Long-Term Efficacy of a Mini-Course in Radiation-Reducing Techniques in Invasive Cardiology

Zusammenfassung

Material und Methoden: Vor, 2 Monate und 2 Jahre nach dem Minikurs (Periode I, II und III) wurden folgende Dosisparameter analysiert, erhoben durch 7 Kardiologen im Verlauf von je 10 Koronarangiografien: Dosisflächenprodukt (DFP), radiografischer und fluoroskopischer DFP-Anteil, Anzahl radiografischer Bilder und Serien, und Durchleuchtungszeit.

Ergebnisse: In Periode I und II betrug das mediane Patienten-DFP 31,4 bzw. 15,8 Gy × cm² und sank bis Periode III nochmals auf 8,5 Gy × cm². Dieser Langzeiteffekt war verbunden mit einer kürzeren medianen Durchleuchtungszeit (180, 172 bzw. 120 Sekunden), weniger (12, 12, 10) und kürzeren medianen Durchleuchtungszeiten (180, 172 bzw. 120 Sekunden), weniger (12, 12, 10) und kürzeren medianen Durchleuchtungszeiten (180, 172 bzw. 120 Sekunden), weniger (12, 12, 10) und kürzeren medianen Durchleuchtungszeiten (180, 172 bzw. 120 Sekunden).

Die multivariate Regressionsanalyse bestätigte die zunehmende Effektivität des Minikurses von Periode II (−44,6%) auf III (−60,7%). Gleichzeitig sank der Einfluss der interventionellen Erfahrung der Untersucher (−8,6% bzw. −4,9% pro 1000 Koronarangiografien, jeweils durchgeführt bis zum Kurs). Die Anzahl der Koronarangiografien nach dem Kurs beeinflusste die Langzeitergebnisse nicht.

Schlussfolgerung: Das vorgestellte Kursprogramm ermöglicht im Langzeitverlauf eine signifikante, nachhaltige und fortschreitende Reduktion der Patientendosis in der diagnostischen Koronarangiografie. Eine selbstkontrollierende Dokumentation relevanter Dosisparameter ist bestens geeignet, individuelle strahlenhygienische Bemühungen der Untersucher zu überwachen und zu verbessern.

Abstract
Purpose: To validate the long-term efficacy of a 90-min, educational mini-course in less-irradiating cardiac interventional techniques.

Materials and Methods: Before, two months after, and two years after the mini-course (periods I, II, and III), we analyzed the following radiation dose parameters for ten coronary angiographies (CA), performed by each of 7 cardiologists: total dose-area product (DAP), radiographic and fluoroscopic DAP fractions, number of radiographic frames and runs, and fluoroscopy time.

Results: The median patient DAP for periods I, II, and III was 31.4, 15.8 and 8.5 Gy × cm², respectively. The long-term effect was related to shorter median fluoroscopy times (180, 172, and 120 s), shorter (57, 52, and 45) and fewer (12, 12, and 10) radiographic runs, consistent collimation and restriction to an adequate image quality. Both radiographic DAP/frame (28.7, 17.0, and 18.4 mGy × cm²) and fluoroscopic DAP/second (45.7, 24.2, and 10.0 mGy × cm²) decreased significantly. The multivariate linear regression analysis confirmed the increasing efficacy of the mini-course itself (−44.6% and −60.7%), and revealed a decreasing influence of the interventionalist’s experience (−8.6% and −4.9% per 1,000 CAs, lifelong performed until the mini-course). The number of CAs performed after the mini-course did not influence the long-term DAP results.

Conclusion: The presented educational mini-course allows a significant, long-lasting, and apparently ongoing reduction of patient radiation exposure due to CA. A self-surveillance documentation of relevant radiation parameters is well suited to monitor and improve each operator’s individual long-term radiation-reducing efforts.
Introduction

The first warnings regarding the "strongly delayed appearance of serious effects" of radiation exposure on tissue were published as early as 1896 [1]. In 1925, Groedel, the founder of both the German Cardiac Society and the American College of Cardiology, summarized the following factors as playing deleterious roles in conjunction with X-ray applications: excessively long fluoroscopic examinations, short focus-skin distances, insufficient filtering of the X-ray beam, and excessive numbers of radiographs made by insufficiently trained operators [2]. These problems have apparently not yet been entirely resolved in invasive cardiology. Current publications from the International Commission on Radiation Protection (ICRP) point out that unfortunately many interventionists are not aware of the potential for injury from cardiac procedures, their occurrence, or the simple methods for decreasing their incidence by utilizing dose control strategies [3].

The mean dose-area product (DAP) received by patients from coronary angiography (CA) and percutaneous coronary intervention (PCI) is high and typically varies to an extremely great degree in invasive cardiology: i.e., 16–106 Gy × cm² and 34–191 Gy × cm², respectively [4, 5]. Actual median (mean) reported levels for DAPs of 741,238 procedures registered from 842 German catheterization laboratories in 2011 were 23.0 (31.3) Gy × cm² for CA and 50.3 (65.9) Gy × cm² for combined PCI [6]. A DAP of 100 Gy × cm² applied in one constant tube angulation to an area of the patient's skin of approximately 40 cm² will induce a calculated entrance skin erythema (>2 Gy) [7] and even deep-skin and musculocutaneous injury (>15 Gy) in conjunction with complex coronary interventions may result [3, 8, 9]. Recent results suggest that hazards include altered cellular redox balance [10], somatic DNA damage [10, 11], and radiation-induced cancer [12, 13] in patients and chronically exposed physicians. In an international survey the average number of cases of radiation-induced cancer per million CAs was calculated as 280 [14]. For these reasons we have developed and validated effective dose-reduction techniques [4, 15] and, without negative implications on diagnostic accuracy and procedural outcome, have achieved a mean patient DAP of 4.2 Gy × cm² for elective CA [16] and 6.7 Gy × cm² for one-vessel PCI [17], equivalent to effective doses of 0.8 and 1.3 mSv, respectively [5].

To "Encourage Lower-Irradiation Cardiac Interventional Techniques" was the primary goal of our ELICIT initiative: the presented mini-course in radiation-reducing techniques that has proved successful and has been validated as effective in the clinical routine [18]. Two years after this training program, we investigated the long-term efficacy of this mini-course and analyzed independent influencing factors such as operators’ interventional experience and patient body mass index (BMI), sex, and age.

Methods

Definitions

Entrance skin dose (unit: Gray [Gy] = J/kg) is the dose to air in the entrance plane of the patient including backscatter and represents the most appropriate quantity for the characterization of deterministic skin lesions. The dose area product (DAP) is the product of the mean dose in air in a given plane perpendicular to the central beam and the irradiated area at this plane (unit: Gy × cm²) and is independent of the distance from the X-ray source. The effective dose (ED) is the sum of all weighted dose equivalents of exposed organs in the body (unit: Sievert [Sv]) and characterizes future stochastic cancer risks. DAP-to-ED conversion factors have been calculated to range between 0.18 and 0.22 mSv/Gy × cm² for the thoracic region in adults [5].

Study design and recommendations for radiation-reducing techniques

All seven interventionists of the Cardiology Center of Greifswald University, Germany, participated. In study period I, each of them consecutively performed ten elective CAs by femoral access and documented the following radiation dose parameters: total DAP, radiographic (DAPΓ) and fluoroscopic (DAPp) fractions, fluoroscopy time, and the number of radiographic frames and runs. DAPΓ/ frame and DAPp/second(s) were calculated to demonstrate the efficacy of collimation to the region of interest. Bypass graft controls, significant valve diseases, and emergency catheterizations were excluded, since they are inhomogeneous [16, 17]. The interactive oral PowerPoint mini-course, a standardized 90-minute workshop, addressed the following dose-reduction principles [4, 15, 18]:

- Essential radiographic runs and frames
- Consistent collimation, i.e. to the ostial region of interest during intubation of coronary arteries
- Fluoroscopy-free collimation or intermittent fluoroscopy by short pedaling
- Adequately low-level acquisition modes
- Less irradiating angulations
- Reduced magnification and full inspiration during radiography whenever possible
- Long source-to-skin and short patient-to-detector distances
- Sufficiently well-rested operators

Periods II and III consisted of repeated measurements approximately two months and two years after the mini-course, with short-term and long-term median (interquartile ranges) results of 67 (54–88) and 654 (603–711) days, respectively (Table 1). All patients and interventionists were encoded. Each interventionist estimated the number of lifelong CAs performed prior to the mini-course as a parameter of his/her individual interventional experience (Fig. 1), used the same equipment before and after the mini-course, received feedback in writing about his/her individual results at baseline, but obtained no further in-house training.

Equipment

We used a digital, single-arm undercouch tube flat panel system (AXIOM Artis dFC; Siemens AG, Erlangen, Germany) with an anti-scatter grid and a 2.9-mm aluminum filter equivalent installed in the X-ray beam. Based on the previous fluoroscopic image, an automatic algorithm determined the thickness of additional variable copper filters within the range of 0.2 to 0.9 mm for fluoroscopy and 0.0 to 0.9 mm for radiography. With a focus-image detector distance of 1.2 m, a 25-mm thick aluminum absorber and a 20-cm detector field format, the detector entrance dose rates were 0.029 μGy/pulse (15 pulses/s) for fluoroscopy and 0.170 μGy/frame for radiographic documentation (15 frames/s). The DAP was measured by a light transparent ionization chamber (K1–S Axiom, PTW, Freiburg, Germany), which was calibrated in-situ. Uncertainty varied between 0% in 90° left lateral oblique (LAO) angulation and <10% overestimation of the real DAP in postero-anterior (PA) angulation (total uncertainty <15%). All technical devices remained unchanged during the study, i.e. did not receive any upgrade with respect to radiation dose parameters.
Values were expressed as median and interquartile range (IQR: 25–75th percentile). Three groups (periods [p] I, II, and III) of non-normally distributed variables were analyzed by the Kruskal-Wallis test. A two-tailed p-value of < 0.05 was regarded as significant to prove the variables to be different. Paired comparisons (period II vs. I, III vs. II, and III vs. I) of significant results were performed using the Mann-Whitney U test. In this case, a Bonferroni adjustment of significance level was required, i.e., p-values lower than 0.05/3 were considered statistically significant. Analyses were performed using the software SAS (V 9.1, SAS Institute Inc., Cary, NC, USA). For categorical data, frequencies were compared by the χ² test. To rule out biasing effects, we performed multivariate linear regression analysis to adjust for patient age, sex, and BMI, as well as for the operator’s interventional experience prior to and two years after the mini-course. The various periods were coded dichotomously for the short-term (period I vs. period II) and long-term (period I vs. period III) analyses. Normal distribution of residuals was tested using the Shapiro-Wilk test at a significance level of 0.05/3 level (Mann-Whitney U test and Bonferroni adjustment) and *0.05 level (χ² test).

### Table 1

<table>
<thead>
<tr>
<th>patient dose parameters</th>
<th>II vs. I</th>
<th>III vs. II</th>
<th>III vs. I</th>
</tr>
</thead>
<tbody>
<tr>
<td>operators [n]</td>
<td>7</td>
<td>7</td>
<td>7</td>
</tr>
<tr>
<td>patients [n]</td>
<td>70</td>
<td>70</td>
<td>70</td>
</tr>
<tr>
<td>age [years]</td>
<td>63.3 (52.6 – 68.9)</td>
<td>64.9 (54.3 – 70.3)</td>
<td>65.4 (56.5 – 71.4)</td>
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<tr>
<td>female [%]</td>
<td>34.3</td>
<td>41.4</td>
<td>52.9</td>
</tr>
<tr>
<td>BMI [kg/m²]</td>
<td>27.1 (24.9 – 30.0)</td>
<td>27.6 (24.6 – 30.1)</td>
<td>26.8 (24.2 – 30.4)</td>
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<tr>
<td>DAP [Gy × cm²]</td>
<td>31.4 (20.0 – 40.6)</td>
<td>15.8 (10.3 – 22.9)</td>
<td>8.5 (6.3 – 13.0)</td>
</tr>
<tr>
<td>DAPR [Gy × cm²]</td>
<td>19.7 (13.0 – 27.8)</td>
<td>11.1 (6.6 – 15.4)</td>
<td>7.5 (4.8 – 11.7)</td>
</tr>
<tr>
<td>DAPF [Gy × cm²]</td>
<td>8.0 (5.0 – 13.0)</td>
<td>4.5 (2.0 – 9.5)</td>
<td>1.3 (0.7 – 2.0)</td>
</tr>
<tr>
<td>fluoroscopy time [s]</td>
<td>180 (126 – 306)</td>
<td>172 (120 – 324)</td>
<td>120 (82 – 174)</td>
</tr>
</tbody>
</table>

### Statistical analysis

Values were expressed as median and interquartile range (IQR: 25–75th percentile). Three groups (periods [p] I, II, and III) of non-normally distributed variables were analyzed by the Kruskal-Wallis test. A two-tailed p-value of < 0.05 was regarded as significant to prove the variables to be different. Paired comparisons (period II vs. I, III vs. II, and III vs. I) of significant results were performed using the Mann-Whitney U test. In this case, a Bonferroni adjustment of significance level was required, i.e., p-values lower than 0.05/3 were considered statistically significant. Analyses were performed using the software SAS (V 9.1, SAS Institute Inc., Cary, NC, USA). For categorical data, frequencies were compared by the χ² test. To rule out biasing effects, we performed multivariate linear regression analysis to adjust for patient age, sex, and BMI, as well as for the operator’s interventional experience prior to and two years after the mini-course. The various periods were coded dichotomously for the short-term (period I vs. period II) and long-term (period I vs. period III) analyses. Normal distribution of residuals was tested using the Shapiro-Wilk test at a significance level of 0.05/3 level (Mann-Whitney U test and Bonferroni adjustment) and *0.05 level (χ² test).
Specific influence of the short-term results

<table>
<thead>
<tr>
<th>Parameter</th>
<th>mini-course</th>
<th>p-value</th>
<th>operator's experience</th>
<th>p-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>DAP</td>
<td>−44.6 %</td>
<td>&lt;0.0001</td>
<td>−8.6 %</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>DAP R</td>
<td>−47.5 %</td>
<td>&lt;0.0001</td>
<td>−7.7 %</td>
<td>&lt;0.0001</td>
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<tr>
<td>DAP F</td>
<td>−44.7 %</td>
<td>&lt;0.0001</td>
<td>−12.2 %</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>Radiographic frames</td>
<td>−10.5 %</td>
<td>0.0072</td>
<td>−4.9 %</td>
<td>0.0170</td>
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<tr>
<td>Frames/run</td>
<td>−11.4 %</td>
<td>&lt;0.0001</td>
<td>−1.0 %</td>
<td>0.0012</td>
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<td>Fluoroscopy time</td>
<td>+0.5 %</td>
<td>0.9660</td>
<td>−2.0 %</td>
<td>0.2905</td>
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<tr>
<td>DAP R/frame</td>
<td>−41.3 %</td>
<td>&lt;0.0001</td>
<td>−2.0 %</td>
<td>0.0549</td>
</tr>
<tr>
<td>DAP F/frame</td>
<td>−43.0 %</td>
<td>&lt;0.0001</td>
<td>−2.0 %</td>
<td>0.0549</td>
</tr>
</tbody>
</table>

Long-term results

<table>
<thead>
<tr>
<th>Parameter</th>
<th>p-value</th>
</tr>
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<tr>
<td>DAP</td>
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<td>DAP R</td>
<td>0.0186</td>
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<tr>
<td>DAP F</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>Radiographic frames</td>
<td>&lt;0.0001</td>
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<tr>
<td>Frames/run</td>
<td>0.0791</td>
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<tr>
<td>Fluoroscopy time</td>
<td>0.0071</td>
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<tr>
<td>DAP R/frame</td>
<td>0.0026</td>
</tr>
<tr>
<td>DAP F/frame</td>
<td>0.6825</td>
</tr>
</tbody>
</table>

BMI: body mass index; DAP: dose area product; DAP R/F: radiographic, fluoroscopic DAP.

Effects of interventional experience, body mass index, age, and sex

The calculated multivariate linear regression model for the short-term efficiency of the mini-course yielded results according the following equation between the natural logarithm (ln) of the dependent variable – for example, the DAP – and the independent influencing factors, i.e., the period ([p]: 0 = period I, 1 = period II), the operator’s lifelong experience ([e]: per 1,000 CAs), patient body mass index ([b]: kg/m²), patient sex ([s]: 0 = female, 1 = male) and patient age ([a]: years):

\[
\ln (\text{DAP}) = 1.003 + (-0.591 \times p) + (-0.00009 \times e) + (0.077 \times b) + (0.203 \times s) + (0.005 \times a)
\]

The natural logarithm of DAP for a 54-year-old male patient (BMI 26.4 kg/m²) and for operator’s lifelong experience of 1,000 CAs was 3.393 and 2.803 for periods I and II, respectively, equivalent to the expected absolute values for DAP of 29.7 Gy × cm² (exp[3.393]) and 16.5 Gy × cm² (exp[2.803]). Under conditions of constancy of all other variables, the patient DAP for a CA performed in period II was exp(−0.591) = 0.554 (−44.6 %), compared to the DAP in period I, indicating the adjusted relative short-term efficacy of our mini-course (Table 2, upper panel). The DAP likewise slightly varied according to exp(−0.009) = 0.914 (−8.6 %) per 1,000 CAs of individual experience (Table 2, upper panel) to exp(−0.0077) = 1.080 (−8.0 %) per one unit of kg/m² BMI and to exp(−0.203) = 1.225 (−22.5 %) for men compared to women.

Adjusted for the operators’ experience and patient sex, BMI and age, the patient DAP for a CA performed in period III was exp(−0.933) = 0.393 (−60.7 %), compared to the DAP in period I. Likewise, the mini-course yielded a significant beneficial long-term effect on all parameters associated with radiation dose (Table 2, lower panel). The operator’s interventional experience, characterized by the number of lifelong CAs performed prior to the mini-course, in ad-
dition to the efficacy of the mini-course indeed slightly influenced various dose parameters. Per 1,000 CAs performed prior to the mini-course, the presented regression model revealed long-term decreases of 4.9%, 3.9%, 8.6%, 3.9%, and 3.9% for DAP, radiographic DAP, fluoroscopic DAP, radiographic frames, and fluoroscopy time, respectively (\( \text{Table 2} \)). Our multivariate regression analysis revealed that the latter long-term DAP results were not significantly influenced by the number of CAs performed after the mini-course. All variables for DAP and for dose intensity, but neither the number of radiographic frames and runs nor the fluoroscopy time, were positively associated with patient BMI.

Discussion

Our mini-course in radiation-reducing techniques in cardiology is the first such effort to be validated in the clinical routine with respect to long-term efficacy. Without further in-hospital training, the reduction of the median patient DAP was not merely significant and lasting, but apparently ongoing. Whereas the short-term efficacy [18, 19] was predominantly due to consistent collimation, the long-term benefits moreover included significant improvements in all radiation-reducing techniques (\( \text{Table 1} \)). Avoidance of lengthy filming – i.e., fewer and shorter (median length 3.0 s) radiographic runs – was an essential step towards significant dose reduction. Bypass grafts, late-filling collateral pathways, and low-flow phenomena, however, should be documented by sufficiently longer series. The interventionists indeed maintained the improved short-term collimation to the region of interest (\( \text{Table 1} \)), i.e., the median fluoroscopic DAP second continued to decrease from 24.2 to 10.0 mGy × cm\(^2\). In accordance with current ICRP recommendations [3], the interventionists obviously reduced the fluoroscopic pulse rate and/or the detector entrance dose level towards an adequate instead of the left main stem and the left anterior descending artery in the typical LAO views for the documentation of the bifurcations of the left main stem and the left anterior descending artery in favor of less irradiating caudal and cranial PA views [4, 15]. As recommended in our course program, interventionists should therefore avoid the typical LAO views for the documentation of the bifurcations of the left main stem and the left anterior descending artery.

The multivariate regression models for both short-term and long-term analyses revealed further details. As expected, the BMI, in a range between 7.5 and 99.9 kg/m\(^2\), is correlated positively with all DAP fractions, but not with the number and length of radiographic runs. Female gender is correlated to lower DAP\(^2\) frame, which could be interpreted to result from the lower overall bone mass of women and their tendency to postmenopausal osteoporosis.

Over and above the predominant short-term and long-term efficacy of the mini-course, which, according our multivariate linear regression analysis, enabled adjusted DAP reductions of 44.6% and 60.7% (\( \text{Table 2} \)), the operator’s interventional experience also slightly but significantly influenced all DAP fractions and the number of radiographic frames and runs. Per 1,000 CAs individually performed lifelong prior to the mini-course, all DAP fractions decreased between 7.7% and 12.2% two months after and between merely 3.9% and 8.6% to baseline two years after the interventional training (\( \text{Table 2} \)). The operators’ experience after the mini-course apparently did not influence the patient DAP. Whereas the short-term effect of the operators’ lifelong experience manifested itself in terms of fewer radiographic frames and better collimation, an additional but merely slight long-term effect was evidently shorter fluoroscopy times. The presented short-term results accordingly confirm that the predominance of fluoroscopy time as the widely considered criterion for operators’ efforts in radiation-reducing techniques should decrease as a logical consequence of the fact that approx. 65 – 70% of the total DAP for a CA is due to radiographic acquisition [4, 17].

The presented mini-course in radiation-reducing techniques resulted in a considerable long-term reduction in patient DAP in the course of elective CA, predominantly due to the efficacy of the mini-course itself (\( \text{Table 2} \)). The resulting median DAP of 8.5 GY × cm\(^2\) is far below typical levels of 23.0 GY × cm\(^2\) according to the National German Quality Registry [6]. With the use of DAP-to-ED conversion factors of approximately 0.20 mSv/GY × cm\(^2\), the biologically relevant patient ED decreased in the course of our presented ELICIT study from 4.6 mSv to 1.7 mSv [5]. The additional individual lifetime attributable risk (LAR) of cancer from a coronary intervention generating 40 mSv (equivalent to 200 GY × cm\(^2\)) is approximately 0.20 to 1.00% [8]. ED values of multi-slice computed tomography coronary angiography (CTCA) with the use of retrospective ECG-gating – representing noninvasive CA – are reported to vary from 8 to 28 mSv [4, 20, 21] and yield the considerable calculated LAR of cancer incidence for a representative 50-year-old subject with a range between 0.10 and 0.17% for males and between 0.23 and 0.37% for females [22]. The use of automatic dose control reduced the ED in chest scans by 36% [23], and in conformity with actual recommendations [24], ECG-triggered, single-and two-heartbeat CTCA enabled lower EDs of 2.2 to 8.6 mSv, without affecting diagnostic performance [21, 25–27]. For the specific subgroup of non-obese patients with a low and stable heart rate, furthermore, values even on the submillisievert level for CTCA have been achieved [28]. ED determined from Monte Carlo simulations of chest CT, however, were reported to underestimate in-vivo ED substantially for both volume and helical CTCA [25]. Several limitations to this study are worthy of discussion. The lower-irradiating tube angulations recommended in our course program were proved effective merely for the specific intervention of elective coronary angiography. They need to be questioned and undeniably are not transferable to the broad variety of fluoroscopy-guided radiological interventions at diverse vascular regions and organs of interest all over the human body. For iliac, femoral and peripheral vascular interventions, it is a consistent postero-anterior (PA) tube angulation that ensures patient diameters as low as reasonably achievable and consequently should be favored over both cranio-caudal PA views and oblique lateral views. Alterations of tube angulation during interventions at hepatic vessels or the biliary system will not considerably reduce the time-adjusted radiation intensity. In addition, the DAP-to-ED conversion factors for the abdominal and pelvis regions are higher than presented here for the cardiac region, and amount to approx. 0.26 and 0.29 mSv/GY × cm\(^2\), respectively [29]. Furthermore, all coronary angiographies were performed using a modern flat panel catheterization system of one manufacturer, which, in contrast to former image intensifier systems, allowed variable pulse rates and detector entrance dose levels in the clinical routine.
The presented mini-course in radiation-reducing techniques in cardiology is the first validated effort that allows in the clinical routine a significant, lasting, and apparently ongoing reduction of patient radiation exposure due to invasive CA to levels far below typical values in modern catheterization laboratories and comparable to or even below those achieved by new techniques in CTCAs. The documentation of DAPρ and DAPρβ and of the number of radiographic frames in addition to fluoroscopy time and total DAP is well suited to monitor – by self-surveillance or by supervisory authorities – each operator’s individual long-term efforts to decrease radiation exposure. The individual strategy of our mini-course conclusively allows considerable occupational dose reductions, confirms recent recommendations on training and supervision in those techniques [8, 30, 31], and indeed promises to be more efficient than current directives for simple documentation and regulative control of upper-limit reference levels for DAP and fluoroscopy time at cardiac centers [6].

References

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