# When will acute stroke interventions be as widely available as primary PCI?



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# Introduction

During the period from 1991 to 1993 I was working in the Netherlands where I underwent research training (Erasmus University Rotterdam) and interventional cardiology training (Thoraxcenter Zwolle). That was the fascinating period when thrombolytic treatment of acute myocardial infarction became the standard of care and the first small randomised trials demonstrated that primary percutaneous coronary intervention (p-PCI) is superior to thrombolysis in the treatment of patients presenting with acute ST-segment elevation myocardial infarction (STEMI)<sup>1-3</sup>. It took nine more years before the first national (Czech) guidelines recommended primary angioplasty (over thrombolysis) as the preferred treatment for all patients with acute STEMI<sup>4</sup>. This was followed one year later by the European guidelines<sup>5</sup> and two years later by the American guidelines<sup>6</sup>. It took another seven years for primary angioplasty in Europe to become widely available for nearly all patients with acute STEMI<sup>7,8</sup>. Thus, the delay between the first randomised evidence for p-PCI and implementation in real-world practice was nearly 20 years... Fortunately, today this is just history, as approximately 80% of European patients (and even close to 100% of those living in large urban areas) now have access to p-PCI. Primary PCI networks exist in all European countries irrespective of the economic strength of the country (interestingly, in rich

countries such as the UK and France, p-PCI was widely implemented later than in many Eastern European countries). Thus, cardiologists are now experienced in treating emergency patients including those after cardiopulmonary resuscitation, in cardiogenic shock or pulmonary oedema. This is why cardiologists are so interested in offerng their skills, time and effort for the benefit of other patients presenting with life-threatening diseases, e.g., acute ischaemic stroke.

In 2015, a similar revolution in the treatment of acute ischaemic stroke (AIS) occurred: mechanical thrombectomy (catheter-based intervention) was clearly shown to be superior to thrombolysis alone in five randomised trials<sup>9-13</sup> published that year. There is one important difference: the neuro community (neurologists, neuroradiologists, neurosurgeons) developed new guidelines very rapidly, so that the nine- to eleven-year delay in the development of guidelines for cardiologists described above was shortened to just one year for acute stroke guidelines14-17. This offers a better prospect for patients with acute ischaemic stroke. But is it realistic? Can we really expect that all suitable (i.e., early presenters with larger AIS) European patients (irrespective of their location) will have access to catheter-based therapy in the near future? There are many obstacles and limitations which may hamper the availability of acute stroke mechanical thrombectomy for the European population, especially in some countries.

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#### THE NATURE OF AIS

Longer patient-related delays (stroke is not painful and many patients are not able to call for help themselves due to acute speech disability) combined with a shorter time window for effective treatment result in a significantly lower proportion of patients indicated for thrombectomy within the optimal time window (ideally <4.5 hours; only in properly selected patients may a longer delay be acceptable).

#### THE COMPLEXITY OF STROKE CARE

While for STEMI patients a single specialist (cardiologist) must be immediately available 24/7/365, the multidisciplinaty nature of acute stroke care requires numerous specialists to be available even more quickly: neurologist, imaging radiologist, endovascular interventionalist, intensivist or anaesthesiologist, sometimes also a neurosurgeon or cardiologist, etc.

### THE LACK OF TRAINED NEUROINTERVENTIONALISTS

There is a lack of trained neurointerventionalists in most countries. There are just a few countries in Europe (a maximum of five) with a sufficient number of endovascular interventionalists to cover the needs of the population for acute stroke care. Even in these best practice countries, the immediate need is to increase the numbers of procedures three to fourfold compared to 2016. In many countries (with acute stroke intervention rates well below 10/million/year) the required increase in acute stroke interventions should be far greater to meet the needs of the population. Because of limited human resources and the duration of training, it may take many years before acute stroke interventions will become available to all European patients to the same extent as primary angioplasty is available for STEMI patients.

#### THE TURF BATTLES

Turf battles exist between many of the different medical specialisations. We see some interventionalists trying to protect their "field of interest" from the influx of "foreigners" (physicians of other specialties), partly with good intent, i.e., to protect patients from complications caused by inexperienced beginners. The result of this protective attitude will be that outcomes may be excellent - but only for a few patients. Acute stroke care is now facing a huge dilemma: A) fast and wide implementation of mechanical thrombectomy for many patients at the price of less excellent results, versus B) slow and selective care for fewer patients with better results in those lucky enough to receive such care.

In recognition of the increasing importance of stroke for cardiac patients, in 2016 the European Society of Cardiology (ESC) founded a new interdisciplinary constituent body – the ESC Council on Stroke<sup>18</sup>. The leadership of this council (https://www.escardio.org/Councils/Council-on-Stroke/esc-council-on-stroke) is truly interdisciplinary: three cardiologists, one stroke neurologist, one interventional neuroradiologist, one vascular surgeon. From the very beginning, the council sought cooperation with other societies

interested in stroke. While there has been a positive response from neurosurgeons and vascular surgeons, the response from stroke neurologists (namely the European Stroke Organisation) and from the neuroradiologists (the European Society for Minimally Invasive Neurological Therapy) has been rather cold. The requirement for two years' full-time training in a neuroradiology department for those interventional cardiologists who wish to treat acute stroke patients is not realistic. No mature interventional cardiologist can leave his/her cathlab for two years. The ESC Council on Stroke is seeking a more open interdisciplinary discussion on this topic. We believe that, for example, experienced interventional cardiologists (with greater than five years of interventional practice), who are also carotid operators (regularly performing carotid stenting procedures), can easily learn to perform acute stroke interventions after a much shorter period (e.g., three months) of neuroradiology training. From the patient's perspective, this is the only way to offer this effective treatment for acute stroke to the broad European population. I do hope that representatives of the respective medical specialties will place patient interests first and foremost and forget about the unnecessary turf battles.

## **Conflict of interest statement**

The author has no conflicts of interest to declare.

## References

- 1. Zijlstra F, de Boer MJ, Hoorntje JC, Reiffers S, Reiber JH, Suryapranata H. A comparison of immediate coronary angioplasty with intravenous streptokinase in acute myocardial infarction. *N Engl J Med.* 1993;328:680-4.
- 2. Grines CL, Browne KF, Marco J, Rothbaum D, Stone GW, O'Keefe J, Overlie P, Donohue B, Chelliah N, Timmis GC, Vlietstra RE, Strzelecki M, Puchrowicz-Ochocki S, O'Neill WW. A comparison of immediate angioplasty with thrombolytic therapy for acute myocardial infarction. The Primary Angioplasty in Myocardial Infarction Study Group. *N Engl J Med.* 1993;328:673-9.
- 3. Gibbons RJ, Holmes DR, Reeder GS, Bailey KR, Hopfenspirger MR, Gersh BJ. Immediate angioplasty compared with the administration of a thrombolytic agent followed by conservative treatment for myocardial infarction. The Mayo Coronary Care Unit and Catheterization Laboratory Groups. *N Engl J Med.* 1993;328:685-91.
- 4. Widimsky P, Janoušek S, Vojáček J; on behalf of the Czech Society of Cardiology. Guidelines for the diagnosis and treatment of acute myocardial infarction (Q-wave/ST elevation/bundle branch block). *Cor et Vasa* 44 (2002) K123-K143.
- 5. Van de Werf F, Ardissino D, Betriu A, Cokkinos DV, Falk E, Fox KA, Julian D, Lengyel M, Neumann FJ, Ruzyllo W, Thygesen C, Underwood SR, Vahanian A, Verheugt FW, Wijns W; Task Force on the Management of Acute Myocardial Infarction of the European Society of Cardiology. Management of acute myocardial infarction in patients presenting with ST-segment elevation. The Task Force on the Management of Acute Myocardial Infarction of the European Society of Cardiology. *Eur Heart J.* 2003;24:28-66.

- 6. Antman EM, Anbe DT, Armstrong PW, Bates ER, Green LA, Hand M, Hochman JS, Krumholz HM, Kushner FG, Lamas GA, Mullany CJ, Ornato JP, Pearle DL, Sloan MA, Smith SC Jr; American College of Cardiology; American Heart Association; Canadian Cardiovascular Society. ACC/AHA guidelines for the management of patients with ST-elevation myocardial infarction-executive summary. A report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines (Writing Committee to revise the 1999 guidelines for the management of patients with acute myocardial infarction). *J Am Coll Cardiol.* 2004;44:671-719.
- 7. Widimsky P, Wijns W, Fajadet J, de Belder M, Knot J, Aaberge L, Andrikopoulos G, Baz JA, Betriu A, Claeys M, Danchin N, Djambazov S, Erne P, Hartikainen J, Huber K, Kala P, Klinceva M, Kristensen SD, Ludman P, Ferre JM, Merkely B, Milicic D, Morais J, Noc M, Opolski G, Ostojic M, Radovanovic D, De Servi S, Stenestrand U, Studencan M, Tubaro M, Vasiljevic Z, Weidinger F, Witkowski A, Zeymer U; European Association for Percutaneous Cardiovascular Interventions. Reperfusion therapy for ST elevation acute myocardial infarction in Europe: description of the current situation in 30 countries. *Eur Heart J.* 2010;31: 943-57.
- 8. Kristensen SD, Laut KG, Fajadet J, Kaifoszova Z, Kala P, Di Mario C, Wijns W, Clemmensen P, Agladze V, Antoniades L, Alhabib KF, De Boer MJ, Claeys MJ, Deleanu D, Dudek D, Erglis A, Gilard M, Goktekin O, Guagliumi G, Gudnason T, Hansen KW, Huber K, James S, Janota T, Jennings S, Kajander O, Kanakakis J, Karamfiloff KK, Kedev S, Kornowski R, Ludman PF, Merkely B, Milicic D, Najafov R, Nicolini FA, Noč M, Ostojic M, Pereira H, Radovanovic D, Sabaté M, Sobhy M, Sokolov M, Studencan M, Terzic I, Wahler S, Widimsky P; European Association for Percutaneous Cardiovascular Interventions. Reperfusion therapy for ST elevation acute myocardial infarction 2010/2011: current status in 37 ESC countries. *Eur Heart J.* 2014; 35:1957-70.
- 9. Berkhemer OA, Fransen PS, Beumer D, van den Berg LA, Lingsma HF, Yoo AJ, Schonewille WJ, Vos JA, Nederkoorn PJ, Wermer MJ, van Walderveen MA, Staals J, Hofmeijer J, van Oostayen JA, Lycklama à Nijeholt GJ, Boiten J, Brouwer PA, Emmer BJ, de Bruijn SF, van Dijk LC, Kappelle LJ, Lo RH, van Dijk EJ, de Vries J, de Kort PL, van Rooij WJ, van den Berg JS, van Hasselt BA, Aerden LA, Dallinga RJ, Visser MC, Bot JC, Vroomen PC, Eshghi O, Schreuder TH, Heijboer RJ, Keizer K, Tielbeek AV, den Hertog HM, Gerrits DG, van den Berg-Vos RM, Karas GB, Steyerberg EW, Flach HZ, Marquering HA, Sprengers ME, Jenniskens SF, Beenen LF, van den Berg R, Koudstaal PJ, van Zwam WH, Roos YB, van der Lugt A, van Oostenbrugge RJ, Majoie CB, Dippel DW; MR CLEAN Investigators. A Randomized Trial of Intraarterial Treatment for Acute Ischemic Stroke. N Engl J Med. 2015;372:11-20.
- 10. Goyal M, Demchuk AM, Menon BK, Eesa M, Rempel JL, Thornton J, Roy D, Jovin TG, Willinsky RA, Sapkota BL, Dowlatshahi D, Frei DF, Kamal NR, Montanera WJ, Poppe AY,

- Ryckborst KJ, Silver FL, Shuaib A, Tampieri D, Williams D, Bang OY, Baxter BW, Burns PA, Choe H, Heo JH, Holmstedt CA, Jankowitz B, Kelly M, Linares G, Mandzia JL, Shankar J, Sohn SI, Swartz RH, Barber PA, Coutts SB, Smith EE, Morrish WF, Weill A, Subramaniam S, Mitha AP, Wong JH, Lowerison MW, Sajobi TT, Hill MD; ESCAPE Trial Investigators. Randomized Assessment of Rapid Endovascular Treatment of Ischemic Stroke. *N Engl J Med*. 2015;372:1019-30.
- 11. Campbell BC, Mitchell PJ, Kleinig TJ, Dewey HM, Churilov L, Yassi N, Yan B, Dowling RJ, Parsons MW, Oxley TJ, Wu TY, Brooks M, Simpson MA, Miteff F, Levi CR, Krause M, Harrington TJ, Faulder KC, Steinfort BS, Priglinger M, Ang T, Scroop R, Barber PA, McGuinness B, Wijeratne T, Phan TG, Chong W, Chandra RV, Bladin CF, Badve M, Rice H, de Villiers L, Ma H, Desmond PM, Donnan GA, Davis SM; EXTEND-IA Investigators. Endovascular therapy for ischemic stroke with perfusion-imaging selection. *N Engl J Med.* 2015;372:1009-18.
- 12. Saver JL, Goyal M, Bonafe A, Diener HC, Levy EI, Pereira VM, Albers GW, Cognard C, Cohen DJ, Hacke W, Jansen O, Jovin TG, Mattle HP, Nogueira RG, Siddiqui AH, Yavagal DR, Baxter BW, Devlin TG, Lopes DK, Reddy VK, de Rochemont RD, Singer OC, Jahan R; SWIFT PRIME Investigators. Stent-retriever thrombectomy after intravenous t-PA vs. t-PA alone in stroke. *N Engl J Med.* 2015;372:2285-95.
- 13. Jovin TG, Chamorro A, Cobo E, de Miquel MA, Molina CA, Rovira A, San Román L, Serena J, Abilleira S, Ribó M, Millán M, Urra X, Cardona P, López-Cancio E, Tomasello A, Castaño C, Blasco J, Aja L, Dorado L, Quesada H, Rubiera M, Hernandez-Pérez M, Goyal M, Demchuk AM, von Kummer R, Gallofré M, Dávalos A; REVASCAT Trial Investigators. Thrombectomy within 8 hours after symptom onset in ischemic stroke. *N Engl J Med*. 2015;372:2296-306.
- 14. Jauch EC, Saver JL, Adams HP Jr, Bruno A, Connors JJ, Demaerschalk BM, Khatri P, McMullan PW Jr, Qureshi AI, Rosenfield K, Scott PA, Summers DR, Wang DZ, Wintermark M, Yonas H; American Heart Association Stroke Council; Council on Cardiovascular Nursing; Council on Peripheral Vascular Disease; Council on Clinical Cardiology. Guidelines for the early management of patients with acute ischemic stroke: a guideline for health-care professionals from the American Heart Association/American Stroke Association. *Stroke*. 2013;44:870-947.
- 15. Powers WJ, Derdeyn CP, Biller J, Coffey CS, Hoh BL, Jauch EC, Johnston KC, Johnston SC, Khalessi AA, Kidwell CS, Meschia JF, Ovbiagele B, Yavagal DR; American Heart Association Stroke Council. 2015 American Heart Association/American Stroke Association Focused Update of the 2013 Guidelines for the Early Management of Patients With Acute Ischemic Stroke Regarding Endovascular Treatment: A Guideline for Healthcare Professionals From the American Heart Association/American Stroke Association. *Stroke*. 2015;46:3020-35.
- 16. Casaubon LK, Boulanger JM, Blacquiere D, Boucher S, Brown K, Goddard T, Gordon J, Horton M, Lalonde J, LaRivière C, Lavoie P, Leslie P, McNeill J, Menon BK, Moses B, Penn M,

Perry J, Snieder E, Tymianski D, Foley N, Smith EE, Gubitz G, Hill MD, Glasser E, Lindsay P; Heart and Stroke Foundation of Canada. Canadian Stroke Best Practice Recommendations: Hyperacute Stroke Care Guidelines, Update 2015. *Int J Stroke*. 2015;10:924-40.

17. Lavine SD, Cockroft K, Hoh B, Bambakidis N, Khalessi AA, Woo H, Riina H, Siddiqui A, Hirsch JA, Chong W, Rice H, Wenderoth J, Mitchell P, Coulthard A, Signh TJ, Phatorous C, Khangure M, Klurfan P, terBrugge K, Iancu D, Gunnarsson T, Jansen O, Muto M, Szikora I, Pierot L, Brouwer P, Gralla J, Renowden S, Andersson T, Fiehler J, Turjman F, White P, Januel AC, Spelle L, Kulcsar Z, Chapot R, Spelle L, Biondi A, Dima S, Taschner C, Szajner M, Krajina A, Sakai N, Matsumaru Y, Yoshimura S, Ezura M, Fujinaka T, Iihara K, Ishii A, Higashi T, Hirohata M, Hyodo A, Ito Y, Kawanishi M, Kiyosue H, Kobayashi E, Kobayashi S, Kuwayama N, Matsumoto Y, Miyachi S, Murayama Y, Nagata I, Nakahara I, Nemoto S, Niimi Y, Oishi H, Satomi J, Satow T, Sugiu K, Tanaka M, Terada T, Yamagami H, Diaz O,

Lylyk P, Jayaraman MV, Patsalides A, Gandhi CD, Lee SK, Abruzzo T, Albani B, Ansari SA, Arthur AS, Baxter BW, Bulsara KR, Chen M, Delgado Almandoz JE, Fraser JF, Heck DV, Hetts SW, Hussain MS, Klucznik RP, Leslie-Mawzi TM, Mack WJ, McTaggart RA, Meyers PM, Mocco J, Prestigiacomo CJ, Pride GL, Rasmussen PA, Starke RM, Sunenshine PJ, Tarr RW, Frei DF, Ribo M, Nogueira RG, Zaidat OO, Jovin T, Linfante I, Yavagal D, Liebeskind D, Novakovic R, Pongpech S, Rodesch G, Soderman M, terBrugge K, Taylor A, Krings T, Orbach D, Biondi A, Picard L, Suh DC, Tanaka M, Zhang HQ. Training Guidelines for Endovascular Ischemic Stroke Intervention: An International Multi-Society Consensus Document. *AJNR Am J Neuroradiol*. 2016;37:E31-4.

18. Widimsky P, Doehner W, Diener HC, Van Gelder IC, Halliday A, Mazighi M, on behalf of the ESC Council on Stroke. The role of cardiologists in stroke prevention and treatment: position paper of the European Society of Cardiology Council on Stroke. *Eur Heart J*, ehx478. 28 August 2017.