EuroCTO Club 2018 meeting: "Experts Live" in Toulouse



Nicolas Boudou¹, MD; Alexandre Avran², MD; Roberto Garbo³, MD; Thierry Lefevre⁴, MD; David Hildick-Smith⁵, MD, FRCP; Nicolaus Reifart⁶, MD; Alfredo Galassi⁷, MD; Alessio Mattesini⁸, MD; Carlotta Sorini Dini⁸, MD; Emmanouil Brilakis⁹, MD, PhD; Masahisa Yamane¹⁰, MD; George Sianos¹¹, MD, PhD; Kambis Mashayekhi¹², MD; Dimitrios Karmpaliotis¹³, MD, PhD; Tony DeMartini¹⁴, MD; Gerard S. Werner¹⁵, MD; Carlo Di Mario^{8*}, MD, PhD

1. CHU Rangueil, Toulouse University Hospital, Toulouse, France; 2. Division of Cardiology, Marseille University Hospital, Marseille, France; 3. Division of Cardiology, San Giovanni Bosco Hospital Torino, Turin, Italy; 4. Institut Cardiovasculaire Paris Sud, Massy, France; 5. Sussex Cardiac Centre, Brighton and Sussex University Hospitals, Brighton, United Kingdom; 6. Department of Cardiology, Krankenhaus Bad Soden, Bad Soden, Germany; 7. Division of Cardiology, University of Catania, Catania, Italy; 8. Interventional Structural Cardiology, Careggi University Hospital, Florence, Italy; 9. Minneapolis Heart Institute, Minneapolis, MN, USA; 10. Cardiology Department, Saitama Sekishinkai Hospital, Saitama, Japan; 11. Division of Cardiology, Thessaloniki University Hospital, Thessaloniki, Greece; 12. Department of Cardiology and Angiology II, University Heart Center Freiburg - Bad Krozingen, Bad Krozingen, Germany; 13. Division of Cardiology, Columbia University Medical Center, New York, NY, USA; 14. Prairie Cardiovascular, Springfield, IL, USA; 15. Division of Cardiology, Klinikum Darmstadt, Darmstadt, Germany

The first EuroCTO Club meeting after the new Eucomed rules became effective was a success, a testimony to the motivation of the 250 active physicians participating, mainly members of the Club, benefiting only in part from the educational grants provided by the generous contribution of the industry. The congress scientific programme always starts with a review of the recent CTO literature, this time including four randomised trials. The absence of changes in magnetic resonance imaging (MRI)-measured left ventricular ejection fraction (LVEF) and end-diastolic volume in the Evaluating XIENCE and Left Ventricular Function in Percutaneous Coronary Intervention on Occlusions After ST-Elevation Myocardial Infarction (EXPLORE) trial¹, randomising 304 patients with a CTO in a non-culprit vessel within a week after STEMI, was confirmed by another smaller randomised trial in a less acute setting, the Recovery of Left Ventricular Function After Stent Implantation in Chronic Total Occlusion of Coronary Arteries (REVASC CTO) study². Mashayekhi et al² found no differences in regional wall thickening, again measured with MRI, in

the 205 CTO patients randomised to receive or not receive revascularisation. In their original publication, the EXPLORE investigators suggested a possible improvement limited to patients with left anterior descending (LAD) occlusion. They now extend this to patients having good collateral flow to the CTO territory³, but still without clear evidence of clinical improvement. In the EXPLORE trial, anginal symptoms improved in the CTO group at one year but became similar at three years with a paradoxical worse cardiac mortality in the treated group (6.0% in the CTO PCI group vs. 1.0% in the CTO no PCI group, p=0.02)4. These results are totally at variance with the evidence confirmed in multiple trials, the last in the special population of 201 cardiogenic shock post-STEMI patients with non-culprit CTO included in the IABP-SHOCK II trial⁵, that CTO determines an independent mortality risk at 12 months after myocardial infarction (HR 1.30, 95% CI: 1.02-1.67, p=0.03). Also, the survivors to the acute phase have a worse prognosis in this trial substudy, with appropriate electrical shocks from the implantable cardioverter defibrillators (ICD) significantly more frequent in the

^{*}Corresponding author: Structural Interventional Cardiology Division, Careggi University Hospital, Largo G. Alessandro Brambilla, 3, 50134 Florence, Italy. E-mail: carlo.dimario@unifi.it

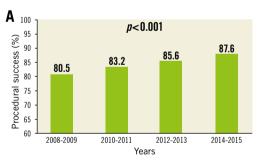
presence of a non-culprit CTO, a finding confirmed in the 240 patients with CTO out of the 720 patients enrolled in the eCTOpyin-ICD registry⁶. The favourable results both in terms of improved LVEF and dyspnoea in the low LVEF cohort of the 839-patient study conducted in Bad Krozingen and Zurich7 were discussed during the meeting by Galassi, who also elaborated on further indications for the use of LV assist devices during PCI. With 834 patients enrolled despite the premature termination, the Drug-Eluting Stent Implantation Versus Optimal Medical Treatment in Patients With Chronic Total Occlusion (DECISION-CTO) trial (ClinicalTrials.gov Identifier: NCT01078051) remains the largest randomised trial of CTO recanalisation (Park SJ. Presented at the American College of Cardiology Congress 2017, Washington DC, USA, March 2017). Its negative outcome (no difference in MACE in the intention-to-treat analysis) stunned the CTO operators at the time of presentation last year. Now we realise that the fundamental bias of allowing non-CTO artery treatment at the time of randomisation is in part responsible for an outcome totally different from the results of a countless number of registries comparing patients with failed vs. successful CTO PCI, the most recent being a registry of 5,496 patients from the PanLondon BCIS database⁸. The DECISION-CTO investigators rightly decided to present a per protocol analysis, considering a crossover of 78/398 (19.6%) from medical treatment to PCI, most of them in the first three days post randomisation, and a crossover in the opposite direction of 7.0% (Lee SW. DECISION-CTO: updated QoL and longterm outcomes. Presented at TCTAP 2018, Seoul, Korea, April 2018). Patients in these two groups had an incidence of combined MACE completely at variance with the incidence observed in the group of initial randomisation, partially invalidating the results of the intention-to-treat analysis. In the light of this high crossover rate, the lack of symptomatic improvement at one year initially reported can also be viewed differently, especially after new data showing that, at three to six and 12 months, all the functional indices improved in the successfully treated group. These results are at variance with all other reports including the marked improvement of quality of life at one month in the recently published individually monitored 1,000-patient Outcomes, Patient Health Status, and Efficiency in Chronic Total Occlusion Hybrid Procedures (OPEN CTO) registry^{9,10}. Martin-Yuste, the top enroller of the EuroCTO trial¹¹, reviewed its results. The recent publication of this trial, surely the greatest achievement of the EuroCTO Club with a highly significant improvement of angina frequency (5.23, 95% CI: 1.75-8.71; p=0.003) and quality of life (6.62, 95% CI: 1.78-11.46; p=0.007) at 12 months in the PCI group, certainly played a major role in eliminating restrictions to CTO recanalisation in the most recent European Society of Cardiology Guidelines on Myocardial Revascularization, with indications now only focused on symptoms and ischaemia, irrespective of lesion complexity¹². In a paragraph dedicated to CTO recanalisation, recommendations include the need to ensure viability in case of regional myocardial impairment, the discouragement to proceed ad hoc and the opportunity that these procedures be reserved to expert CTO operators.

Other highlights of the congress included an update on the CASTLE score¹³ (**Table 1**), an acronym for the five elements (previous CABG, age, stump, tortuosity, length, extent of calcification) affecting success in the EuroCTO registry, a report prior to publication of the trends in CTO recanalisation from the 2008-2015 registry¹⁴ (**Figure 1**), and many technical lectures on the characteristics and mode of use of recently introduced dedicated devices, from guidewires to microcatheters, dual lumen catheters,

Table 1. EuroCTO CASTLE score.

| EuroCTO CASTLE score | | |
|---|--------------------|--|
| Cases in derivation data set (n) | 14,882 | |
| Cases in validation data set (n) | 5,745 | |
| Endpoint | procedural success | |
| Variables in multivariate analysis (n) | 21 | |
| Score variables (n) | 6 | |
| Score range | 0-6 | |
| | Score | OR in multivariate |
| Score variables | points | analysis |
| Score variables CABG history | | |
| | points | analysis |
| CABG history | points 1 | analysis 1.42 (1.25-1.61) |
| CABG history Age (>70 years) | points 1 1 | analysis 1.42 (1.25-1.61) 1.28 (1.09-1.49) |
| CABG history Age (>70 years) Stump (blunt or invisible) | points 1 1 1 | analysis 1.42 (1.25-1.61) 1.28 (1.09-1.49) 1.64 (1.47-1.84) |
| CABG history Age (>70 years) Stump (blunt or invisible) Tortuosity degree (severe* or unseen) | points 1 1 1 1 | analysis 1.42 (1.25-1.61) 1.28 (1.09-1.49) 1.64 (1.47-1.84) 2.28 (2.00-2.60) |

*Severe tortuous CTO: either 2 or more pre-occlusive bends of >90° or at least 1 bend of >120°.



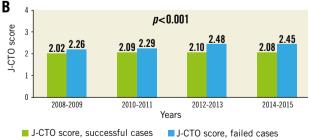


Figure 1. Procedural success trend in the EuroCTO Registry and the importance of the J-CTO score. A) Procedural success and B) J-CTO score. Adapted from the presentation by G. Sianos at EuroCTO 2018, Toulouse, France. https://www.eurocto.eu/events/it-IT/858/10th-experts-live-cto-workshop/presentation

trapping devices alone or in combination with guide extension, and low-profile balloons.

As in the last five years, some of the most experienced operators in the Club demonstrated new techniques and material during live cases, working with the support of colleagues from the Rangueil University Hospital coordinated by N. Boudou and D. Carrié (https://www.eurocto.eu/events/it-IT/858/10th-experts-live-ctoworkshop/video). The cases selected truly reflected the target audience - extremely complex CTOs requiring all the skills, tricks and attention of the operators to overcome truly ambiguous stumps, extreme tortuosity and calcification, very long occlusions, curly invisible septal and epicardial collaterals, very poor LVEF requiring Impella support during the procedure. The absence of complications in all cases and the success in 10/11 CTOs approached (90.9%) confirms the steady progress in this field. The EuroCTO Club meeting is unique because it offers the possibility to see at work US (Karmpaliotis, De Martini, Brilakis), Japanese (Yamane, Nasu) and European operators (Werner, Garbo, Boudou, Lefevre, Di Mario, Avran, Sianos, Galassi, Bufe, Mashayekhi, Ledermann, Levesque, Quillet, Hovasse, Drogoul, Tchetche, Weilenmann). In past editions differences were clear, with Japanese operators stressing careful wire manipulation and the importance of avoiding subintimal tracking or limiting it to the occluded segment, and US operators focusing on rapid change of procedural strategy following the hybrid algorithm, ready to move to anterograde dissection re-entry (ADR) or retrograde strategies for long lesions or ambiguous caps. This year the general impression was that the best of the two worlds joined in this European blend, taking full advantage of the extraordinary steerability of modern dual-core guidewires but also accepting knuckling and other aggressive strategies in the worst anatomies. Very trackable microcatheters, dual lumen catheters parked in a side branch originating at the occlusion site to penetrate the occlusion or re-enter a side branch at the distal end or facilitate the parallel wire technique, trapping balloons and combined trapping-guide extension devices were certainly instrumental in reducing the procedure duration and minimising contrast and X-ray exposure (no procedure went above 350 ml and 3 Gy).

EuroCTO Experts Live will be back in Berlin on 13-14 September 2019, to showcase once again the new advances offered by the science and technique of CTO recanalisation.

Conflict of interest statement

R. Garbo is a consultant for Terumo, Volcano and IMDS. D. Hildick-Smith is an advisor/speaker for Abbott, Boston, Medtronic and Terumo. The other authors have no conflicts of interest to declare.

References

1. Henriques JP, Hoebers LP, Råmunddal T, Laanmets P, Eriksen E, Bax M, Ioanes D, Suttorp MJ, Strauss BH, Barbato E, Nijveldt R, van Rossum AC, Marques KM, Elias J, van Dongen IM, Claessen BE, Tijssen JG, van der Schaaf RJ; EXPLORE Trial Investigators. Percutaneous Intervention for Concurrent Chronic

Total Occlusions in Patients With STEMI: The EXPLORE Trial. J Am Coll Cardiol. 2016;68:1622-32.

- 2. Mashayekhi K, Nührenberg TG, Toma A, Gick M, Ferenc M, Hochholzer W, Comberg T, Rothe J, Valina CM, Löffelhardt N, Ayoub M, Zhao M, Bremicker J, Jander N, Minners J, Ruile P, Behnes M, Akin I, Schäufele T, Neumann FJ, Büttner HJ. A Randomized Trial to Assess Regional Left Ventricular Function After Stent Implantation in Chronic Total Occlusion: The REVASC Trial. *JACC Cardiovasc Interv.* 2018;11:1982-91.
- 3. van Dongen IM, Elias J, van Houwelingen KG, Agostoni P, Claessen BEPM, Hoebers LP, Ouweneel DM, Scheunhage EM, Delewi R, Piek JJ, Råmunddal T, Laanmets P, Eriksen E, Bax M, Suttorp MJ, van der Schaaf RJ, Tijssen JGP, Henriques JPS. Impact of collateralisation to a concomitant chronic total occlusion in patients with ST-elevation myocardial infarction: a subanalysis of the EXPLORE randomised controlled trial. *Open Heart.* 2018;5: e000810
- 4. Elias J, van Dongen IM, Råmunddal T, Laanmets P, Eriksen E, Meuwissen M, Michels HR, Bax M, Ioanes D, Suttorp MJ, Strauss BH, Barbato E, Marques KM, Claessen BEPM, Hirsch A, van der Schaaf RJ, Tijssen JGP, Henriques JPS, Hoebers LP; EXPLORE investigators. Long-term impact of chronic total occlusion recanalisation in patients with ST-elevation myocardial infarction. *Heart*. 2018;104:1432-8.
- 5. Saad M, Fuernau G, Desch S, Eitel I, de Waha S, Pöss J, Ouarrak T, Schneider S, Zeymer U, Thiele H. Prognostic impact of non-culprit chronic total occlusions in infarct-related cardiogenic shock: results of the randomised IABP-SHOCK II trial. *EuroIntervention*. 2018;14:e306-13.
- 6. van Dongen IM, Yilmaz D, Elias J, Claessen BEPM, Delewi R, Knops RE, Wilde AAM, van Erven L, Schalij MJ, Henriques JPS. Evaluation of the Impact of a Chronic Total Coronary Occlusion on Ventricular Arrhythmias and Long-Term Mortality in Patients With Ischemic Cardiomyopathy and an Implantable Cardioverter-Defibrillator (the eCTOpy-in-ICD Study). *J Am Heart Assoc.* 2018 May 2;7(10).
- 7. Galassi AR, Boukhris M, Toma A, Elhadj Z, Laroussi L, Gaemperli O, Behnes M, Akin I, Lüscher TF, Neumann FJ, Mashayekhi K. Percutaneous Coronary Intervention of Chronic Total Occlusions in Patients With Low Left Ventricular Ejection Fraction. *JACC Cardiovasc Interv.* 2017;10:2158-70.
- 8. Jones DA, Rathod KS, Pavlidis AN, Gallagher SM, Astroulakis Z, Lim P, Sirker A, Knight CJ, Dalby MC, Malik IS, Mathur A, Rakhit R, Redwood S, MacCarthy PA, Baker C, Desilva R, Di Mario C, Weerackody R, Hill J, Wragg A, Smith EJ. Outcomes after chronic total occlusion percutaneous coronary interventions: an observational study of 5496 patients from the Pan-London CTO Cohort. *Coron Artery Dis.* 2018;29:557-63.
- 9. Sapontis J, Salisbury AC, Yeh RW, Cohen DJ, Hirai T, Lombardi W, McCabe JM, Karmpaliotis D, Moses J, Nicholson WJ, Pershad A, Wyman RM, Spaedy A, Cook S, Doshi P, Federici R, Thompson CR, Marso SP, Nugent K, Gosch K, Spertus JA, Grantham JA. Early Procedural and Health Status Outcomes After

Chronic Total Occlusion Angioplasty: A Report From the OPEN-CTO Registry (Outcomes, Patient Health Status, and Efficiency in Chronic Total Occlusion Hybrid Procedures). *JACC Cardiovasc Interv.* 2017;10:1523-34.

- 10. Riley RF, Sapontis J, Kirtane AJ, Karmpaliotis D, Kalra S, Jones PG, Lombardi WL, Grantham JA, McCabe JM. Prevalence, predictors, and health status implications of periprocedural complications during coronary chronic total occlusion angioplasty. *EuroIntervention*. 2018;14:e1199-206.
- 11. Werner GS, Martin-Yuste V, Hildick-Smith D, Boudou N, Sianos G, Gelev V, Rumoroso JR, Erglis A, Christiansen EH, Escaned J, di Mario C, Hovasse T, Teruel L, Bufe A, Lauer B, Bogaerts K, Goicolea J, Spratt JC, Gershlick AH, Galassi AR, Louvard Y; EUROCTO trial investigators. A randomized multicentre trial to compare revascularization with optimal medical therapy for the treatment of chronic total coronary occlusions. *Eur Heart J.* 2018:39:2484-93.
- 12. Neumann FJ, Sousa-Uva M, Ahlsson A, Alfonso F, Banning AP, Benedetto U, Byrne RA, Collet JP, Falk V, Head SJ, Jüni P, Kastrati A, Koller A, Kristensen SD, Niebauer J, Richter DJ, Seferovic PM, Sibbing D, Stefanini GG, Windecker S, Yadav R,

- Zembala MO; ESC Scientific Document Group. 2018 ESC/EACTS Guidelines on myocardial revascularization. *Eur Heart J.* 2019;40: 87-165.
- 13. Szijgyarto Z, Rampat R, Werner GS, Ho C, Reifart N, Lefevre T, Louvard Y, Avran A, Kambis M, Buettner HJ, Di Mario C, Gershlick A, Escaned J, Sianos G, Galassi A, Garbo R, Goktekin O, Meyer-Gessner M, Lauer B, Elhadad S, Bufe A, Boudou N, Sievert H, Martin-Yuste V, Thuesen L, Erglis A, Christiansen E, Spratt J, Bryniarski L, Clayton T, Hildick-Smith D. Derivation and Validation of a Chronic Total Coronary Occlusion Intervention Procedural Success Score From the 20,000-Patient EuroCTO Registry: The EuroCTO (CASTLE) Score. *JACC Cardiovasc Interv.* 2019;12:335-42.
- 14. Konstantinidis NV, Werner GS, Deftereos S, Di Mario C, Galassi AR, Buettner JH, Avran A, Reifart N, Goktekin O, Garbo R, Bufe A, Mashayekhi K, Boudou N, Meyer-Geßner M, Lauer B, Elhadad S, Christiansen EH, Escaned J, Hildick-Smith D, Carlino M, Louvard Y, Lefèvre T, Angelis L, Giannopoulos G, Sianos G; Euro CTO Club. Temporal Trends in Chronic Total Occlusion Interventions in Europe. *Circ Cardiovasc Interv.* 2018;11:e006229.