Impact of first medical contact to revascularisation time on long-term clinical outcomes in ST-segment elevation myocardial infarction patients

Roux Olivier\textsuperscript{a}, Schweighauser Andreas\textsuperscript{b, c}, Schukraft Sara\textsuperscript{c}, Stauffer Jean-Christophe\textsuperscript{d}, Goy Jean-Jacques\textsuperscript{c}, Wenaweser Peter\textsuperscript{e}, Togni Mario\textsuperscript{c}, Windecker Stephan\textsuperscript{d}, Cook Stéphane\textsuperscript{e}, Arroyo Diego\textsuperscript{c}, Puricel Serban\textsuperscript{c}

\textsuperscript{a} Department of Cardiology, Stadtspital Triemli, Zurich, Switzerland
\textsuperscript{b} Department of Internal Medicine, Kantonsspital Chur, Switzerland
\textsuperscript{c} Department of Cardiology, University and Hospital Fribourg, Switzerland
\textsuperscript{d} Department of Cardiology, Inselspital, Bern, Switzerland

Summary

INTRODUCTION: We sought to identify predictors for a prolonged delay from first medical contact to revascularisation (FMC-R) in ST-segment elevation myocardial infarction (STEMI) patients at our institution and to assess the impact of a prolonged treatment delay on 3-year clinical outcome.

METHODS: EVALFAST is a prospective and retrospective registry enrolling all patients admitted directly from pre-hospital care to the catheterisation laboratory at Fribourg Hospital for suspected STEMI, starting in June 2008. Relevant patient and procedural data were collected retrospectively and prospectively. Clinical follow-up was performed by phone or clinic visit. Patients were divided into two groups: FMC-R interval <90 minutes (short) and FMC-R delay of ≥90 minutes (long). The primary clinical endpoint was major adverse cardiac events (MACE) at 3-year follow-up. Secondary clinical endpoints were all-cause death and peak creatine kinase levels. Clinical outcome was compared between the two patient groups (short vs long) using Cox regression analysis.

RESULTS: Of the 406 patients enrolled between 2008 and 2014, 187 (46\%) were treated with a short delay and 219 (54\%) with a long delay. Age at presentation was the only predictor associated with prolonged delay (per additional year: odds ratio 1.03, 95\% confidence interval 1.01–1.05, p = 0.001). The primary clinical endpoint occurred in 15\% (n = 28) of patients in the short group, and 25\% (n = 54) in the long group (p = 0.02). This difference was driven by higher rates of cardiac death (p = 0.08) and the need for repeat revascularisation (p = 0.11).

CONCLUSION: Increased age impacts the FMC-R delay in patients with STEMI. Patients with shorter treatment delays (<90 minutes after FMC) have significantly lower MACE rates at 3 years.

Keywords: ST-segment elevation myocardial infarction (STEMI), pre-hospital delay, revascularisation time, major adverse cardiac events

Introduction

Incidence rates of ST-segment elevation myocardial infarction (STEMI) are decreasing. The advent of primary percutaneous coronary intervention (PCI), potent platelet inhibitors and the implementation of STEMI networks have substantially decreased mortality after STEMI\textsuperscript{[1–3]}. However, in-hospital mortality remains high, at an estimated 4–12\%, and 1-year mortality as assessed in angiography registries is estimated at around 10\% \textsuperscript{[4–6]}.

Several factors may impact clinical outcomes after STEMI. Time delays are one factor that can be modified by healthcare providers. Based on data from large registries,
From 2008 to 2009, clopidogrel was administered as a secondary treatment was given as soon as STEMI was suspected. Patients with confirmed STEMI were transferred directly to the catheterisation laboratory of the University and Hospital Fribourg, bypassing the emergency room (ER). Medication and antiplatelet agent in addition to aspirin. From 2009, ticagrelor and prasugrel were available, and these were preferentially prescribed over clopidogrel [8, 9]. As the patients were enrolled between June 2008 and December 2014, primary PCI was performed according to the 2008 ESC recommendations for the management of acute myocardial infarction in patients presenting with persistent ST-segment elevation [10]. Manual thrombectomy was performed whenever deemed necessary by the interventional cardiologist [11].

Endpoints
The primary clinical endpoint was survival free from major adverse cardiac events (MACE), defined as the composite of cardiac death, nonfatal myocardial infarction and any unplanned revascularisation. Secondary endpoints were all-cause mortality, the individual components of the primary endpoint, peak creatine kinase (CK) and creatine kinase muscle brain type (CK-MB), and length of stay in days.

Definitions
The onset of pain was defined as the beginning of symptoms. FMC was defined as the first assessment by a medical professional. Revascularisation time was defined as the time point of the guidewire passage through the culprit lesion. Total ischaemic time was defined as the time from symptom onset to revascularisation. Death was classified as either cardiac or non-cardiac, according to the Academic Research Consortium definition [6]. Deaths that could not be classified were considered cardiac. Unplanned revascularisation was defined as any repeat percutaneous or surgical revascularisation, regardless of location. Stent thrombosis was either definite or probable, according to the Academic Research Consortium definition [6].

Statistical analysis
Analyses were performed using SPSS software 26.0 (SPSS Inc, Chicago, IL, USA) and STATA 14 MP. Continuous variables are expressed as mean ± standard deviation or median with interquartile range (IQR). Categorical variables are expressed as counts and percentages. For continuous variables, histograms were checked for normal distribution. For the univariate analysis, baseline and procedural characteristics as well as clinical outcome were compared between patients presenting a delay from first medical contact to revascularisation (FMC-R) of <90 minutes (short) and patients presenting a delay of ≥ 90 minutes (long) using a chi-square test for categorical variables, an unpaired t-test for continuous variables with a normal distribution, and non-parametric tests such as the Wilcoxon rank sum test or the Mann-Whitney U-test for continuous variables with a non-Gaussian distribution.

To find the correlation of normally distributed variables, the Pearson product-moment correlation coefficient was calculated. For variables with a non-Gaussian distribution, Spearman’s rank correlation coefficient was computed. Crude estimates of clinical outcomes were assessed using the Kaplan-Meier method. Cox regression was used to adjust for baseline imbalances between patients with an FMC-R interval <90 minutes and patients with an FMC-R delay ≥90 minutes. To identify independent predictors of
the occurrence of MACE, a statistical model was computed using Cox proportional hazards. All variables that showed a significant difference between the groups, as well as clinically relevant variables (age, gender, diabetes, treatment during working hours, three-vessel disease, presentation at ER of PCI centre, transfer from GP, total ischaemic time) were forced into the model. In an explorative analysis, we created age categories. Patients in the same decade were considered as belonging to the same age category. Patients <40 years were pooled into the same category, as were patients >90 years.

Results

A total of 447 patients were admitted with STEMI, of whom 406 were included in the present study. The patient flowchart is depicted in figure 1. Patients were divided into two groups according to their FMC-R delay: those treated after <90 minutes (“short” group, n = 187) and those with a delay ≥90 minutes (“long” group, n = 219). Baseline characteristics are presented in table 1. All patients were considered for primary PCI. No patient received thrombolysis. Mean age was 63 ± 12 years and was significantly lower in the short vs the long group (61 ± 12 vs 65 ± 12, p = 0.001), 75% (n = 306) were men (short: 78% (n = 145) vs long: 74% (n = 161), p = 0.36). Diabetes mellitus was present in 13% (n = 25) of patients in the “short” group and in 19% (n = 41) of patients in the “long” group (p = 0.14). Patients with previous myocardial infarction were equally distributed in both groups (short: 11% (n = 20) vs long: 19% (n = 22), p = 0.83). Patients admitted during working hours were less likely to be treated early (short: 41% (n = 76) vs long: 59% (n = 130), p <0.001).

Delays

Treatment delays are outlined in supplementary table S1 in appendix 1. Overall, the median FMC-R time was 94 minutes (IQR 71–120 minutes). The shortest FMC-R time was seen in patients presenting directly to the ER of the PCI centre (86 minutes [IQR 58–110 minutes]), whereas patients referred from their GP had the longest FMC-R interval (129 minutes [IQR 74–164 minutes]). Total ischaemic time was shortest in patients referred by emergency medical services (152 minutes [IQR 108–235 minutes]). Median total ischaemic time was significantly shorter in patients with a short compared to patients with a long FMC-R delay (148 minutes [IQR 110–220 minutes] vs 226 minutes [IQR 154–345 minutes], p <0.001).

Peak CK and length of stay

There were no differences in peak CK or CK-MB levels between the groups (table S2). Median length of stay was 5 days (IQR 2–7), and similar between short and long FMC-R delays.
Lesion and procedural characteristics

There were no differences in lesion or procedural characteristics between the two groups (table S3). Overall, 546 lesions were treated in 406 patients. Predilatation was performed in 438 cases (80%) (short vs long: n = 194 (79%) vs n = 244 (81%), p = 0.39). There were 404 (74%) complex lesions (short v. long: n = 185 (76%) vs n = 219 (73%), p = 0.51). Thrombus aspiration was performed in 210 (38%) lesions (short vs long: n = 103 (42%) vs n = 107 (36%), p = 0.13). Post-procedural TIMI 3 flow was present in 517 cases (94%) (short vs long: n = 234 (96%) and n = 283 (94%), p = 0.44).

Predictors for a prolonged FMC-R delay

Overall, binary logistic regression identified age as the only predictor of prolonged FMC-R delay (per additional year, odds ratio: 1.03, 95% confidence interval [CI]: 1.01 – 1.05, p = 0.001; table 2).

Clinical outcome

Clinical outcomes are provided in table 3. At 3 years, MACE rates were significantly higher in patients with “long” FMC-R intervals (24.6% vs 15.0%, hazard ratio [HR] 1.80, 95% CI 1.10–2.94; p = 0.02). This difference was driven by a higher rate of repeat revascularisation (14.2% vs 10.2%, HR 1.65, 95% CI 0.89–3.0; p = 0.11), and an increased incidence of cardiac death (7.8% vs 3.2%, HR 2.55, 95% CI 0.91–7.15; p = 0.08). Kaplan-Meier curves for the primary endpoint are presented in figure 2.

Table 1: Baseline patient characteristics.

| Variable                                | All patients (n = 406) | Short (n = 187) | Long (n = 219) | p-value |
|-----------------------------------------|------------------------|----------------|---------------|---------|
| Age (years), mean ± SD                  | 63 ± 12                | 61 ± 12        | 65 ± 12       | 0.001   |
| Male, n (%)                             | 306 (75)               | 145 (76)       | 161 (74)      | 0.36    |
| BMI (kg/m²), mean ± SD                  | 26.8 ± 4.1             | 26.6 ± 4.0     | 27.0 ± 4.1    | 0.27    |
| Diabetes mellitus, n (%)                | 66 (16)                | 25 (13)        | 41 (19)       | 0.14    |
| Smoking, n (%)                          | 181 (45)               | 86 (46)        | 95 (43)       | 0.62    |
| Hypertension, n (%)                     | 190 (47)               | 92 (49)        | 98 (45)       | 0.37    |
| Dyslipidemia, n (%)                     | 153 (38)               | 69 (37)        | 84 (38)       | 0.76    |
| Family history, n (%)                   | 94 (23)                | 42 (23)        | 52 (24)       | 0.76    |
| Previous MI, n (%)                      | 42 (10)                | 20 (11)        | 22 (10)       | 0.83    |
| Previous PCI, n (%)                     | 59 (15)                | 30 (16)        | 29 (13)       | 0.42    |
| Previous CABG, n (%)                    | 8 (2)                  | 2 (1)          | 6 (3)         | 0.23    |
| Renal failure, n (%)                    | 17 (4)                 | 7 (4)          | 10 (5)        | 0.68    |
| Admitted during working hours, n (%)    | 208 (51)               | 76 (41)        | 130 (59)      | <0.001  |
| Cardiogenic shock, n (%)                | 21 (5)                 | 10 (5)         | 11 (5)        | 0.88    |
| Coronary artery disease                 |                        |                |               |         |
| – Single-vessel, n (%)                  | 160 (39)               | 78 (42)        | 82 (37)       | 0.38    |
| – Two-vessel, n (%)                     | 135 (34)               | 64 (34)        | 71 (32)       | 0.70    |
| – Three-vessel, n (%)                   | 111 (27)               | 45 (24)        | 66 (30)       | 0.17    |
| Culprit lesion                          |                        |                |               |         |
| – Left main, n (%)                      | 3 (1)                  | 2 (1)          | 1 (1)         | 0.48    |
| – LAD, n (%)                            | 179 (44)               | 84 (45)        | 95 (43)       | 0.76    |
| – LCX, n (%)                            | 50 (12)                | 20 (11)        | 30 (14)       | 0.34    |
| – RCA, n (%)                            | 170 (42)               | 81 (43)        | 89 (41)       | 0.64    |
| Provenance                              |                        |                |               |         |
| – Walk-in to ER of PCI centre, n (%)    | 72 (18)                | 41 (22)        | 31 (14)       | 0.05    |
| – Transfer Riaz, n (%)                  | 56 (14)                | 25 (13)        | 31 (14)       | 0.81    |
| – Transfer Tafers, n (%)                | 40 (10)                | 19 (10)        | 21 (10)       | 0.84    |
| – Transfer Payeme, n (%)                | 56 (14)                | 22 (12)        | 34 (16)       | 0.27    |
| – Transfer Meyriez, n (%)               | 11 (3)                 | 3 (2)          | 8 (4)         | 0.20    |
| – Transfer Estavayer, n (%)             | 2 (1)                  | 1 (1)          | 1 (1)         | 0.90    |
| – Referral from GP, n (%)               | 22 (5)                 | 7 (4)          | 15 (7)        | 0.16    |
| – EMS, n (%)                            | 120 (30)               | 52 (28)        | 68 (31)       | 0.47    |
| – Other, n (%)                          | 9 (2)                  | 5 (3)          | 4 (2)         | 0.56    |
| – Unknown, n (%)                        | 16 (4)                 | 11 (8)         | 5 (2)         | 0.06    |

BMI = body mass index; CABG = coronary artery bypass graft; EMS = emergency medical services; ER = emergency room; GP = general practitioner; LAD = left anterior descending artery; LCX = left circumflex artery; MI = myocardial infarction; PCI = percutaneous coronary intervention

Table 2: Predictors for first medical contact to revascularisation >90 minutes.

| Variable                                | Odds ratio (95% confidence interval) | p-value |
|-----------------------------------------|--------------------------------------|---------|
| Walk-in to ER of PCI centre             | 0.61 (0.36–1.03)                     | 0.06    |
| Age (per additional year)               | 1.03 (1.01–1.05)                     | 0.001   |

ER = emergency room; PCI = percutaneous coronary intervention
One patient in the “short” group died 607 days after STEMI from sudden death at home, which was considered a cardiac death. One patient in the “long” group died suddenly 233 days after STEMI, which was also considered a cardiac death. No other ambiguous or otherwise unclassified deaths occurred.

Information on all-cause mortality is also provided in Table 3. Mortality at 1 year was higher in the “long” than in the “short” group (8.2% vs 2.1%, HR 3.90, 95% CI 1.33–11.5; p = 0.01). However, at the 3-year follow-up, the difference in mortality was no longer statistically significant (11.9% vs 7.0%, HR 1.79, 95% CI 0.92–3.48; p = 0.08; fig. 2).

Discussion

The main findings of the present study are [1]: age negatively impacts the delay between first medical contact and revascularisation in STEMI patients [2]; the system-dependent delay in our center complies with the new ESC quality target in 50% of patients [3]; patients revascularised within 90 minutes after first medical contact have lower MACE rates at 3 years.

Predictors

We identified age as the only significant predictor for a treatment delay exceeding 90 minutes. Likewise, in a registry including 3832 patients, Ruiz et al. also identified age as an independent predictor for an FMC-R delay of >120 minutes [11]. This may be due to several factors. Firstly, atypical symptoms are more frequent with advancing age, thus rendering the diagnosis of STEMI more difficult [12]. Moreover, communication between the patient and the first medical responder may be impaired in the elderly. Secondly, comorbidities and frailty might prolong the time from diagnosis to treatment [13]. Lastly, in the very old, conservative treatment instead of primary PCI may be considered before activating established STEMI network pathways. Increased treatment delays for patient subsets presenting with atypical symptoms have been found previously. Roswell and colleagues found prolonged FMC-R delays in women and explained their finding by postulating atypical symptoms as the cause of this association [14–18].

In analysing 14,518 US patients with first medical contact at non-PCI-capable hospitals and requiring transfer for primary PCI, Dauerman et al. found a prolonged system-dependent delay in patient subsets with a propensity for atypical clinical presentation (women, elderly and diabetics). Interestingly, other potent predictors for short treatment delays in their analysis were white race and higher annual STEMI volume [19].

With an increasingly elderly population, more elderly patients are likely to be treated for STEMI, increasing the risk of misdiagnoses and adverse clinical outcomes. Future clinical research in this elderly population should not only concentrate on medical parameters such as risk factors, Table 3: Primary clinical end point and overall mortality at 30 days, 1 year and 3 years according to treatment delay.

|                  | Total (n = 406) | Short (n = 187) | Long (n = 219) | HR (95% CI)        | p-value |
|------------------|----------------|----------------|---------------|-------------------|---------|
| 30-day MACE, n (%) | 23 (5.7)       | 5 (2.7)        | 18 (8.2)      | 2.61 (0.91–7.49)  | 0.07    |
| – Cardiac death, n (%) | 15 (3.7)       | 4 (2.1)        | 11 (5.0)      | 2.41 (0.63–7.29)  | 0.22    |
| – Nonfatal MI, n (%) | 1 (0.2)        | 0 (0.0)        | 1 (0.5)       | –                 | –       |
| – Repeat revascularisation, n (%) | 7 (1.7)        | 1 (0.5)        | 6 (2.7)       | 4.79 (0.46–49.59) | 0.19    |
| 1-year MACE, n (%)   | 55 (13.5)      | 16 (8.5)       | 39 (17.8)     | 2.05 (1.11–3.80)  | 0.02    |
| – Cardiac death, n (%) | 19 (4.7)       | 4 (2.1)        | 15 (6.9)      | 2.64 (0.84–8.26)  | 0.09    |
| – Nonfatal MI, n (%) | 3 (0.7)        | 0 (0.0)        | 3 (1.4)       | –                 | –       |
| – Repeat revascularisation, n (%) | 33 (8.1)       | 12 (6.4)       | 21 (9.6)      | 1.76 (0.82–3.81)  | 0.15    |
| 3-year MACE, n (%)    | 82 (20.2)      | 28 (15.0)      | 54 (24.6)     | 1.80 (1.10–2.94)  | 0.02    |
| – Cardiac death, n (%) | 23 (5.7)       | 6 (3.2)        | 17 (7.8)      | 2.55 (0.91–7.15)  | 0.08    |
| – Nonfatal MI, n (%) | 9 (2.2)        | 3 (1.6)        | 6 (2.7)       | 1.46 (0.31–6.86)  | 0.64    |
| – Repeat revascularisation, n (%) | 50 (12.3)      | 19 (10.2)      | 31 (14.2)     | 1.65 (0.89–3.04)  | 0.11    |
| 30-day mortality n (%) | 12 (3.0)       | 3 (1.6)        | 9 (4.1)       | 2.57 (0.69–9.51)  | 0.15    |
| 1-year mortality, n (%) | 22 (5.4)       | 4 (2.1)        | 18 (8.2)      | 3.9 (1.33–11.5)   | 0.01    |
| 3-year mortality, n (%) | 39 (9.6)       | 13 (7.0)       | 26 (11.9)     | 1.79 (0.92–3.48)  | 0.08    |

CI = confidence interval; HR = hazard ratio; MACE = major adverse cardiac events; MI = myocardial infarction

Figure 2: Kaplan-Meier curves for survival free from the occurrence of major adverse cardiac events and death.
clinical and angiographic presentations, management and prognosis, but also, and most importantly, on ethical considerations, including patient autonomy and expectations.

Delays
Our analysis shows that the ESC-recommended system-dependent treatment delay of <90 minutes was achieved in approximately 50% of patients treated between 2008 and 2014. The maximum expected delay before choosing PCI over fibrinolysis (120 minutes) was achieved in 75% of patients. Many PCI centres, regional STEMI networks and national societies have reported their system-dependent treatment delay, showing large differences. In a cross-sectional, multicentre study including seven PCI-capable hospitals in the Netherlands, Tra and colleagues reported reaching the guideline-recommended target of <90 minutes in 78.7% of patients [20]. However, they defined the beginning of the system-dependent delay as the time of the first electrocardiogram and not the time of first medical contact, which likely explains their overall performance. Our results are in line with the treatment delays reported in GRACE and the HORIZONS-AMI trial, where reperfusion delays from first medical contact or from first hospital door threshold of <90 minutes were achieved in roughly 50% of patients [21, 22]. Short system-dependent delays have been recommended for a long time [23]. Interestingly, the 2012 ESC Guidelines recommendations of delays <90 and at most <120 minutes are derived from a post-hoc analysis of the DANAMI-2 trial and an analysis of the National Cardiovascular Data Registry in the US [24, 25].

While both studies suggested that shorter time delays were beneficial in terms of clinical outcomes, the question of whether or not the recommended delays were feasible was not addressed. The median system-dependent delay in DANAMI-2 was 127 minutes (IQR 98–157 minutes) for patients undergoing primary PCI. Nonetheless, achieving optimal treatment delays should be the ultimate objective of every STEMI network. Comparisons between countries, geographical regions, healthcare systems and cultures may be subject to biases. It is unclear whether there would be much room for improvement in extensive rural areas, remote from primary PCI-capable centres, in which access to primary healthcare is restricted. This may, in part, explain data from the United States, where only minimal improvements in treatment delays were observed from the late 1990s to the mid-2000s [26, 27]. Indeed, feasibility seems to be a key issue in this context. The overall treatment delay from symptom onset to reperfusion is dependent on a large number of variables. The patient-dependent component, governing the time from symptom onset to first medical contact, could possibly be modulated through public service announcements; these would have to be provided repeatedly. Geographic location, i.e., the distance to the nearest healthcare provider, seems to play an important role and is likely an unchangeable factor. The system-dependent component should be constantly analysed. Malfunctioning processes should be amended based on thorough analyses. Future research might focus on STEMI networks that perform well and on trying to understand whether the key reasons for a network’s success are transferrable to other systems with distinct socio-geographical attributes.

Clinical outcomes
It is generally accepted that total ischaemic time and age are some of if not the most influential predictors for clinical outcome after STEMI. There is ample evidence of fewer mid- and long-term MACE in patients with shorter door-to-balloon or FMC-R times [24, 25]. In this analysis, we found significantly lower rates of MACE in patients with an FMC-R delay <90 minutes compared to patients with a delay exceeding 90 minutes. This difference was largely driven by lower rates of cardiac death occurring within the first year. At 3 years, the overall MACE rate was 20% and all-cause mortality was 9.6% for patients treated in the STEMI network of Fribourg.

Data on 3-year mortality after STEMI are limited. Hosseiny and colleagues recently reported an overall 3-year mortality rate of 11.6% in 1313 Australian STEMI patients [28]. All-cause mortality at 1 year has been reported at 11.4% in 2804 consecutive patients treated with primary PCI for STEMI in Denmark [4]. In contrast, 1-year mortality in our registry was 5.4%. This difference might be explained by the fact that patients in the Danish registry were included between 1998 and 2008, with inherent differences in the medical devices used (e.g., PCI with bare metal or first-generation drug-eluting stents). One-year all-cause mortality in the SCAAR study was reported at 9.4% between 2009 and 2010 [29]. In 2017, Garcia et al. reported a 2-year mortality of 8.5% in 1268 Catalan patients treated between 2002 and 2013 [30].

Limitations
Our study has several limitations. The data collection from 2008 to 2010 was retrospective and may be subject to misclassification or information bias. Information on first medical contact was collected retrospectively and was missing in a minority of patients. Given our limited sample size, the statistical power for identifying predictors for a long FMC-R delay was likely low.

The external validity and general transferability of our results are limited, as geographical, political and social factors are intertwined, and the results likely reflect the care provided in a small, high-income country with universal access to health care.

Conclusion
Increased age prolongs the delay to revascularisation after first medical contact in patients with STEMI. Patients with treatment delays of <90 minutes have significantly lower MACE rates at 3 years.

Financial disclosure
This trial was an investigator-initiated study supported by an unrestricted grant from the Fonds Scientifique Cardiovasculaire (Fribourg, Switzerland). The funding sources had no role in the design of the study, data collection, data monitoring, data analysis, data interpretation, or writing of the report.

Potential competing interests
The authors declare no conflict of interest.

References
1 Puymirat E, Simon T, Steg PG, Schiele F, Guéret P, Blanchard D, et al.; USIK USIC 2000 Investigators; FAST MI Investigators. Association of changes in clinical characteristics and management with improvement in
Appendix 1

**Supplementary data**

Table S1: Provenance and delay.
Table S2: Peak CK and length of stay.
Table S3: Lesion and procedural characteristics.
Table S4: FMC-R delay according to age category.

Figure S1: Distribution of age according to short or long treatment delays.
Figure S2: Proportion of patients with long treatment delays for each age category.

The appendix is available as a separate file at https://smw.ch/article/doi/smw.2020.20368.