

Clinical and economic impact of platelet function testing in neurointervention

Balancing procedure-related and therapeutic risks in neurointervention

Flow diversion—the preferred treatment modality for many intracranial aneurysms—has risks¹

- Thromboembolic complications occurred in 9.2% of procedures²
- Intracranial hemorrhage occurred in up to 6.2% of patients³

Dual antiplatelet therapy (DAPT) (e.g., aspirin, clopidogrel) is used more than 90% of the time before intracranial stent placement⁴

 While DAPT reduces the risk of thromboembolic complications, major hemorrhagic complications have been reported in 11.1% of patients⁵⁻⁷



Individual response to clopidogrel varies8

- Up to 50% of patients demonstrate variable response to clopidogrel⁹
- In a flow diversion multicenter study, 28.8% of patients were identified as clopidogrel nonresponders²



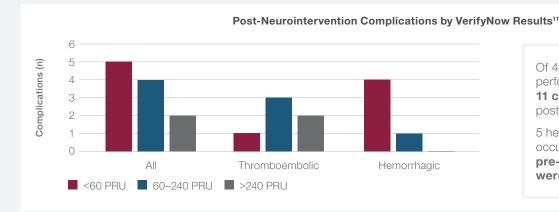
Clopidogrel hypo-responders have a 60% higher risk of thrombosis.¹⁰



Clopidogrel hyper-responders have an 11% higher risk of hemorrhage.¹⁰

Predicting patient bleeding and thromboembolic risk⁶

VerifyNow results have been demonstrated to be a predictor of periprocedural hemorrhagic and thromboembolic complications in patients undergoing flow diversion.^{9,11}



Of 48 flow diversion procedures performed on 44 patients, **11 complications** occurred post-procedure (22.9%).

5 hemorrhagic complications occurred, and in 4 of them, pre-procedural results were <60 PRU.

Several publications have reviewed treatment algorithms for P2Y12 inhibitor management in the setting of flow diversion

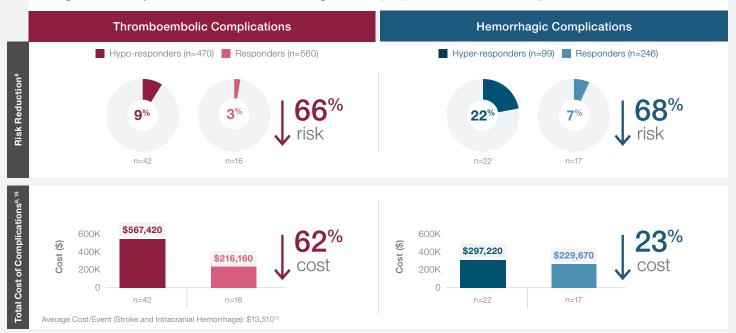
- An example algorithm using a target range of 60–240 PRU resulted in identification of clopidogrel hyper-responders, which required dose adjustment in 71% of patients^{11–13}
- Another study using a result of >208 PRU to indicate clopidogrel nonresponders resulted in less thromboembolic complications
 when a dose adjustment was made (9.8% vs. 51.9%)²

"Platelet function tests, particularly point-of-care tests like VerifyNow, have been increasingly adopted to assess patient responses, anticipate complications, and better manage these patients perioperatively." 14

Predicting patient thromboembolic and bleeding risks with VerifyNow testing demonstrated reduced risk⁶

A meta-analysis, which included 15 studies, evaluated the usefulness of a VerifyNow result as a tool to predict periprocedural complications. Three categories of patient cohorts were defined: hypo-responders (>240 PRU), responders and hyper-responders (<175 PRU).

Using the meta-analysis data, a simulation model utilizing estimates (US\$) was built to calculate complication risk and cost reduction.¹⁵



References

- Walcott B, Stapleton C, Choudhri O, Patel AB. Flow diversion for the treatment of intracranial aneurysms. JAMA Neurology. 2016;73(8):1002–1008.
- Adeeb N, Griessenauer C, Foreman P, Moore JM, Shallwani H, Motiei-Langroudi R, et al. Use
 of platelet function testing before pipeline embolization device placement: a multicenter cohort
 study. Stroke. 2017;48(5):1322–1330.
- Al-Mufti F, Amuluru K, Gandhi C, Gandhi CD, Prestigiacomo CJ, et al. Flow diversion for intracranial aneurysm management: a new standard of care. Neurotherapeutics. 2016;13(3):582– 589.
- Faught R, Satti S, Hurst R, Pukenas BA, Smith MJ. Heterogeneous practice patterns regarding antiplatelet medications for neuroendovascular stenting in the USA: a multicenter survey. J NeuroIntervent Surg. 2014;6(10):774–779.
- Goh C, Churilov L, Mitchell P, Yan B. Clopidogrel hyper-response and bleeding risk in neurointerventional procedures. Am J Neuroradiol. 2013;34(4):721–726.
- Kim H, Oh J, Park S, Yoon SM, Ahn HS, Kim BT. The efficacy of P2Y12 reactive unit to predict the periprocedural thromboembolic and hemorrhagic complications according to clopidogrel responsiveness and safety of modification of dual antiplatelet therapy: a meta-analysis. J Korean Neurosurg Soc. 2020;63(5):539–549.
- Nishi H, Nakahara I, Matsumoto S, Hashimoto T, Ohta T, Sadamasa N, et al. Platelet reactivity and hemorrhage risk in neurointerventional procedures under dual antiplatelet therapy. J NeuroIntervent Surg. 2016;8:949–953.
- Yi H, Hwang G, Lee B. Variability of platelet reactivity on antiplatelet therapy in neurointerventional procedure. J Korean Neurosurg Soc. 2019;62(1):3–9.

- Tonetti D, Jankowitz B, Gross B. Antiplatelet therapy in flow diversion. Neurosurgery. 2020;86(S1):S47–52.
- 10. Delgado Almandoz J, Kadkhodayan Y, Crandall B, Scholz JM, Fease JL, Tubman DE. Variability in initial response to standard clopidogrel therapy, delayed conversion to clopidogrel hyper-response, and associated thromboembolic and hemorrhagic complications in patients undergoing endovascular treatment of unruptured cerebral aneurysms. J NeuroIntervent Surg. 2014;6(1):727-722.
- 11. Delgado Almandoz J, Crandall B, Scholz J, Fease JL, Anderson RE, Kadkhodayan Y, et al. Last-recorded P2Y12 reaction units value is strongly associated with thromboembolic and hemorrhagic complications occurring up to 6 months after treatment in patients with cerebral aneurysms treated with the pipeline embolization device. Am J Neuroradiol. 2014;35(1):128–135.
- Kayan Y, Delgado Almandoz J, Fease J, Milner AM, Scholz JM, Mulder M. Efficacy of a two-test protocol for achieving a therapeutic response to clopidogrel prior to elective endovascular intracranial aneurysm treatment and an 'induced' postoperative hyper-response. J NeuroIntervent Surg. 2017;9:792–814.
- Kim K, Fraser J, Grupke S, Cook AM. Management of antiplatelet therapy in patients undergoing neuroendovascular procedures. *J Neurosurg*. 2018;129(4):890–905.
 Ajadi E, Kabir S, Cook A, Grupke S, Alhajeri A, Fraser JF. Predictive value of platelet reactivity unit
- Ajadi E, Kabir S, Cook A, Grupke S, Alhajeri A, Fraser JF. Predictive value of platelet reactivity unit (PRU) value for thrombotic and hemorrhagic events during flow diversion procedures: a meta-analysis. J NeuroIntervent Surg. 2019;11(11):1123–1128.
 Fuller R, McCullough E, Bao M, Averill RF. Estimating the costs of potentially preventable hospital
- Fuller R, McCullough E, Bao M, Averill RF. Estimating the costs of potentially preventable hospital acquired complications. Health Care Financing Review. 2009;30(4):17–32.

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