

ANTIPLATELET THERAPY

Agents at a glance · mechanisms · dosing · precautions

teal = agent · red = caution / interacting drug / contraindicated · green = reversible / preferred

Agent	Mechanism	Onset · Reversibility	Typical dosing	Precautions
Aspirin COX-1 inhibitor	Irreversibly acetylates COX-1 → ↓ thromboxane A ₂ .	Onset <1 h (chewed). Irreversible (~7–10 d).	Acute stroke 81–300 mg ; then 81 mg daily.	GI bleeding (dose-related); NSAIDs blunt its effect — take ASA first (see table below).
Clopidogrel (Plavix) P2Y ₁₂ — prodrug	Irreversible P2Y₁₂ ; prodrug — needs CYP2C19 activation.	Onset 2–6 h. Irreversible (~5–7 d).	Load 300–600 mg ; then 75 mg daily.	Avoid if CYP2C19 slow metabolizer or P2Y ₁₂ test > 180 PRU (test 8 h after loading, or day 7 if no load). Avoid CYP inhibitors ¹ .
Ticagrelor (Brilinta) P2Y ₁₂ — direct	Reversible, direct-acting P2Y ₁₂ (not a prodrug).	Onset 0.5–2 h. Reversible (~3–5 d, BID).	Load 180 mg ; then 90 mg BID .	Avoid CYP3A drugs ² . Caution: digoxin, opioids, simvastatin / lovastatin. Dyspnea common.
Prasugrel (Effient) P2Y ₁₂ — prodrug	Irreversible P2Y ₁₂ ; faster/more reliable than clopidogrel.	Onset ~0.5 h. Irreversible (~7–10 d).	Load 60 mg; then 10 mg daily (cardiology).	CONTRAINDICATED with prior stroke / TIA — excess intracranial hemorrhage.
Dipyridamole + ASA (Aggrenox) PDE / adenosine	Dipyridamole inhibits PDE & adenosine reuptake → ↑ cAMP; + aspirin.	Onset hours. Aspirin component irreversible.	ASA 25 mg / ER-dipyridamole 200 mg BID .	Headache (common, limits tolerability); caution in CAD.
Cilostazol PDE3 inhibitor	Inhibits PDE3 → ↑ cAMP; antiplatelet + vasodilator.	Onset hours. Reversible.	100 mg BID.	Contraindicated in heart failure . Avoid in acute coronary syndromes. Caution with CYP3A4 & CYP2C19 inhibitors.
IV agents cangrelor · GP IIb/IIIa	Cangrelor = IV reversible P2Y ₁₂ . Abciximab / eptifibatide / tirofiban = GP IIb/IIIa blockers.	Onset minutes. Rapid on/off .	Weight-based IV infusion.	Peri-procedural in neuro-intervention (stenting, thrombectomy rescue). Bridging when oral route unavailable.
Vorapaxar (Zontivity) PAR-1 antagonist	Blocks thrombin receptor PAR-1 on platelets.	Very long-acting. Effectively irreversible .	2.08 mg daily (add-on).	CONTRAINDICATED with prior stroke / TIA / ICH — intracranial bleeding.

¹ CYP inhibitors (avoid with clopidogrel): omeprazole, esomeprazole, fluconazole, voriconazole, fluoxetine, fluvoxamine.

² CYP3A drugs (avoid with ticagrelor): ketoconazole, clarithromycin, rifampin, phenytoin, phenobarbital.

⚠ Key Interactions — Medications to Avoid

Combination	Why it matters	What to do
Aspirin + ibuprofen / NSAIDs	Blunts aspirin's antiplatelet effect & ↑ GI bleeding.	Take aspirin first; prefer acetaminophen.
Clopidogrel + omeprazole (or esomeprazole)	↓ clopidogrel efficacy (CYP2C19 inhibition).	Use pantoprazole.
Clopidogrel + CYP2C19 loss-of-function	Poor activation → ↓ efficacy.	Prefer ticagrelor.
Ticagrelor + CYP3A4 inducers (carbamazepine, phenytoin, phenobarbital)	↓ ticagrelor efficacy.	Use a non-enzyme-inducing AED.
Ticagrelor + strong CYP3A4 inhibitors (clarithromycin, ketoconazole)	↑ ticagrelor levels → ↑ bleeding.	Avoid; switch antibiotic / antifungal.
Prasugrel or Vorapaxar + prior stroke / TIA	Markedly ↑ intracranial hemorrhage.	Contraindicated — use clopidogrel, ticagrelor, or ASA.
Any antiplatelet + SSRIs / SNRIs	Additive bleeding risk (esp. GI).	Weigh need; consider GI protection; counsel on bleeding.
Dual therapy (antiplatelet + anticoagulant)	Markedly ↑ risk of bleeding.	Minimize duration; avoid long-term use.