepatic disease associated with coagulopathy and clinically relevant bleeding risk. Not recommended in patients with severe hepatic impairment. Use with caution in patients with mild or moderate hepatic impairment (Child Pugh A or B). No dose adjustment is required in patients with mild or moderate hepatic impairment. Use with caution in patients with elevated liver enzymes (ALT/AST  $> 2 \times$  ULN) or total bilirubin  $\geq 1.5 \times$  ULN. Prior to initiating Eliquis, liver function testing should be performed. Catheter ablation (NVAf): Patients can continue Eliquis use while undergoing catheter ablation. Cardioversion (NVAf): Eliquis can be initiated or continued in NVAf patients who may require cardioversion. See SmPC for further details. Patients with NVAf and acute coronary syndrome (ACS) and/or percutaneous coronary intervention (PCI): There is limited experience of treatment with apixaban at the recommended dose for NVAf patients when used in combination with antiplatelet agents in patients with ACS and/or undergoing PCI after haemostasis is achieved. See SmPC for further details. Paediatric population: Eliquis is not recommended in children and adolescents below the age of 18.

**CONTRAINDICATIONS (SPC section 4.3):** Hypersensitivity to active substance or to excipients, active clinically significant bleeding, hepatic disease associated with coagulopathy and clinically relevant bleeding risk, lesion or condition if considered a significant risk factor for major bleeding, see SmPC for further details. Concomitant treatment with any other anticoagulant agent except under specific circumstances of switching anticoagulant therapy or when unfractionated heparin (UFH) is given at doses necessary to maintain an open central venous or arterial catheter or when UFH is given during catheter ablation for atrial fibrillation, see SmPC for further details.

**WARNINGS AND PRECAUTIONS (SPC section 4.4):** Haemorrhage risk: Carefully observe for signs of bleeding. Use with caution in conditions with increased risk of haemorrhage. Discontinue administration if severe haemorrhage occurs. An agent to reverse the anti-factor Xa activity of apixaban is available. For information on reversal and managing bleeding, see SmPC for further details. Interaction with other medicinal products affecting haemostasis: Concomitant treatment with any other anticoagulant is contraindicated (see contraindications). Concomitant use of Eliquis with antiplatelet agents increases the risk of bleeding. Care with concomitant SSRIs, SNRIs or NSAIDs, including acetylsalicylic acid. Following surgery, other platelet aggregation inhibitors are

not recommended concomitantly with Eliquis. In patients with atrial fibrillation and conditions that warrant mono or dual antiplatelet therapy, a careful assessment of the potential benefits against the potential risks should be made before combining this therapy with Eliquis. A clinical trial enrolled patients with atrial fibrillation with ACS and/or undergoing PCI and a planned treatment period with a P2Y12 inhibitor, with or without ASA, and oral anticoagulant (either apixaban or VKA) for 6 months. Concomitant use of ASA increased the risk of ISTH (International Society on Thrombosis and Hemostasis) major or CRNM (Clinically Relevant Non-Major) bleeding in apixaban-treated subjects. See SmPC for further details. Use of thrombolytic agents for the treatment of acute ischemic stroke: Limited experience. Patients with prosthetic heart valves: safety and efficacy of Eliquis have not been studied in patients with prosthetic heart valves, with or without atrial fibrillation. Therefore, the use of Eliquis is not recommended in this setting. Patients with antiphospholipid syndrome: Direct acting Oral Anticoagulants (DOACs), including Eliquis, are not recommended for patients with a history of thrombosis who are diagnosed with antiphospholipid syndrome (see SmPC for further details). Surgery and invasive procedures: Discontinue at least 48 hours prior to elective surgery or invasive procedures with a moderate or high risk of bleeding. Discontinue at least 24 hours prior to elective surgery or invasive procedures with a low risk of bleeding. If surgery or invasive procedures cannot be delayed, appropriate caution should be exercised, taking into consideration an increased risk of bleeding. Eliquis should be restarted after the invasive procedure or surgical intervention as soon as possible provided the clinical situation allows and adequate haemostasis has been established. For patients undergoing catheter ablation for atrial fibrillation, Eliquis treatment does not need to be interrupted. Temporary discontinuation: Discontinuing anticoagulants, including Eliquis, for active bleeding, elective surgery, or invasive procedures places patients at an increased risk of thrombosis. Lapses in therapy should be avoided and if anticoagulation with Eliquis must be temporarily discontinued for any reason, therapy should be restarted as soon as possible. Spinal/epidural anaesthesia or puncture: Patients treated with antithrombotic agents for prevention of thromboembolic complications are at risk of developing an epidural or spinal haematoma which can result in long-term or permanent paralysis. The risk of these events may be increased by the post-operative use of indwelling epidural catheters or the concomitant use of medicinal products affecting haemostasis. Indwelling epidural or intrathecal catheters must be removed at least 5 hours prior to the first dose of Eliquis. The risk may also be increased by traumatic or repeated epidural or spinal puncture. Patients are to be frequently monitored for signs and symptoms of neurological impairment (e.g., numbness or weakness of the legs, bowel or bladder dysfunction). If neurological compromise is noted, urgent diagnosis and treatment is necessary. Prior to neuraxial intervention the physician should consider the potential benefit versus the risk in anticoagulated patients or in patients to be anticoagulated for thromboprophylaxis. There is no clinical experience with the use of Eliquis with indwelling intrathecal or epidural catheters. See SmPC for further details. Haemodynamically unstable PE patients or patients who require thrombolysis or pulmonary embolectomy: Eliquis is not recommended as an alternative to unfractionated heparin in patients with pulmonary embolism who are haemodynamically unstable or may receive thrombolysis or pulmonary embolectomy since the safety and efficacy of Eliquis have not been established. Patients with active cancer: Patients with active cancer can be at high risk of both venous thromboembolism and bleeding events. When apixaban is considered for DVT or PE treatment in cancer patients, a careful assessment of the benefits against the risks should be made. Renal impairment: see dosage and administration section. Elderly patients: Increasing age may increase haemorrhagic risk. Also, the co-administration of Eliquis with ASA in elderly patients should be used cautiously because of a potentially higher bleeding risk. Body weight: Low body weight ( $< 60$  kg) may increase haemorrhagic risk. Hepatic impairment: see dosage and administration section. Interaction with Inhibitors of CYP3A4 and P-gp: Not recommended with strong inhibitors of both CYP3A4 and P-gp. These medicinal products may increase Eliquis exposure by 2-fold or greater in the presence of additional factors that increase Eliquis exposure (e.g. severe renal impairment) see SmPC for further details. Interaction with Inducers of CYP3A4 and P-gp: Eliquis should not be used for the treatment of DVT and PE in patients receiving concomitant systemic treatment with strong inducers of both CYP3A4 and P-gp since efficacy may be compromised. Concomitant systemic treatment with strong inducers of both CYP3A4 and P-gp, Eliquis should be used with caution for the prevention of VTE in elective hip or knee replacement surgery, for the prevention of stroke and systemic embolism in patients with NVAf and for the prevention of recurrent DVT and PE, though no dose adjustment for Eliquis is required during concomitant therapy with such medicinal products. Hip fracture surgery: Eliquis has not been studied in clinical trials in patients undergoing hip fracture surgery. Therefore, it is not recommended in these patients. Laboratory parameters: Clotting tests (PT, INR, and aPTT) are affected by the mechanism of action of apixaban. Changes observed at the expected therapeutic dose are small and subject to a high degree of variability, see SmPC for further details. Information about excipients: Eliquis contains lactose. Patients with galactose intolerance, total lactase deficiency or glucose-galactose malabsorption should not take Eliquis.

**DRUG INTERACTIONS (SPC Section 4.5):** Eliquis should be used with caution when co-administered with SSRIs/SNRIs, NSAIDs, ASA and/or P2Y12 inhibitors because these medicinal products typically increase the bleeding risk. There is limited experience of co-administration with other platelet aggregation inhibitors (such as GPIIb/IIIa receptor antagonists, dipyridamole, dextran or sulfapyrazone) or thrombolytic agents. As such agents increase the bleeding risk, co-administration of these products with Eliquis is not recommended. See SmPC for further details. Due to an increased bleeding risk, concomitant treatment with any other anticoagulants is contraindicated, except under specific circumstances of switching anticoagulant therapy, when UFH is given at doses necessary to maintain an open central venous or arterial catheter or when UFH is given during catheter ablation for atrial fibrillation. Administration of activated charcoal reduces Eliquis exposure. Also see contraindications and special warnings and

precautions section; Consult SmPC (contraindications, special warnings and precautions and drug interactions) for full details on interactions.

**PREGNANCY AND LACTATION (SPC section 4.6):** Pregnancy: As a precautionary measure, it is preferable to avoid the use of apixaban during pregnancy. Breastfeeding: A decision must be made whether to discontinue breast-feeding or to discontinue/abstain from apixaban therapy taking into account the benefit of breast-feeding for the child and the benefit of therapy for the woman.

**UNDESIRABLE EFFECTS (SPC section 4.8):** Increased risk of occult or overt bleeding from any tissue or organ, which may result in post haemorrhagic anaemia. The signs, symptoms, and severity will vary according to the location and degree or extent of the bleeding. Frequencies: common ( $\geq 1/100$  to  $< 1/10$ ); uncommon ( $\geq 1/1,000$  to  $< 1/100$ ); rare ( $\geq 1/10,000$  to  $< 1/1,000$ ); very rare ( $< 1/10,000$ ); not known (cannot be estimated from the available data). **Prevention of VTE in adult patients who have undergone elective hip or knee replacement surgery (VTEp):** Common: anaemia; haemorrhage\*; haematoma\*; nausea; contusion. Uncommon: thrombocytopenia\*; epistaxis\*; haematochezia\*;

liver function test abnormal (including blood bilirubin increased\*); haematuria\*; specific haemorrhage such as gastrointestinal\*, abnormal vaginal\*, urogenital\*, post procedural\*, wound secretion\*, incision site\*, operative\*. Rare: hypersensitivity\*; anaphylaxis\*; haemoptysis\*; gingival bleeding\*; specific haemorrhage such as eye (including conjunctival)\*, rectal\*, muscle\*. Not known: angioedema\*; specific haemorrhage such as brain (encompassing intracranial, intraspinal)\*, intra-abdominal\*, respiratory tract\*, haemorrhoidal\*, mouth\*, retroperitoneal\*, traumatic\*, erythema multiforme\*. **Prevention of stroke and systemic embolism in adult patients with NVAf, with one or more risk factors (NVAf):** Common: anaemia; haemorrhage\*; haematoma\*; hypotension (including procedural hypotension); epistaxis\*; nausea; gingival bleeding\*; gamma-glutamyltransferase increased; haematuria\*; contusion; specific haemorrhage such as eye (including conjunctival)\*, gastrointestinal\*, rectal\*. Uncommon: thrombocytopenia\*; hypersensitivity\*; anaphylaxis\*; haemoptysis\*; haematochezia\*; liver function test abnormal (including blood bilirubin increased\*); specific haemorrhage such as brain (encompassing intracranial, intraspinal)\*, intra-abdominal\*, haemorrhoidal\*, mouth\*, abnormal vaginal\*, urogenital\*, post procedural\*, wound secretion\*, incision site\*, operative\*, traumatic\*. Rare: specific haemorrhage such as respiratory tract\*, retroperitoneal\*, muscle\*. Very Rare: erythema multiforme\*. Not known: angioedema\*.

**Treatment of DVT and PE, and prevention of recurrent DVT and PE (VTEt):** Common: anaemia; thrombocytopenia\*; haemorrhage\*; haematoma\*; epistaxis\*; nausea; gingival bleeding\*; gamma-glutamyltransferase increased; alanine aminotransferase increased; skin rash; haematuria\*; contusion; specific haemorrhage such as gastrointestinal\*, mouth\*, rectal\*, abnormal vaginal\*, urogenital\*. Uncommon: hypersensitivity\*; anaphylaxis\*; haemoptysis\*; haematochezia\*; liver function test abnormal (including blood bilirubin increased\*); specific haemorrhage such as eye (including conjunctival)\*, haemorrhoidal\*, muscle\*, post procedural\*, wound secretion\*, incision site\*, operative\*, traumatic\*. Rare: specific haemorrhage such as brain (encompassing intracranial, intraspinal)\*, respiratory tract\*. Not Known: angioedema\*; specific haemorrhage such as intra-abdominal\* and retroperitoneal\*, erythema multiforme\*.

\*Denotes serious adverse reaction

**Refer to SmPC for all other adverse events**

**LEGAL CATEGORY:** POM.

**MARKETING AUTHORISATION NUMBER (SPC section 8):**

EU/1/11/691/002-3, EU/1/11/691/008, EU/1/11/691/014

**PACKAGE QUANTITIES:** Carton of 20 film-coated tablets 2.5 mg, 60 film-coated tablets 2.5 mg, 56 film-coated tablets 5 mg, 28 film-coated tablets 5 mg.

**MARKETING AUTHORISATION HOLDER (SPC section 7):** Bristol-Myers Squibb/Pfizer EEIG, Plaza 254, Blanchardstown Corporate Park 2, Dublin 15, D15 T867, Ireland

**FOR FURTHER INFORMATION CONTACT:** medical.information@bms.com or 1 800 749 749 (Ireland)

**DATE OF PREPARATION:** April 2021

**ADDITIONAL INFORMATION AVAILABLE ON REQUEST**

**Approval Code:** 432-IE-2100041

**Adverse events should be reported. Reporting forms and information can be found at:**

**Ireland - via HPRa Pharmacovigilance at [www.hpra.ie](http://www.hpra.ie)**

**Adverse events should also be reported to Bristol-Myers Squibb via [medical.information@bms.com](mailto:medical.information@bms.com) or**

**1 800 749 749 (Ireland)**

**Date of preparation:** June 2023

**Job code:** 432-IE-2300057