Scatter correction on its own increases image contrast in TI-201 myocardium perfusion scintigraphy, but does it also improve diagnostic accuracy?


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Scatter correction on its own increases image contrast in Tl-201 myocardium perfusion scintigraphy, but does it also improve diagnostic accuracy?

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INTRODUCTION

Visual reading of 201Tl myocardium perfusion images is hampered by low image contrast resulting from poor and distant dependent spatial resolution of the gamma camera, movement of the heart and, above all, inclusion of scattered photons. These scattered photons may originate from the 135 keV and 167 keV photo peaks of thallium-201, but may also derive from other radionuclides, such as 99mTc when for instance a dual isotope protocol is used for data acquisition.1–3 Low contrast will mask (small) defects with only a mildly reduced perfusion. It also hampers the outlining of the myocardium for further processing. Scatter correction methods may reduce this image distortion by removing (most of) the scattered
photons and thus will result in a higher image contrast.

Several methods for scatter correction have been proposed during the past years.\(^4\)\(^{-}\)\(^{18}\) The triple energy window scatter correction method (TEW) presented by Ogawa et al.\(^{15}\) has been shown to perform as one of the best.\(^{19,20}\) TEW estimates the contribution of scattered photons to the acquisition data by means of two additional narrow energy windows (scatter windows, SWs) placed adjoined to the main energy window (MW) (Fig. 1). The scatter contribution to the MW data is estimated through a linear interpolation and, after filtering to remove some of the noise, subtracted from it.

In this study, we assessed the single influence of TEW scatter correction on contrast and image interpretation of clinical myocardium perfusion studies. Contrast was assessed between the myocardium and the left ventricular cavity. Moreover, defect severity was assessed. As the subtraction process, which is part of TEW scatter correction, may degrade the signal-to-noise ratio (SNR), we also assessed changes in interpretation of images and the appreciation of the quality of the scintigrams by the reader.

**MATERIAL AND METHODS**

*Data acquisition*

Stress and rest myocardium perfusion studies of 30 consecutive patients were used for this investigation. All patients were admitted for the evaluation of undiagnosed chest pain. Patients underwent either ergometric stress or received pharmacological stress test using dipyridamole (Table 1). At maximum stress, 75 MBq \(^{201}\)Tl was injected followed by a re-injection of 37 MBq \(^{201}\)Tl at rest.\(^{21}\) Immediately after stress and 60 minutes after re-injection of the radiopharmaceutical SPECT projection data were acquired using a three head rotating gamma camera (GCA 9300/hg, Toshiba, Tokyo, Japan) equipped with low-energy high-resolution collimators.

A 20% main energy window was centered on the main photo peak (~70 keV) in the energy spectrum of thallium-201. Two 7% wide energy windows, used to collect information on the contribution of scattered photons to the MW data, were placed adjoined the main energy window. Projection data were acquired over 360° with the step-and-shoot method (3 × 30 projections, 4° steps) without zoom in a 128\(^3\) matrix (3.125 mm/pixel).

*Data processing*

Prior to SPECT reconstruction, the MW data and the SW data were filtered separately by an order-8 Butterworth filter with cut-off frequencies equal to 0.12 cycles per pixel and 0.07 cycles per pixel respectively. During tomographic reconstruction only a Ramp filter was applied.

After reorientation of the data 8 short axis slices of almost 1 cm thickness each were obtained with the left ventricle (LV) centered in it (Fig. 2). To facilitate evaluation of myocardial perfusion, bull’s-eye displays were also generated.

*Contrast analysis*

Contrast between the myocardium and LV cavity as well as contrast between normally and abnormally perfused myocardium was quantitatively assessed. Apical and basal regions were excluded, resulting in the analysis of only

**Table 1** Patient data

<table>
<thead>
<tr>
<th>Patients</th>
<th>30 (19 male, 11 female)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Mean age (range)</td>
<td>60 yr (39–81 yr)</td>
</tr>
<tr>
<td>Myocardial infarction</td>
<td>11 (10 male, 1 female)</td>
</tr>
<tr>
<td>Stress modality</td>
<td>ergometric test: 25</td>
</tr>
<tr>
<td></td>
<td>pharmacological test: 5</td>
</tr>
</tbody>
</table>

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**Fig. 1** Window settings in Triple Energy Window scatter correction. The main window is centered on the peak in the photon energy spectrum of the isotope; the windows used to estimate the scatter contribution are placed adjoined to the main window.

**Fig. 2** Region of interest selection. Four mid-ventricular short axis slices were selected for contrast assessment, with apical and basal slices excluded. On the bull’s-eye display a corresponding, band-shaped region was selected to determine the minimum and maximum count densities.
the four mid-ventricular slices. A comparable region on the bull’s-eye display was selected (Fig. 2). Contrast values were determined in each of the four short axis slices separately.

Contrast was defined as

\[ C = \frac{\text{max} - \text{min}}{\text{max} + \text{min}}, \]

where \( \text{max} \) is the maximum count density found in the myocardium, either in each of the short axis slices or in the bull’s-eye display, and \( \text{min} \) is the minimum count density found either in the LV cavity of the same short axis slice or in an area with reduced perfusion in the bull’s-eye display. Contrast values were computed for each of the four analyzed short axis slices separately and for the bull’s-eye display.

Defect size assessment
The size of exercise induced perfusion defects was assessed from the bull’s-eye displays. According to the model of Maddahi,\(^2\) we allocated on the bull’s-eye display non-overlapping perfusion areas to the three main coronary arteries: LAD, LCx and RCA. A significant redistribution of the radiopharmaceutical on the rest image was defined as an increase of the relative count density of at least 10% of the maximum. For each of the three areas the size and the mean change of count density were determined with and without TEW.

Image reading
All studies were analyzed by an experienced nuclear medicine physician. To blind the reader to patient information and reconstruction differences, all studies were coded and randomly presented in six daily sessions of 10 patient studies each.

Data were visually assessed from the computer screen, which was showing the short axis slices and the horizontal and vertical long axis slices, as well as the bull’s-eye displays. Stress and the re-injection studies were shown simultaneously. Myocardial perfusion was assessed in nine segments: ANT, AL, LAT, IL, INF, IS, SEP, AS, and Apex. A four-point scale was used, ranging from absolutely abnormal (0), via probably abnormal (1) and probably normal (2) to absolutely normal (3). Changes in the visual grading between stress and rest images were confirmed in the bull’s-eye displays. In addition, the quality of each rest and stress study was graded as poor, moderate, acceptable or good (0–3).

After completion of the reading sessions the results of the scatter-corrected data were compared to those obtained with non-corrected data. An additional defect was present when the segment score changed from absolutely normal without scatter correction to probably or absolutely abnormal after TEW.

Statistical analysis
To assess the significance of the change in contrast, the paired-samples two-tailed t-test was used. The measurements of the contrast between the myocardium and the LV-cavity in each of the four slices of each patient study were assumed to be independent.

The agreement between scatter corrected and non-corrected studies was assessed through Cohen’s kappa (Ck).
RESULTS

Contrast improvement

Figure 3 shows a typical example of two short axis slices of a patient, with and without scatter correction. The contrast improvement can be assessed visually.

Figures 4 and 5 show the contrast changes induced by TEW scatter correction. All the points of the graphs lie above the line of identity, which means that the contrast increased in all slices of all studies, as well as in the bull’s-eye displays. In the stress studies, the mean contrast between the LV cavity and the myocardium without scatter correction was 0.54 (range 0.20–0.96). After scatter correction this contrast increased significantly to 0.76 (0.29–1.51). In the rest studies, contrast improved from 0.52 (0.32–1.08) to 0.78 (0.34–1.75). The average increase of contrast was 43% for stress studies and 48% for rest studies.

The mean contrast between normally and abnormally perfused myocardium as assessed from the bull’s-eye display of a single study, increased from 0.33 (0.10–0.56) to 0.42 (0.17–0.72) for stress studies and from 0.31 (0.12–0.64) to 0.41 (0.16–0.82) for rest studies. The average increase of the contrast was equal to 25% and 32% for stress and rest studies, respectively. All contrast changes were significant at a level p < 0.001.

Table 2 Changes in vascular area and severity of resolving perfusion defects induced by TEW

<table>
<thead>
<tr>
<th>artery</th>
<th>area increase</th>
<th>area decrease</th>
<th>severity increase</th>
<th>severity decrease</th>
</tr>
</thead>
<tbody>
<tr>
<td>LAD</td>
<td>17</td>
<td>7</td>
<td>23</td>
<td>4</td>
</tr>
<tr>
<td>RCA</td>
<td>10</td>
<td>16</td>
<td>12</td>
<td>15</td>
</tr>
<tr>
<td>LCx</td>
<td>11</td>
<td>11</td>
<td>18</td>
<td>7</td>
</tr>
<tr>
<td>All arteries</td>
<td>20</td>
<td>8</td>
<td>20</td>
<td>8</td>
</tr>
</tbody>
</table>

Table 3 Comparison of the quality scores with (+) and without (−) TEW

<table>
<thead>
<tr>
<th>TEW−</th>
<th>0</th>
<th>1</th>
<th>2</th>
<th>3</th>
<th>Total</th>
</tr>
</thead>
<tbody>
<tr>
<td>0</td>
<td>1</td>
<td>4</td>
<td>0</td>
<td>0</td>
<td>5</td>
</tr>
<tr>
<td>1</td>
<td>0</td>
<td>1</td>
<td>3</td>
<td>4</td>
<td>8</td>
</tr>
<tr>
<td>2</td>
<td>0</td>
<td>1</td>
<td>2</td>
<td>2</td>
<td>5</td>
</tr>
<tr>
<td>3</td>
<td>0</td>
<td>0</td>
<td>0</td>
<td>11</td>
<td>11</td>
</tr>
<tr>
<td>Total</td>
<td>1</td>
<td>6</td>
<td>5</td>
<td>17</td>
<td>29</td>
</tr>
</tbody>
</table>

Table 4 Comparison of segmental scores in stress studies with (+) and without (−) TEW. The additional defects found after scatter correction are shown in italics

<table>
<thead>
<tr>
<th>TEW−</th>
<th>0</th>
<th>1</th>
<th>2</th>
<th>3</th>
<th>Total</th>
</tr>
</thead>
<tbody>
<tr>
<td>0</td>
<td>70</td>
<td>11</td>
<td>5</td>
<td>3</td>
<td>89</td>
</tr>
<tr>
<td>1</td>
<td>1</td>
<td>5</td>
<td>9</td>
<td>8</td>
<td>23</td>
</tr>
<tr>
<td>2</td>
<td>0</td>
<td>2</td>
<td>10</td>
<td>20</td>
<td>32</td>
</tr>
<tr>
<td>3</td>
<td>1</td>
<td>1</td>
<td>7</td>
<td>108</td>
<td>117</td>
</tr>
<tr>
<td>Total</td>
<td>72</td>
<td>19</td>
<td>31</td>
<td>139</td>
<td>261</td>
</tr>
</tbody>
</table>

Table 5 Comparison of segmental scores in rest studies with and without TEW. The additional defects found after scatter correction are shown in italics

<table>
<thead>
<tr>
<th>TEW−</th>
<th>0</th>
<th>1</th>
<th>2</th>
<th>3</th>
<th>Total</th>
</tr>
</thead>
<tbody>
<tr>
<td>0</td>
<td>41</td>
<td>13</td>
<td>6</td>
<td>6</td>
<td>66</td>
</tr>
<tr>
<td>1</td>
<td>0</td>
<td>1</td>
<td>6</td>
<td>9</td>
<td>16</td>
</tr>
<tr>
<td>2</td>
<td>1</td>
<td>0</td>
<td>11</td>
<td>38</td>
<td>50</td>
</tr>
<tr>
<td>3</td>
<td>0</td>
<td>1</td>
<td>2</td>
<td>126</td>
<td>129</td>
</tr>
<tr>
<td>Total</td>
<td>42</td>
<td>15</td>
<td>25</td>
<td>179</td>
<td>261</td>
</tr>
</tbody>
</table>
RCA artery showed the opposite behavior: after TEW the defects were in general, smaller and less severe.

**Image reading**

In one TEW corrected patient study, visual analysis was considered impossible due to poor quality. Without correction, the quality of this study was judged as moderate. This study is not included in the following results.

Despite the increased image contrast in all studies, image quality was rated lower in 13 studies and did not change in 15. Quality was improved in only one study. Before scatter correction, the image quality of 22 of 29 studies was rated as good or acceptable (Table 3). After TEW, 15 of 22 studies were still rated as good or acceptable, and quality worsened in 9 studies, but was never downgraded to poor.

Tables 4 and 5 summarize the segmental stress and rest scores before and after TEW. The mean segmental score decreased from 1.91 to 1.68 and from 2.31 to 1.93 for exercise and rest studies respectively. The segmental perfusion score changed through TEW: for exercise studies, 56 segments received a lower score and 12 were judged to have a better perfusion. In rest studies the number of changes was even larger, 78 segments were rated lower and in only 4 segments an improved perfusion was decided to be present. The TEW procedure yielded 11 additional defects (1 × 3; 1 × 2) in 8 stress studies. In rest studies, 15 additional defects were scored in 12 studies that were judged as normal without TEW. Small, non-significant changes were observed in 20 myocardial segments of stress studies and in 38 segments of rest studies. A moderate agreement between the rating before and after TEW was found: Cohen’s kappa was equal to 0.60 and 0.48 for exercise and rest studies, respectively. The maximum relative perfusion change between stress and rest studies, as obtained from the bull’s-eye displays increased after applying TEW in 28 of 29 patient studies. In 3 patient studies this difference changed from a value below to above the threshold of 10%.

**DISCUSSION**

In this study, we have shown considerable image contrast improvements by only applying TEW in clinical 201Tl perfusion studies. In all slices of all studies, as well as in the corresponding bull’s-eye displays contrast increased. This increased contrast may facilitate the reading of the patient studies. But, at the same time, the scatter correction process, subtracting two noisy signals, will lead to a reduced signal and an increased noise. The final result will be a reduction of the SNR, with noise being the major degrading factor. The worsening of the SNR may nullify the improved readability of the images. Therefore, to keep the SNR at a reasonable level, we applied low pass filters to the MW data as well as to the estimate of the scatter contribution. In a previous phantom study, we have optimized the filter parameters to obtain maximum SNR values. We used these optimized parameter values in the present study. Nevertheless, after scatter correction, we still obtained sometimes poor images, but only in those studies that already had a (visually) low initial SNR and were qualified as poor. Only one study rendered unreadable through TEW.

A few minutes after injection, the amount of circulating 201Tl will be negligible. So, for the contrast between the myocardium and the LV cavity after scatter correction, one may expect values close to 1.00. In a few studies we have found very high contrasts (>1.00). Negative pixel values resulting from reconstruction filter over and under shoots explain these high contrasts, even before scatter correction. Negative numbers are not set to zero in our Toshiba computer system, but will be displayed as pixels with zero values. Overcompensation of the scatter may cause additional (too) high values. Contrast values much smaller than 1.00 even after TEW may be also due to partial volume effects and the motion of the ventricle during acquisition. ECG gating during acquisition was not applied in this study. Also, attenuation will affect the contrast values. This influence, however, will be the same for studies processed with and without scatter correction.

Another cause for overcompensation of the scatter component in the main energy window may be found in the window settings. In daily clinical practice, we center the 20% main energy window on the peak (~70 keV) in the photon energy spectrum. By doing so, not only some "good" 201Tl photons with energies around 80 keV are excluded, but also characteristic lead X-rays. These X-rays (in the 80–90 keV range) are caused by interactions of the high energy photons (135 keV and 167 keV) of 201Tl with the collimator. These X-rays do not contain the direction information of the interacting photons and should not be included in the image. To be able to compare scatter corrected with non-corrected data using a single acquisition, the main window setting was not changed in this study. Therefore, the upper scatter window that is positioned adjoined to the main window may contain too many counts, resulting in an overestimation of the scatter contribution to the main window.

In this study, we only assessed the contrast defined as the maximum differences in count densities. This type of measurement may be noise sensitive. However, the filtering of the data removes much of the noise. In fact, the data are not obtained from a single pixel, as it combines information from many neighboring pixels. Due to the noise, the increase of contrast between myocardium and LV cavity, ranging from 43% to 48%, may be a bit overestimated. When creating the bull’s-eye displays an additional smoothing is involved, resulting in a lower increase of contrast between defects and normally perfused myocardium.

Because the scatter fraction is spatially variant, the correction will be different from point to point. So, the
maximum contrast may be found at different locations before and after scatter correction. We did not assess the location of the minimum and maximum values. The change in maximum contrast does not necessarily reflect the local changes, which may be different.

We measured the image contrast in two ways, between the LV cavity and myocardium as well as between normally and abnormally perfused myocardium. Optimal contrast between the myocardium and its cavity will facilitate the delineation of the myocardium. Increased contrast within the myocardium will make perfusion defects easier to assess. One can also use dedicated color scales aimed to visualize small differences in count densities. This method, however, does not correct for the errors (inclusion of the scattered photons), but only makes small differences easier to assess.

Another way to increase visual image contrast is the subjective change of the brightness and contrast settings either of the monitor used to display the image, or by software. Such a contrast adaptation, however, will introduce a global change of the image and is difficult to reproduce. TEW will change the contrast locally by correcting point by point for the estimated local scatter contribution.

To our knowledge, no other study focussing solely on contrast changes in clinical myocardium perfusion studies has been published up to now. Yang et al. studied count ratios in dual isotope myocardial studies using $^{123}$I-labeled fatty acid (BMIPP) and $^{201}$Tl.24 To measure the count densities they used profiles through a perfusion defect and the center of the LV-cavity. Thus, they compared defect counts with the count density in the opposite myocardium, which was assumed to be normally perfused. In a Monte Carlo and phantom study, El Fakhri et al. analyzed methods to correct for several image degrading factors including scatter.25 They also found that scatter correction significantly improved contrast, but reduced SNR. In two other studies, contrast and signal-to-noise ratios were analyzed for brain and liver images respectively. Gustafsson et al. found comparable results in a Monte Carlo brain study: “...contrast was improved, but the contrast(signal)-to-noise ratio was made worse.”26

In a phantom study, Perisinakis et al. found a mean contrast increase of 1.7 but at the cost of a degradation of the SNR.27 This increase of contrast is comparable with the increase noted in our study.

In the present study, we measured contrast changes induced by TEW. Although contrast values significantly improved in all studies we could not show a significantly different clinical outcome. Cohen’s kappa, reflecting the similarity of the results before and after TEW, was around 0.5 for rest as well as for stress. Thus, there was a moderate agreement between scatter corrected and non-corrected studies.

Image reading was performed in a similar way for scatter corrected and non-corrected images, the readers were not trained to judge the scatter corrected studies. To discriminate between real and artificial defects, the same mental thresholds were applied to scatter corrected and non-corrected data. Afterwards reading results were compared to the available clinical data of the patients. The additional defects found after scatter correction were significant in size and severity and were not in conflict with the clinical status of the patients. Obviously, no golden standard was available for this clinical study. Different (visual) thresholds to distinguish perfusion defects from normally perfused myocardium may have to be applied for corrected and non-corrected data. Training of the operators, however, would have required a considerable number of studies with well-defined abnormalities. Such a study was not considered relevant within the scope of our investigations. However, it will be required when changing any imaging protocol in clinical use.

Overall we have found more and more severe defects after scatter correction. The perfusion territories of the RCA (inferior), however, showed the opposite behavior compared to the myocardial segments perfused by the LCx and LAD arteries (anterior and antero-septal). The differences in depth of these myocardial segments in the patient body will result in different attenuation of particularly the scattered lower energetic photons and so in the relative contribution of scattered photons.

In this single isotope study the scattered photons originate mainly from the higher energy photons emitted by $^{201}$Tl. In dual-isotope studies, to image stress and rest perfusion in one acquisition, the amount of scatter may be more prominent. Retraining of the operator to deal with these images, whether they are scatter corrected or not, will be required.

**CONCLUSIONS**

Triple Energy Window scatter correction significantly improves the contrast between the LV myocardium and the LV cavity, as well as between normally perfused myocardium and regions with reduced perfusion. This improved contrast makes the images easier to read and should result in an improved clinical outcome of an individual patient study. However, improved accuracy could not be observed in the present study with a limited number of patients, which may be due to a decreased signal-to-noise ratio.

**REFERENCES**

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