Original paper

Improvement of early detection of breast cancer through collaborative multi-country efforts: Medical physics component


ARTICLE INFO

Keywords:
IAEA
Mammography
Breast cancer
Quality control

ABSTRACT

Purpose: The International Atomic Energy Agency (IAEA) through a Coordinated Research Project on “Enhancing Capacity for Early Detection and Diagnosis of Breast Cancer through Imaging”, brought together a group of mammography radiologists, medical physicists and radiographers; to investigate current practices and improve procedures for the early detection of breast cancer by strengthening both the clinical and medical physics components. This paper addresses the medical physics component.

Methods: The countries that participated in the CRP were Bosnia and Herzegovina, Costa Rica, Egypt, India, Kenya, the Frmr. Yug. Rep. of Macedonia, Mexico, Nigeria, Pakistan, Philippines, Slovenia, Turkey, Uganda, United Kingdom and Zambia. Ten institutions participated using IAEA quality control protocols in 9 digital and 3 analogue mammography equipment. A spreadsheet for data collection was generated and distributed. Evaluation of image quality was done using TOR MAX and DMAM2 Gold phantoms.

Results: QC results for analogue equipment showed satisfactory results. QC tests performed on digital systems showed that improvements needed to be implemented, especially in thickness accuracy, signal difference to noise ratio (SDNR) values for achievable levels, uniformity and modulation transfer function (MTF). Mean glandular dose (MGD) was below international recommended levels for patient radiation protection. Evaluation of image quality by phantoms also indicated the need for improvement.

Conclusions: Common activities facilitated improvement in mammography practice, including training of medical physicists in QC programs and infrastructure was improved and strengthened; networking among medical physicists and radiologists took place and was maintained over time. IAEA QC protocols provided a uniformed approach to QC measurements.


© 2018 Associazione Italiana di Fisica Medica. Published by Elsevier Ltd. This is an open access article under the CC BY-NC-ND license (http://creativecommons.org/licenses/BY-NC-ND/4.0/).
1. Introduction

In 2012, there were an estimated 1670000 cases of breast cancer diagnosed worldwide [1] making breast cancer the second most common cancer in the world and the most common cancer in women. According to the International Agency for Research on Cancer (IARC), over half of breast cancers occurred in the less developed countries. Incidence rates vary between regions, with the lowest rate in Central Africa (27/100000) and the highest in Belgium (111.9/100000). Deaths from breast cancer are disproportionately higher in less developed countries with 62% of total deaths occurring in these countries. As countries develop breast screening programs it is likely that breast cancer incidence rates will increase due to its early detection.

At the moment, X-ray mammography is the only technique that has proven the ability to detect breast cancer at an early stage, before the cancer is palpable and is the basis of the most organized breast screening programs to detect breast cancer in a non-symptomatic population. For mammography to be effective at detecting breast cancer at an early stage, adequate differentiation of small masses and microcalcifications is required, which in principle can only produce subtle contrast differences in mammography images. These imaging requirements place high technical demands on the imaging equipment and require high image quality and rigorous Quality Assurance (QA) to maintain these standards. Additional constraints are placed as the radiation dose should be kept at the lowest possible level, given the size average breast composition or average breast glandularity of the non-symptomatic target group, and the radiosensitivity of the breast. Routine performance testing of mammography imaging equipment by competent medical physicists is an essential component of a comprehensive QA program for mammography screening [2,3]. Additionally, it is apparent that all involved medical professionals have to be properly trained and highly acquainted with the mammographic procedure.

In 2012, the International Atomic Energy Agency (IAEA) started a Coordinated Research Project (CRP): IAEA CRP E1.30.39 Enhancing Capacity for Early Detection and Diagnosis of Breast Cancer through Imaging, which grouped together mammography radiologists, medical physicists and radiographers from 15 different countries. During the 4-year period of the project many activities were undertaken with the intention to investigate current practices, aiming to improve early detection of breast cancer by strengthening both the clinical and medical physics components. In this context, as part of the CRP activities participants received additional training in several components of the QA process. The overall objective of the CRP, from the Medical Physics perspective, was to contribute to the improvement in diagnosis and detection of breast cancer following the application of international standards of best practice in mammography. The CRP activities were specifically designed to: familiarize participants with the assessment of image quality and dosimetry requirements for mammography; training medical physicists and radiographers on performing measurements and collecting data, improving the provision of QA processes; collecting comprehensive Quality Control (QC) results in participating institutions using the relevant IAEA protocols; analyzing and evaluating these QC results and comparing them with internationally established requirements and tolerances for corrective actions [2,4]; evaluating image quality in mammography units in a standardized way, using a common phantom and centralized analysis of the images, and finally create a network of medical physicists and radiologists that can support each other on the technical aspects of mammography.

There are a number of differences between the IAEA protocol and that of the European protocol. The main difference is that in the IAEA approach, measurement of contrast detail performance uses the same test object, but with an automated reading system, which is how many individuals apply the EC test object/protocol these days. In addition, there are slight differences in the specification of the position for the test object for assessment of SNR values, but which would not cause there to be a significant difference between the IAEA protocol and that of the EC. The main difference is that the specifications for MTF measurements is more detailed in the IAEA protocol than in the EC’s one.

2. Materials and methods

2.1. Participants

Countries participating in the IAEA CRP had very different levels of implementation of their breast screening programs and large deviations in the available mammography equipment and corresponding conformance with established quality assurance programs. To remove this local bias in terms of the level of QA implementation, and standardize the practices, all groups followed a common methodology for the QC test. IAEA Human Health Series No.2 [4] and No.17 [2] were agreed as references (for screen-film and digital mammography, respectively), allowing uniform collections of basic QC metrics and assessment of the performance of participating mammography equipment. A total of 15 countries participated in different phases of the CRP (Bosnia and Herzegovina, Costa Rica, Egypt, India, Kenya, the Fmr. Yug. Rep. of Macedonia, México, Nigeria, Pakistan, Philippines, Slovenia, Turkey, Uganda, United Kingdom, Zambia), whereas 9 agreed to take part in this equipment testing inter-comparison. Table 1 shows information about participating institutes in terms of mammography equipment (analogous/digital), institution, existence of established screening program at the commencement of the CRP, number of clinical procedures during 2015 and implementation of organized QC programs before and after this CRP. Analogous equipment with its screen/film combination that participated in the study were: Planned Nuance Classic (CAWO MAMMO R200/Kodak MIN R), Siemens Balance (Kodak MIN R 2000/AGFA HT) and Metaltronica Flat SE (Agfa HD Mamory/Coordinated Research Project (CRP): IAEA CRP E1.30.39 Enhancing Capacity for Early Detection and Diagnosis of Breast Cancer through Imaging, which grouped together mammography radiologists, medical physicists and radiographers from 15 different countries. During the 4-year period of the project many activities were undertaken with the intention to investigate current practices, aiming to improve early detection of breast cancer by strengthening both the clinical and medical physics components. In this context, as part of the CRP activities participants received additional training in several components of the QA process. The overall objective of the CRP, from the Medical Physics perspective, was to contribute to the improvement in diagnosis and detection of breast cancer following the application of international standards of best practice in mammography. The CRP activities were specifically designed to: familiarize participants with the assessment of image quality and dosimetry requirements for mammography; training medical physicists and radiographers on performing measurements and collecting data, improving the provision of QA processes; collecting comprehensive Quality Control (QC) results in participating institutions using the relevant IAEA protocols; analyzing and evaluating these QC results and comparing them with internationally established requirements and tolerances for corrective actions [2,4]; evaluating image quality in mammography units in a standardized way, using a common phantom and centralized analysis of the images, and finally create a network of medical physicists and radiologists that can support each other on the technical aspects of mammography.

There are a number of differences between the IAEA protocol and that of the European protocol. The main difference is that in the IAEA approach, measurement of contrast detail performance uses the same test object, but with an automated reading system, which is how many individuals apply the EC test object/protocol these days. In addition, there are slight differences in the specification of the position for the test object for assessment of SNR values, but which would not cause there to be a significant difference between the IAEA protocol and that of the EC. The main difference is that the specifications for MTF measurements is more detailed in the IAEA protocol than in the EC’s one.

2. Materials and methods

2.1. Participants

Countries participating in the IAEA CRP had very different levels of implementation of their breast screening programs and large deviations in the available mammography equipment and corresponding conformance with established quality assurance programs. To remove this local bias in terms of the level of QA implementation, and standardize the practices, all groups followed a common methodology for the QC test. IAEA Human Health Series No.2 [4] and No.17 [2] were agreed as references (for screen-film and digital mammography, respectively), allowing uniform collections of basic QC metrics and assessment of the performance of participating mammography equipment. A total of 15 countries participated in different phases of the CRP (Bosnia and Herzegovina, Costa Rica, Egypt, India, Kenya, the Fmr. Yug. Rep. of Macedonia, México, Nigeria, Pakistan, Philippines, Slovenia, Turkey, Uganda, United Kingdom, Zambia), whereas 9 agreed to take part in this equipment testing inter-comparison. Table 1 shows information about participating institutes in terms of mammography equipment (analogous/digital), institution, existence of established screening program at the commencement of the CRP, number of clinical procedures during 2015 and implementation of organized QC programs before and after this CRP. Analogous equipment with its screen/film combination that participated in the study were: Planned Nuance Classic (CAWO MAMMO R200/Kodak MIN R), Siemens Balance (Kodak MIN R 2000/AGFA HT) and Metaltronica Flat SE (Agfa HD Mamory/
Agfa Mamoray HDR & Fuji AD-M or KODAK Min R 2190/Fuji AD-M & Agfa Mamoray HDR). No Computed Radiology equipment participated in this study. Digital equipment (FFDM) that was studied: GE Senographe Essential, GE Senographe 2000D, Siemens Inspiration, Hologic Selenia and Fuji Amulet s.

2.2. Testing protocols

In order to harmonize the application of the testing protocols, training was provided to the associated staff during a CRP meeting in Slovenia in November 2013, where medical physicists, radiographers and radiologists were present. During a review meeting in Vienna in 2016 the results obtained by the participants were reviewed and discussed, and the common spreadsheet was also reviewed and fine-tuned.

To demonstrate the process and the benefit of the CRP in the clinical practice, relevant QC tests both for analogue and digital equipment were agreed to be performed within each institution twice; one in the early phases of the CRP and one near completion.

Participating institutes with analogue equipment were asked to submit information for the following:

- Kilovoltage accuracy and radiation output at 28 kVp,
- Automated exposure control (AEC) performance,
- Half value layer (HVL at 28 kVp),
- Mean glandular dose (MGD) and
- Film processor sensitometry.

Since many countries were new to digital QC, and in order to standardize the data collection process, the University of Costa Rica (UCR) put together a spreadsheet for data collection based on the IAEA methodology [2]. By means of built-in macros, the spreadsheet automatically checks for tolerances and limiting values, facilitating easy evaluation of performance and compliance of the CRP participants. The CRP team created a group in a popular mobile application, allowing almost real-time support and guidance to the participants, led by the medical physics personnel from the UCR.

The spreadsheet, that automatically generates a printable report with the main results, covered the following tests, (shown in parenthesis are the sections from the protocol):

- Unit assembly (8.2.1),
- Geometry: compression force and thickness accuracy indicator (8.3.1) and collimation system (8.9),
- X Ray tube: half value layer (8.7.1),
- Automatic exposure control performance: SNR and SDNR (8.4),
- Image quality: spatial linearity and geometric distortion (8.5.3),
- Ghosting (8.5.4), uniformity and artefact evaluation (8.5.5) and modulation transfer function (8.6.1),
- Dosimetry: mean glandular dose (8.8.1).

Main points to mention are: test object for SDNR was a 0,2 mm thick square of aluminium of 10 mm on a side placed on a 20 mm PMMA slab, MTF test tool was placed on top of 45 mm PMMA and for dosimetry mean glandular dose is obtained from incident kerma in air using the formula used in Europe and United Kingdom with variable glandular content. Detector response function and noise characteristics were not evaluated in this study.

2.3. Image quality assessment

For the evaluation of the image quality, two different types of comprehensive phantoms were procured by the IAEA and distributed to all participants under the framework of CRP project, depending on the available equipment:

Analogical equipment: the TOR MAX phantom (Fig. 1) by Leeds Test Objects [5], was utilized for the evaluation of sensitometry (ten-step grey-scale), high contrast resolution (1,0 to 20,0 lp/mm), low contrast resolution (1,8–5 lp/mm), low-contrast large-detail detectability (12 details, 5,6 mm diameter) and high-contrast small-detail detectability (11 objects, 0,5 mm and 0,25 mm diameter); analysis of the TOR MAX image was performed by local physicists.

Digital equipment: image quality for digital units is commonly defined in terms of the threshold detectability of small gold discs within a contrast-detail mammography phantom, such as the CDMM [6] used in EUREF protocols [7]. In this research project the DMAM2 Gold phantom was used [8] (Fig. 2) containing 78 threshold contrast details in 6 sizes, with diameters ranging from 0,1 mm to 2,0 mm and with contrast range from 0,49% to 27,56%. This phantom also meets the standards for the European Guidelines for quality assurance in breast cancer screening and diagnosis [3].

To avoid ambiguity in the analysis of the images generated by the participants, DMAM2 Gold images were analysed centrally at the UCR using a dedicated software provided by the phantom vendor, AutoPIA (Automatic Phantom Image Analysis) by Cyberqual Srl (Gorizia, IT), which automatically localizes phantom details [9]. Software validation has been documented [10].

The phantom was placed between two slabs of 20 mm PMMA and total thickness was 53 mm. Exposure for each unit was made with automatic exposure control (technique factors for each equipment are listed on Table 2). A total of 16 “raw” (or “unprocessed”) images, were collected for each equipment and sent to the UCR for AutoPIA software analysis.

Once the software automatically recognizes the phantom type, it localizes the groups of details within the image and the individual details within each group. For each detail, the image quality is evaluated utilizing the contrast-to-noise ratio (CNR) defined, according to the Rose’s model, as the signal difference between the detail of interest and the surrounding background, divided by the noise of the background [11]. For “large-area” details, the software calculates the CNR based on a circular ROI inside the detail from which it calculates the mean pixel value to get the detail signal, then with the use of a circular ring surrounding the detail, it calculates from it the mean pixel value and standard deviation to get the background signal and noise, respectively. For “small details” (0,5 mm and 0,25 mm) the signal produced by the detail is calculated by weighting the individual differences between the pixel value of each pixel belonging to the detail and the surrounding background [12].

Analysis of the modulation transfer function (MTF) has also been carried out centrally by UCR medical physicists, using ImageJ plug in for performing physical characterization and quality checks known as “COQ” [15,16].

2.4. Dosimetry

To assess the mean glandular dose (MGD), the methodology of Dance et al. [17] was utilized, as described in IAEA publications [2,13,14] using the following equation:
where, $K_{ci,t}$ is the incident air kerma at each PMMA phantom thickness, the $g_{t}$-factor converts air kerma to MGD for a breast of glandularity of 50% and the $c_{t}$-factor corrects for the difference in breast composition from 50% glandularity and finally, s is the factor which corrects for the target filter combination of the X-ray tube.

3. Results and discussion

Table 3, shows the results for the 3 analogue mammography equipment from the annual QC test for 2013 and 2015. Facilities reported implementation of sensitometry QC; general parameters like kVp accuracy, output and HVL were within recommended tolerances in all systems. MGD was also below the recommended 2.5 mGy value for 45 mm PMMA in all systems, except in system A1 in 2015.

Tables 4a, 4b and 4c; show the initial results for the QC metrics that allowed the CRP group to evaluate the overall performance of the 9 digital mammography systems that participated in this study. It should be noted that system D2, does not meet the standard for some indices, is not being used clinically.

In Fig. 4, the Figure of Merit (FOM), as the ratio of SDNR squared divided by MGD, permits a quantitative assessment of the overall performance in terms of image quality and patient doses [18].

The spatial resolution properties of participating digital imaging systems are shown by means of the MTF graphs in Fig. 5. COQ Plug in
Note the achievable level. The squares associated for this detail, which was then compared with the acceptable and visible detail identified by the software gave the gold thickness associated with green colour show until which value (Nyquist) frequency, not allowing determination of MTF at 20%, for units D3 and D4 both from same manufacturer. Most systems did not pass the recommended values given in Human Health No.17 publication, this illustrates one of the weaknesses in using MTF as a QA measure.

Finally, image quality based on threshold contrast visibility was evaluated with the DMAM2 Gold phantom and software. The last visible detail identified by the software gave the gold thickness associated for this detail, which was then compared with the acceptable and achievable values of section 2b.2.4 from European guidelines [3].

Fig. 6, shows for each equipment the performance for objects of different sizes. The squares filled with green colour show until which value the system was able to resolve. The acceptable and achievable values are marked with solid and dash lines respectively. Results show that only systems D2, D3, D4 and D9 performed according to the acceptable value of threshold contrast visibility. Optimization of image quality can be implemented in all equipment since very few have green squares that run until the dash line (achievable value). It is to be noted that for the object with diameter 0,5; equipment D1, D5, D6, D7 and D8 were not able to meet the standard for this size but were able to show better results for smallest objects.

After the presented results of the QC test were discussed among all participants and individual efforts and corrective actions were implemented in different facilities. The following list gives an overview of the main interventions that resulted from this common exercise.

- One of the analogue equipment with poor performance in this study was replaced by a digital one (not included in this paper). Additional training activities were organized with the active support of the radiologists and through IAEA assistance.
Fig. 3. Mean glandular dose (mGy) for 20, 45 and 70 mm PMMA.

Fig. 4. Figure of Merit for 20, 45 and 70 mm PMMA.

Fig. 5. Modulation transfer function.
were trained in quality control programs.

- Equipment A1 that performed poorly at the beginning was replaced.
- Equipment D3 even though it passed all QC tests, its detector was changed and overall image quality was improved.
- Equipment D5 had its detector changed and subsequently passed all tests.
- Equipment D6 had service intervention and issues in thickness accuracy and HVL were corrected.
- Equipment D7 showed artefacts with W/Ag combination while imaging thick breast and hence the Ag filter was replaced to improve overall image quality.

4. Conclusions

Improvement in terms of quality practices for mammography has commenced in some countries as a result of the medical physics activities of the CRP. Prior to the implementation of the CRP mammography QC programs among participating countries varied significantly. Analogue QC has been done for many years but the transition to digital mammography imposes new aspects and important challenges in the establishment of a complete QC program. Image quality metrics were the ones that showed the need for more attention, advocating the need for comprehensive quality assessment, rather than fragmented efforts of sporadic performance testing.

Human resources, especially concerning medical physicists, were limited at the beginning of the CRP, and in some countries, it was difficult to find and involve medical physicists. Various centres have addressed this need by training medical physicists from other disciplines within medical physics, with further specialized training on mammography occurring during the course of the CRP. Among the group, the radiologists’ understanding of the role and need for medical physicists has improved and the involvement of physicists in multidisciplinary teams has been strengthened. Radiographic QC practices in mammography were improved. Furthermore, as an additional benefit to the human resources and infrastructure improvement that resulted out of the CRP, networking among medical physicists and radiologists took place and was maintained over time at the national and international level.

The use of IAEA QC protocols provided a uniformed approach to QC measurements. The spreadsheet developed for the project allowed an adequate collection of data, as well as, a review against tolerances in a more expeditious and validated way. Centralized image analysis using computational tools also allowed greater uniformity of methodology and objectivity in results.
Regarding QC test results, accessing “raw” images proved to be a challenge at the beginning, while compliance with the limiting values has been an issue in some centres that was addressed through immediate corrective actions. It must be noted that the primary aim of mammography is to provide diagnostically adequate image quality for the purposes of the examination, and radiation protection has to be considered only when this requirement is met. However, the lack of medical physicist who can properly evaluate the image quality, in combination with a strong regulatory emphasis on radiation protection of patients, as reported in other studies [19], create a framework of very tightly supervised radiation dose in isolation from image quality, jeopardizing the diagnostic findings. Medical physicists and service personnel need to conduct optimization actions where doses can be increased to obtain better overall image quality and radiologists know that their subjective evaluation of image quality alone, is not enough to provide the optimum service to their patients.

Another important issue pointed out was that many countries do not have much experience on how to fix discrepancies outside routine service protocols; and many do not have a preventive maintenance contract for their mammography units.

Some of the elements of quality control that are presented in this study were completely missing in certain participating institutes and the application of more comprehensive testing methodologies provided an additional tool for improvement of the mammography QC practice. The results of this study represent a valuable confirmation of the need for comprehensive quality assurance programs in mammography, but also in diagnostic radiology. Sporadic performance testing of a limited number of parameters, such as kVp and HVL, sometimes only as part of the licensing process, are no longer adequate to investigate in depth the performance of modern mammography systems and point out possible improvements. The active involvement of the medical physicist in the design and the application of such QA programs is mandatory to decrypt all the information that can be derived from the QC testing, bringing added value to the clinical use of the systems and facilitating improved care for the patients.

Acknowledgements

Jerry Soto, Centro de Investigación en Ciencias Atómicas, Nucleares y Moleculares, Universidad de Costa Rica, Costa Rica.
Manindra Bhushan, Rajiv Gandhi Cancer Institute and Research Centre, Delhi, India.
Eduardo López-Pineda and César Ruiz-Trejo; Instituto de Física, Universidad Nacional Autónoma de México, Ciudad de México, México.
Sidrah Mahmood and Muhammad Arif, Multan Institute of Nuclear Medicine and Radiotherapy, Multan, Pakistan.
Zusana Ćosić, Nebojša Vulin, Igor Janković and Marko Arežina; University Clinical Centre of the Republic of Srpska, Department of Clinical Radiology, Banja Luka, Bosnia and Herzegovina.

CRP participants are thankful to our CRP coordinator, Dr. Ravi Kashyap, from the Nuclear Medicine and Diagnostic Imaging Section (NMDS) of the Division of Human Health at the International Atomic Energy Agency (IAEA) who passed away during the course of the project. His enthusiasm and motivation moved the entire group in achieving our initial goals.

Spreadsheets used during this CRP are available to all Medical Physicists, please request to corresponding author.

References