# ECONOMIC IMPACT OF RIVAROXABAN USE FOR THROMBOEMBOLIC EVENTS PREVENTION AFTER HIP OR KNEE REPLACEMENT IN SPAIN

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### BACKGROUND AND AIM

- Patients undergoing elective total hip or knee replacement (THR or TKR) surgery present risk of venous thromboembolism (VTE). VTE is considered as one of the most common preventable hospital-related cause of death.<sup>1</sup> Anti-thrombotic prophylaxis is routinely used for decreasing the risk of VTE by 50-75%.<sup>2,3</sup> In Spain, this is performed with low-molecular-weight heparins (LMWHs) in approximately 90% of the patients, enoxaparin being the most widely used.<sup>2,3</sup>
- Enoxaparin has relevant drawbacks like risk of heparin-induced thrombocytopenia, need of patient training for post-discharge self-administration or home administration by a health professional, the treatment should begin 12 hours before surgery, and needs dose adjustment in case of extreme weight values.<sup>2</sup> Conversely, new oral anticoagulants, like rivaroxaban, offer several advantages over enoxaparin.<sup>4</sup>
- Rivaroxaban (factor Xa inhibitor) 10 mg once-daily dose, a direct oral anticoagulant, has demonstrated superior efficacy and similar safety profile to enoxaparin in preventing VTE in patients undergoing elective THR or TKR surgery in randomized clinical trials (RECORD program<sup>5</sup>), as well as, in the international observational XAMOS (Xarelto® for VTE Prophylaxis After Hip or Knee Arthroplasty)<sup>6</sup> study in the clinical routine practice.
- The aim of this study was to estimate the potential cost-savings from an increased use of rivaroxaban with respect to enoxaparin in the prophylaxis of VTE in patients undergoing elective THR or TKR surgery for the Spanish National Health System (NHS) perspective.

#### **METHODS**

- A budget impact model (BIM) was performed based on the efficacy data of the Spanish cohort of the XAMOS study.<sup>6</sup>
- The BIM estimates the difference in the annual treatment cost between 2 scenarios:
  - ✓ Current scenario: the use of rivaroxaban vs. enoxaparin is estimated in a 30-to-70 ratio.³
  - ✓ Alternative scenario: rivaroxaban use increases progressively with respect to enoxaparin by 10%, 20% and 30% in the years 2017, 2018 and 2019.
- The population included in the BIM was the Spanish patients undergoing THR or TKR surgery estimated based on the annual incidence of the 2 surgeries in Spain<sup>7</sup> applied to the total Spanish population<sup>8</sup> (Table 1).

Table 1. Target population of the study

	Surgery	Number of patients undergoing THR or TKR surgery				
	prevalence	2017	2018	2019		
THR	0,043%	19,654	19,542	19,427		
TKR	0,073%	33,366	33,175	32,981		

• Treatment effectiveness (thromboembolism and major bleeding rates) and healthcare resource use (anaesthesia and nurse care) for patients undergoing elective THR or TKR surgery in the routine practice were derived from data from the Spanish cohort of the XAMOS study<sup>6</sup> (Table 2).

Table 2. Event rates and healthcare resource use associated with rivaroxaban and Standard of Care (i.e. enoxaparin) in the Spanish Cohort of the XAMOS study

Type of event or healthcare resource		Treatment			
Type of event or near	ncare resource	Rivaroxaban	SoC enoxaparin 0.80%		
Thromboembolic event		0.20%			
A	General	18%	23%		
Anesthesia	Local	82%	77%		
Major bleeding (EMA Guideline	€)	0.70%	1.30%		
Post-discharge treatment	Home	3.70%	8.20%		
administration by a nurse	Healthcare facility	0.2%	2.60%		

- Relevant direct costs (pharmacologic prophylaxis, thromboembolic events, anaesthesia, major bleeding events and post-discharge treatment administration by a nurse) were estimated from the Spanish NHS perspective (€ 2014). Unit costs were derived from the corresponding Spanish official source<sup>9,10,11</sup>, except for the cost of major bleeding which is derived from a similar Spanish publication.³ Unit health care resource use and event costs are detailed in Table 3.
- The pharmacological cost was estimated based on the daily consumer price with VAT, the 8/2010 Royal Decree<sup>12</sup> discount was considered as well as co-payment (NHS assuming the 90% of the treatment cost), and applying the mean duration of the treatment after THR and TKR surgery (30 days with both treatment options) from hospital discharge, resulting in a mean cost per patient of €75.65 and €103.46 for rivaroxaban and enoxaparin, respectively.
- The mean cost per patient was estimated as average cost weighted on the incidence or percentage use of the event or resource. The cost per patient was equal for the 2 surgeries (THR or TKR).
- The annual cost was estimated based on the mean cost per patient and the annual use of rivaroxaban and enoxaparin in both scenarios (current and alternative) over the 3 years.

Table 3. Healthcare resource use and event unit costs

Type of cost		Definition	Unit cost	Reference
Dharman	Rivaroxaban	Daily cost of Xarelto® 10mg (consumer price	€2.52	9
Pharmacy cost	Enoxaparin	Daily cost of Clexane® 40 mg (4000 UI) (consumer price)	€3.45	9
Thromboembolic event		Cost per event calculated as a weighted average (6:4) of the cost of the Pulmonary Embolism (Diagnosis-Related Group; DRG78) and Deep Vein Thromboembolism (DRG128)	€3,373.04	10
A 41 : -	General	Unit cost of general anaesthesia	€207.21	11
Anesthesia	Local	Unit cost of local anaesthesia	€51.80	11
Major bleeding		Unit cost Diagnosis-Related Group	€3,718.68	3
Post-discharge treatment administration by a nurse	Home	Nurse for administration of post-discharge antithrombotic therapy	€16.30	11
	Healthcare facility		€7.12	11

#### RESULTS

 The difference in per-patient cost undergoing THR or TKR surgery with rivaroxaban versus enoxaparin was -€105 (Table 4).

Table 4. Mean cost per patient

2017

	Pharmacy costs	Event costs	Anesthesia costs	Major bleeding costs	Nursing costs	Total costs	Mean savings per patient
Rivaroxaban	€75.65	€6.75	€79.77	€26.03	€18.52	€206.72	£105.26
Enoxaparin	€103.46	€26.98	€87.54	€48.34	€45.65	€311.98	<b>-€</b> 105.26

The impact of progressively increasing the use of rivaroxaban in the THR surgery cohort, was
 -€206,890, -€411,413, and -€613,506 for 2017, 2018, and 2019, respectively (Figure 1).

Figure 1. Annual savings associated with an increased use of rivaroxaban in the VTE prophylaxis in undergoing THR surgery

2018

2019

2019

2019

■ Total cost (alternative scenario)

**Accumulated cost** 

Accumulated cost

**Accumulated cost** 

2017

Current	Rivaroxaban use	30%	30%	30%	<b></b>
scenario	Total cost	€5,511,048	€5,479,519	€5,447,428	€16,437,995
Alternative	Rivaroxaban use	40%	50%	60%	
scenario	<b>scenario</b> <i>Total cost</i> €5,304,158 €5,068,106		€4,833,922	€15,206,186	
€ 18,000,000		THR	surgery cos	sts	Cumulative savings: €1,231,809
€ 14,000,000					
€ 10,000,000	Annual saving: €206,890	Annual sa €411,41	•	Annual saving: €613,506	
€ 10,000,000 € 6,000,000 € 2,000,000			•		

In the TKR cohort, the impact of progressively increasing the use of rivaroxaban was -€351,232, - €698,446, and -€1,041,533, in 2017, 2018, 2019, respectively (Figure 2).

2018

■ Total cost (current scenario)

2017

Figure 2. Annual savings associated with an increased use of rivaroxaban in the VTE prophylaxis in patients undergoing TKR surgery

2018

Current	Rivaroxaban use	30%	30%	30%	
scenario	Total cost	€9,355,966	€9,302,439	€9,247,959	€27,906,364
Alternative	Rivaroxaban use	40%	50%	60%	
scenario	Total cost	€9,004,734	€8,603,994	€8,206,426	€25,815,154
80,000,000		TKR su	rgery costs		Cumulative savings: €2,091,210
20,000,000	Annual saving: €351,232	Annual saving €698,445		nual saving: €1,041,533	
0,000,000	COO 1,202				
U —	2017	2018		2019	Accumulated cost
	■ Total cos	st (current scenario)	■ Total cost (al	ternative scenario)	

## CONCLUSIONS

- The analysis showed that, according to effectiveness data from the XAMOS study, an international, non-interventional, observational, open-label study in unselected patients undergoing elective hip or knee surgery in routine practice, rivaroxaban was more effective and less costly than the current SoC, enoxaparin, in the prevention of VTE in Spain.
- The study demonstrated that rivaroxaban increased use would lead in significant cost-savings for the Spanish NHS in the VTE prophylaxis in patients undergoing elective THR and TKR surgery.

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