

Appendix 7: Association of anticoagulation intensity with risk of events in studies that included patients with atherosclerotic diseases only (a), with atrial fibrillation only (b) and with venous thromboembolism only (c)

Appendix 7a: Association of anticoagulation intensity with risk of events in studies that included only patients with atherosclerotic diseases

International normalized ratio	Hemorrhagic event (n = 54 207)		Thromboembolic event (n = 54 207)		Combined events (n = 54 207)	
	Relative risk (95% CI)	Absolute risk (95% CI)	Relative risk (95% CI)	Absolute risk (95% CI)	Relative risk (95% CI)	Absolute risk (95% CI)
< 2	1.4 (0.6–3.4)	1.8 (0.6–5.7)	3.4 (2.5–4.6)	11.3 (6.8–18.7)	2.8 (2.3–3.4)	11.8 (7.4–18.7)
2–3	1.0 (Ref)	1.3 (0.7–2.5)	1.0 (Ref)	3.3 (2.3–4.7)	1.0 (Ref)	4.2 (2.8–6.3)
3–5	2.7 (1.7–4.2)	3.5 (1.8–6.6)	0.6 (0.4–1.1)	2.1 (1.4–3.0)	1.2 (0.7–2.0)	5.1 (3.5–7.4)
> 5	19.8 (9.4–41.5)	26.0 (12.2–55.1)	2.2 (0.9–5.1)	7.2 (2.5–20.9)	7.2 (3.5–14.9)	30.2 (15.7–58.0)

Appendix 7b: Association of anticoagulation intensity with risk of events in studies that included only patients with atrial fibrillation

International normalized ratio	Hemorrhagic event (n = 414 531)		Thromboembolic event (n = 414 531)		Combined events (n = 414 531)	
	Relative risk (95% CI)	Absolute risk (95% CI)	Relative risk (95% CI)	Absolute risk (95% CI)	Relative risk (95% CI)	Absolute risk (95% CI)
< 2	0.3 (0.1–0.9)	0.4 (0.3–0.5)	4.8 (3.2–7.1)	6.1 (3.1–12.0)	2.3 (1.1–5.0)	4.9 (2.8–8.4)
2–3	1.0 (Ref)	1.3 (0.5–3.4)	1.0 (Ref)	1.3 (0.6–2.7)	1.0 (Ref)	2.1 (0.9–4.8)
3–5	4.8 (2.4–9.7)	6.1 (1.7–22.5)	1.6 (0.8–3.4)	2.0 (0.6–6.3)	2.2 (1.3–3.8)	4.7 (1.9–11.4)
> 5	20.3 (10.7–38.3)	25.7 (9.7–67.9)	3.9 (3.0–5.1)	4.5 (2.6–7.9)	15.3 (9.5–24.7)	32.3 (13.2–79.1)

Appendix 7c: Association of anticoagulation intensity with risk of events in studies that included only patients with venous thromboembolism

International normalized ratio	Hemorrhagic event (n = 22 041)		Thromboembolic event (n = 32 096)		Combined events (n = 22 041)	
	Relative rate (95% CI)	Absolute rate (95% CI)	Relative rate (95% CI)	Absolute rate (95% CI)	Relative rate (95% CI)	Absolute rate (95% CI)
< 2	1.3 (0.6–2.7)	1.1 (0.2–6.0)	4.1 (2.0–8.5)	14.6 (4.1–52.3)	2.4 (2.1–2.7)	10.5 (2.0–54.6)
2–3	1.0 (Ref)	1.0 (0.4–2.2)	1.0 (Ref)	3.6 (1.0–13.0)	1.0 (Ref)	4.4 (1.0–20.2)
3–5	5.4 (1.5–19.4)	4.9 (3.4–6.9)	0.6 (0.2–1.9)	2.2 (0.4–11.7)	1.6 (0.6–4.1)	7.0 (3.9–12.4)
> 5*						

Note: CI = confidence interval.

*Studies did not report events and patient observation time that could be grouped into the international normalized ratio category > 5.