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The influence of electron multiplication and internal X-ray fluorescence on the performance of a scintillator-based gamma camera

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Abstract

When considering the 'standard' gamma-camera, one might picture an array of photo-multiplier tubes or a similar array of small-area detectors. This array of imaging detectors would be attached to a corresponding array of scintillator modules (or a solid layer of scintillator) in order to give a high detection efficiency in the energy region of interest, usually 8-140 keV. Over recent years, developments of gamma-cameras capable of achieving much higher spatial resolutions have led to a new range of systems based on Charge-Coupled Devices with some form of signal multiplication between the scintillator and the CCD in order for one to distinguish the light output from the scintillator above the CCD noise. The use of an Electron-Multiplying Charge-Coupled Device (EM-CCD) incorporates the gain process within the CCD through a form of 'impact ionisation', however, the gain process introduces an 'excess noise factor' due to the probabilistic nature of impact ionisation and this additional noise consequently has an impact on the spatial and spectral resolution of the detector. Internal fluorescence in the scintillator, producing K-shell X-ray fluorescence photons that can be detected alongside the incident gamma-rays, also has a major impact on the imaging capabilities of gamma-cameras. This impact varies dramatically from the low spatial resolution to high spatial resolution camera system. Through a process of simulation and experimental testing focussed on the high spatial resolution (EM-CCD based) variant, the factors affecting the performance of gamma-camera systems are discussed and the results lead to important conclusions to be considered for the development of future systems. This paper presents a study into the influence of the EM-CCD gain process and the internal X-ray fluorescence in the scintillator on the performance of scintillator-based gamma cameras (CCD-based or otherwise), making use of Monte Carlo simulations to demonstrate the aspects involved, their influence on the imaging system and the hypotheses previously discussed in experimental studies.

Key words: Gamma camera, CCD, EM-CCD, scintillator, resolution, fluorescence

1 1 Introduction

There are many applications for gamma-cameras in the energy regime from 2 8-140 keV, from medical imaging to synchrotron-based research. It is gener-3 ally possible to split current gamma-camera technology into two groups: low 4 spatial resolution and high spatial resolution systems. Here, low-resolution gamma-cameras are defined as those with a resolution of a few hundred micrometers or greater. High-resolution gamma-cameras are defined here as those with a resolution of better than 100 µm. The grouping occurs in this way due 8 to the technologies behind the detectors available for such camera systems. 9 Gamma-cameras can be made from arrays of imaging detectors, each detector 10 creating a 'single pixel' and generally measuring a few hundred micrometers 11 across. Such camera systems generally have lower resolutions in the spatial 12 regime but comparatively better spectral resolutions [1]. Alternatively, one 13 can manufacture a high-resolution gamma-camera from a single imaging de-14 vice for which each pixel is a few tens of micrometers in size [2]. Although 15 such detector systems can have much higher spatial resolutions, the spectral 16 resolution generally suffers (Sections 3 and 4). 17

In order to create the highest-resolution gamma-cameras, sub-pixel imaging 18 is required and can be achieved through photon-counting imaging techniques 19 and centroiding. The low numbers of photons recorded per event when using 20 photon-counting techniques can be lost beneath the readout noise floor of a 21 standard Charge-Coupled Device (CCD). If one uses an Electron-Multiplying 22 Charge-Coupled Device (EM-CCD), then the effective readout noise can be 23 reduced to the sub-electron level, dramatically increasing the effective signal-24 to-noise level. However, the gain process ('impact ionisation') required to in-25 crease the effective signal-to-noise ratio introduces an additional noise factor, 26 the so called 'gain noise'. This additional noise factor acts to reduce the spec-27 tral resolution and can be studied analytically, but the effect of the additional 28 noise on the centroiding accuracy is more complex and hence a simulation 29 has been produced to ascertain the level of impact of the gain process on the 30 ability to achieve sub-pixel imaging in comparison to a similar CCD system 31 to allow a spatial imaging performance comparison to be made. 32

When using a silicon-based detector for gamma-ray imaging, it is generally preferential to increase the detection efficiency through the use of a scintillator, either directly coupled to the detector or through a fibre-optic system. With a scintillator based detection system, experimental results suggest the K-shell fluorescence X-rays that can be generated in the scintillator from the incident gamma-rays (provided they are of energy greater than the K-shell binding

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energy) can be reabsorbed at another location in the scintillator, acting to 39 decrease the spatial resolution of the system. Although thresholding can be 40 used to a certain extent, this brings a dramatic reduction in the detection 41 efficiency as, for the example of the scintillator CsI(Tl), approximately 90% 42 of 'true' events can be rejected. Using a series of new simulations, validated 43 against previously reported experimental results [2-5], to look at both the 44 spatial and spectral capabilities of the detector systems, the impact on the 45 resolutions of the internal fluorescence is explored for both 'high' and 'low' 46 resolution gamma cameras. 47

This work develops on theory and experimental results taken from [2-5] and 48 the PhD thesis by the author [6], with the theory in Sections 2 and 5 taken from 49 this work. Through new additional simulations and analysis this study aims 50 to confirm the hypotheses presented in the previous experimental work and to 51 place the results in a wider context, developing the scope to include limitations 52 on the camera system. For every application of the gamma-camera the desired 53 specifications may change. Through consideration of the systems as a whole, 54 taking into account, for example, the limitations on the spatial resolution due 55 to the use of a collimator, the choice of system can be considered. The first 56 choice to be made is between that of a spatially or spectrally preferential 57 system. The choice of one detector over another is discussed and possible 58 improvements to the detector systems inferred. 59

60 2 The scintillation process

Scintillators have been dominant in the field of ionising radiation detection 61 for over one hundred years. Solid scintillation was first observed by Elster and 62 Geitel in 1903, where the presence of an alpha-emitting source led to individ-63 ual light flashes in a ZnS screen [7]. Over the last century, developments in 64 the understanding of the scintillation process and the discovery of new scin-65 tillating materials has led to many new uses throughout high-energy physics 66 and astrophysics, along with the continual development for medical imaging 67 applications from the first X-ray film through to modern dental CCD imagers. 68

A scintillator converts the energy from the absorption of ionising radiation into 69 a flash of photons of a much longer wavelength, usually in the visible region 70 of the electromagnetic spectrum. In the case of the gamma-ray detection, the 71 combination of the larger number of output photons compared to the incident 72 flux and the lower energy of the photons produced means that scintillators 73 are a near ideal choice for coupling to a silicon based imager (such as a CCD 74 or CMOS device). As most higher energy X-rays will pass straight through 75 the silicon of the device (with no scintillator present) the detection efficiency 76 for high energy photons is much reduced. Through the inclusion of a thicker 77

⁷⁸ scintillating layer, the detection efficiency can be greatly increased.

79 2.1 Inorganic scintillators

The scintillator acts to convert a single high-energy quantum into many lower energy quanta. The reduction in energy of the quanta to be detected leads to a much higher efficiency of detection than would be possible with the higher energy quantum.

- ⁸⁴ The scintillation process can be described in five stages as detailed in [8]:
- (1) Creation of electron-hole pairs through the absorption of ionising radia tion.
- Relaxation of primary e-h pairs, producing multiple secondary electrons,
 holes, photons, phonons and other electronic excitations.
- (3) Thermalisation of secondary e-h pairs through interactions with the vi brations of the environment.
- (4) Energy transfer to the luminescence centres.
- (5) Emission of energy from luminescence centres in the form of lower energy photons.
- At the energies considered in this study, the photo-electric effect dominates due 94 to a larger interaction cross-section than the Compton interactions, Figure 1. 95 Through the photoelectric effect, a hole is created in the inner electron shell of 96 the atom (K-shell). This leaves an ionised atom and a free electron with energy 97 equal to $h\nu$ minus the binding energy of the electron. The ionised atom in the 98 lattice may relax through the emission of a photon, as another electron drops to 99 fill the hole, or through the Auger effect, where further electrons are released. 100 The electrons then lose energy through further scattering or the emission of 101 photons. This process continues until ionisation is no longer possible. Electrons 102 lose excess energy through inelastic scattering until only low energy excitations 103 in the lattice remain. 104

When the energy of the excited electrons is below the ionisation threshold, 105 the electrons begin to interact with vibrations in the environment: the process 106 of thermalisation. The holes move to the top of the valence band, whilst the 107 electrons move to the bottom of the conduction band, leaving many electron-108 hole pairs each separated by the band gap energy E_q . This stage leaves N_{eh} 109 electron-hole pairs, Equation 1, where ε is the average energy required to create 110 a single electron-hole pair and E_{γ} is the energy of the incident absorbed photon 111 [8]. 112

113
$$N_{eh} = \frac{E_{\gamma}}{\varepsilon}$$
(1)



Fig. 1. The linear absorption coefficient for CsI(Tl) against incident gamma-ray energy. The photo-electric effect dominates in the region of interest in this study (below 200 keV).

The electronic excitations then transfer their energy to the luminescence centres in the scintillator (such as the Tl⁺ ions in CsI(Tl) that are involved in the electronic recombination). The luminescence centres can be excited through either the consecutive capture of an electron then a hole (hole recombination luminescence) or a hole then an electron (electron recombination luminescence) [8]. The emission of photons follows the relaxation of the excited luminescence centres to their ground state.

121 2.2 Thallium-doped caesium iodide

Thallium doped Caesium Iodide is a popular scintillator choice for current gamma-cameras due to the high emission yield (54 photons per keV gamma), the similar refractive index to that of glass in a fibre-optic plate (1.79 at emission maximum) and the low self-absorbance (re-absorbance of emitted photons) for the slow component $(3.4\pm0.5 \ \mu s)$, despite the higher self-absorbance of the fast component ($600\pm50 \ ns$), although this will depend on the crystal



Fig. 2. An SEM of the scintillator CsI(Tl). The scintillator can be seen to be formed of columns of diameter 5-6 μ m. The sample used in this study was produced by e2v technologies with a thickness of 70 μ m [11].

dimensions [9,10]. The scintillator can also be grown in a columnar form that, although not perfect, reduces the spread in the light emitted in the scintillator, Figure 2. In a similar way to a fibre-optic plate, the scintillator acts to channel the light to the CCD along the columnar structures, acting to reduce the light spread and increase the peak in intensity of the Gaussian-like profile observed at the CCD surface.

The scintillator produces approximately 60 photons per keV of incident energy, forming an approximately linear dependence on energy over the range of interest. Through the analysis of the number of photons emitted, it is in theory possible to relate this figure to the energy of the incident photons [4,8].

138 2.3 Internal X-ray fluorescence

Interactions occurring in the scintillator due to gamma-ray irradiation must be studied in greater detail in order to ascertain the imaging capabilities of a scintillator-based camera system. The interaction process inside the scintillator does not simply supply a series of identical scintillation flashes all occurring at the incident gamma-ray energy. At low energies (below the binding energies of elements in the scintillator) the interaction process is simplified and the number of lower energy photons produced is proportional to the incidentphoton energy, with one reaction point for each incident quantum.

For caesium iodide, the scintillator used as a demonstrator in this study, the binding energies of importance are those for caesium at 35.99 keV and iodine at 33.17 keV. Incident photons of energies below approximately 30-35 keV are not of interest here as these will provide a single spectral peak. For the imaging of harder X-rays, those above approximately 30-35 keV, one has to consider the impact of K-shell fluorescence on the imaging capabilities of the system.

Considering the case of a ²⁴¹Am source providing incident gamma-rays at 154 59.5 keV, one can discuss the internal X-ray fluorescence further. Approxi-155 mately 90% of interactions in the caesium iodide at 59.5 keV will be due to 156 the photo-electric effect. Of these primary interactions, the fluorescence yield 157 is approximately 88-90%, resulting from electrons falling from higher energy 158 levels to fill the hole left by the ejected electron [12]. In this case, the differ-159 ence between the interaction probability with the caesium and iodine atoms 160 is negligible. 161

At 59.5 keV, the incident photons have sufficient energy to knock an electron 162 from the inner shell of the caesium and iodine atoms in the scintillator, releas-163 ing an electron of energy equal to the incident photon energy (59.5 keV in this 164 case) minus the binding energy of the atom (either caesium or iodine in this 165 case). The ejected electron has insufficient energy to travel far from the initial 166 interaction point, but may traverse several microns. Remaining excitations in 167 the atom, along with the scintillation caused by the ejected electron, provides 168 the flash of photons at (or very close to) the initial interaction position. 169

Following this process, approximately 10% of the relaxations of the atoms follow the Auger process. The ejected electrons will cause scintillation surrounding the initial interaction position and the total sum of the number of lower energy photons produced at this point will be proportional to the incident photon energy.

The remaining 90% of relaxations result in K-shell X-ray fluorescence, where 175 an outer shell electron falls to fill the hole left by the ejected electron, releas-176 ing the difference in energy in the form of a characteristic X-ray. $K\alpha$ emission 177 results from approximately 90% of these interactions, with the remaining 10%178 resulting from $K\beta$ emission. The emitted characteristic fluorescence X-rays 179 may travel through the scintillator and leave the material undetected. In this 180 case, the number of lower energy photons produced relates to the energy of the 181 emitted electron. If, however, the fluorescence X-ray interacts with the scintil-182 lator, a secondary interaction site will occur away from the initial interaction 183 point, with the distance between the two interaction sites determined by the 184

probability of interaction at the energy of the fluorescence X-ray. It is this 185 distance between the initial interaction and the secondary event that affects 186 the imaging capabilities of the device. This process was observed in [13,14] as 187 a "separately resolved primary interaction and the secondary K X-ray interac-188 tion", giving an 'extra' event outside the line of a slit placed over the camera. 189 Here we aim to determine the impact of the detection of these 'extra' events 190 on the spatial and spectral resolution using simulations designed to confirm 191 and explore the results detailed in the experimental studies and the affect the 192 re-absorbed fluorescence on 'high' and 'low' spatial resolution gamma cameras 193 [2,13,14].194

¹⁹⁵ 3 Gamma cameras based on photo-multiplier tubes

Until recently, a standard gamma-camera consisted of an array of Photo-196 Multiplier Tubes (PMTs) coupled to a thick scintillating layer [1]. When an 197 X-ray photon interacts in the scintillator a flash of photons is produced that 198 can be detected as a series of voltages from the PMT array. The coincidence 199 of signals from a sub-array of PMTs close to the interaction location allow 200 a weighted mean position to be calculated from the relative signals between 201 the PMTs in the sub-array. The total sum of the signal from all detectors in 202 the PMT array is proportional to the energy deposited in the interaction of 203 the incident photon. In the past, PMT arrays have been limited to providing 204 intrinsic detector spatial resolutions of the order of 3-5 mm depending on the 205 crystal type and thickness, PMT dimensions and algorithms used [1]. However, 206 current systems are capable of sub-millimetre precision down to a suggested 207 limit of 0.5 mm [15], although it is noted that the spatial and energy resolution 208 depend strongly on the scintillator used and the corresponding light output, 209 limiting the FWHM energy resolution to approximately 9-11% for NaI(Tl) 210 [1,15].211

Performance in such detectors varies, dependent on the scintillator used and 212 detector arrangements. Energy resolutions of 22-24% have been reported for 213 CsI(Tl), in comparison to an improved performance of 14% for NaI(Tl) for the 214 same detector formulation [16], with a best case energy resolution of 10.8%215 FWHM achieved with NaI(Tl) [17]. Spatial resolutions were also recorded at 216 0.6-0.9 mm and 1.16 mm respectively, reaching the sub-millimetre level, but 217 still a long way from the sub-100 µm levels achievable with high resolution 218 camera systems (Section 4). Another example is that described in [18], in 219 which a 3 mm NaI(Tl) crystal was coupled to a PMT array ("Hamamatsu 220 H5900-00-C12, 1-inch square cross-wire readout type") to create a small FOV 221 gamma-camera. The spatial resolution in this case was limited to 1.5 mm, 222 although more recent developments of this system have led to an improvement 223 in spatial resolution to 0.75 mm at 122 keV using a 2 mm thick $LaBr_3(Ce)$ 224

scintillator [19]. The use of a scintillator with improved light output has in this case also brought improvements in the energy resolution to 8.9% and 13.4% at 122 keV and 60 keV respectively. Similar results of 0.9 mm FWHM spatial resolution have been reported in [20] using a LaBr₃(Ce) scintillator and position sensitive PMT.

230 4 CCD-based gamma cameras

Following from the description above of a 'low spatial resolution' gamma-231 camera, one can consider what is required to image at a much higher spatial 232 resolution. One of the main limiting factors of the aforementioned systems is 233 the use of an array of PMTs. The limit on the physical size of each 'pixel' in 234 the PMT array provides an optimal resolution limit above which such a system 235 could not deliver. One way of improving the spatial resolution of a 'standard' 236 gamma-camera is to replace the PMT array with an 'array of detectors' for 237 which the individual element size is much reduced - a CCD or CMOS/hybrid 238 based system. 239

In order to cope with applications in which a high-flux rate is required, the 240 readout speed of the detector must be increased to enable the use of 'photon-241 counting' techniques (Section 8.1). Unfortunately, an increase in the readout 242 speed of a CCD leads to an increase in the readout noise. This increase in 243 readout noise would not allow the detector to distinguish low signal levels 244 above the random fluctuations in the 'zero-level' of the image. Without any 245 readout noise and no signal present (of any type) then one would expect a 246 flat field image of zeros (a true 'zero-level'). However, with read noise, this 247 flat field of zeros is altered such that the 'zero-level' fluctuates due to the 248 random noise measured at the output of the camera system [21]. The increase 249 in readout noise can, however, be counteracted through the use of off-chip 250 or on-chip amplification of the signal - increasing the signal level before the 251 addition of the readout noise component. 252

The early development of the 'BazookaSPECT' system demonstrates the use 253 of off-chip signal amplification through the use of a second generation imaging 254 intensifier and lens system to amplify the signal from the scintillator before 255 reaching the CCD [22,23]. In comparison with the systems described previ-256 ously, very high spatial resolutions can be achieved down to 50 µm. As the 257 name of the system suggests, however, there are physical detector system di-258 mension limitations that may only be overcome through the use of on-chip 259 signal amplification. 260

²⁶¹ 5 On-chip signal amplification

The number of signal electrons in a charge packet in a CCD can be increased through the process of 'impact ionisation' [21]. When a controllable high voltage is placed over a CCD electrode, creating a high electric field between this and neighbouring electrodes, the 'impact ionisation' process can be controlled. With an increase in the signal charge and no subsequent increase in readout noise, the 'effective readout noise' (compared to original signal level) can be reduced to the sub-electron level.

269 5.1 The Electron-Multiplying CCD

The Low-Light-Level camera uses an Electron-Multiplying CCD (EM-CCD) 270 [24] to provide all light level imaging from bright sunlight down to shadowed 271 overcast starlight. The variable gain allows the device to be run as a standard 272 CCD in sunlight with unity gain and also down to very low light levels by 273 increasing the gain level. The multiplication of the signal before readout ef-274 fectively reduces the readout noise which can be reduced to the sub-electron 275 level [25]. The very low readout noise allows the detection of signals of only a 276 few photons, signal levels which would otherwise be lost beneath the readout 277 noise of a standard CCD. 278

EM-CCDs have many uses, from 24-hour surveillance to military imaging at night. The flexibility of the camera has led to the technology being exploited in night-time surveillance [26], scientific imaging [27] and medical imaging [28]. The EM-CCD shares the same basic structure as a standard frametransfer CCD. The main difference between the EM-CCD and a standard frame-transfer CCD is the addition of a gain register following on from the standard readout register.

286 5.2 The gain process

When an electron passes through a region of high-electric field it can be accelerated. If an electron passes through a region of high electric-field in silicon then the electron can gain sufficient kinetic energy to effectively slam into the lattice, breaking the silicon-silicon covalent bonds, the process of 'impact ionisation' [21]. The generated electrons can in turn gain enough kinetic energy to break further bonds, creating extra electron-hole pairs and causing further impact ionisation.

²⁹⁴ Although this process can generate spurious signal through the formation of

electron-hole pairs from non-signal electrons, the process can be controlled through the application of specific voltages to a specially designed gain structure in the CCD where the probability of impact ionisation increases as the electric-field increases in magnitude. Varying the voltage applied to the multiplication electrode structure alters the electric-field and allows the gain process to be controlled.

³⁰¹ Due to the stochastic nature of the multiplication gain, during the gain process ³⁰² each signal electron can be assumed to behave independently and may generate ³⁰³ a different number of avalanche electrons. The excess noise factor, F, a measure ³⁰⁴ of the ratio of the noise on the signal at the input to the gain register compared ³⁰⁵ to that at the output (for optical photons) where G is the total multiplication ³⁰⁶ gain, $\sigma_{n_{in}}^2$ is the variance on the signal before the gain process and $\sigma_{n_{out}}^2$ is the ³⁰⁷ variance of the output signal, is defined as [29]:

308
$$F^2 = \frac{\sigma_{n_{out}}^2}{G^2 \sigma_{n_{in}}^2}$$
 (2)

For a large number of transfers across the gain register (as found in the EM-CCDs produced by e2v technologies), this formula can be solved [6] to produce:

311
$$F^2 = \frac{2G-1}{G}$$
 (3)

To a first approximation, F tends to $\sqrt{2}$ for high gain factors (tens to hun-312 dreds) [29]. For optical photons the variance on the signal levels measured, 313 assuming Poisson statistics are valid, is equal to the mean signal level, there-314 fore although gain process allows low-signal levels to be increased above the 315 readout noise, the noise on the signal level itself increases [29]. For very low 316 signal levels where high levels of gain are required, the gain process effectively 317 doubles the variance on the signal level. The impact of this increase in noise 318 will be considered further in Section 8.2. It is worth noting that this is only the 319 case for optical photons. If direct detection of X-rays was to be used, where 320 the Fano factor must come into consideration, the noise on the gain process 321 becomes more complicated [30]. 322

323 6 EM-CCD based gamma-cameras

The use of off-chip signal amplification can be replaced by the use of an Electron-Multiplying CCD (Section 5.1), such as in the early development of the Ultra Gamma Camera (UGC) [31]. In this case, a 1 mm thick CsI(Tl) layer was coupled to an EM-CCD through the use of a fibre-optic taper. The EM-CCD used contained rectangular pixels $(20 \times 30 \ \mu\text{m})$, resulting in different spatial resolutions in the two dimensions of 60 μm and 100 μm at 122 keV. Using the peak signal in each event detected, 'energy peaks' can be observed at 122 keV and 28 keV using two different sources, although the broadening of the spectrum to lower energies causes an overlap with the 28 keV peak such that one cannot determine the origin of photons measured at lower energies (if both sources were present).

Each X-ray interacting in the scintillator will do so at a different depth and will therefore generate a signal at the EM-CCD with a differing 'spread': the so called 'depth of interaction' (DOI) effect. Early attempts at producing an energy spectrum from interacting events detailed in [31] use the peak signal in an event as a measure of the energy of the X-ray interacting. Whilst this gives 'energy peaks', the DOI effects broaden the spectrum such that the broad peaks overlap on a large scale.

Developments in energy detection techniques in an attempt to remove scat-342 tered events produce improvements in the spectrum, but energy resolutions 343 of 40 keV and 42 keV at 122 keV and 28 keV respectively [32] show that the 344 technique is limited in comparison to the previously discussed detectors. A 345 higher detector resolution combined with energy discrimination capabilities 346 has been said to be essential for future Single Photon Emission Computed 347 Tomography (SPECT) systems [33], but measurements of the energy resolu-348 tion of 33 keV at 140 keV left a requirement for further study, despite the 349 improvements in the spatial resolution. More recently, a spatial resolution of 350 59.4 µm was reported using a CCD97 [34]. 351

The method used to extract the energy spectrum from the raw X-ray inter-352 action events in the scintillator recorded in the EM-CCD has a major im-353 pact on the energy resolution achieved. Developments to incorporate binning 354 of the signal in each event begin to take into account the DOI effects that 355 are ignored in simple 'peak signal' techniques. However, DOI effects can be 356 approached from two further directions. A Maximum Likelihood Estimation 357 (MLE) technique was used in [14], incorporating a calibration stage in the de-358 vice development, aiming to use recorded events of known DOI to determine 359 the DOI of events recorded when imaging. The second approach makes use of 360 'scale-space', a form of Wavelet transform, developed from algorithms used in 361 computer vision [35], adapted to improve the resolution of gamma-cameras [3]. 362 Applications of these techniques to improve the resolution of cameras showed 363 great promise when applied to simulated data [3] and have been further devel-364 oped with respect to experimental data [4] using the camera described below. 365 The same technique has also since been detailed in [36] in which it was found 366 to offer significant improvements over the techniques described above (as in 367 [32], although still limiting the energy resolution a best-case of over 40% 368 energy resolution at 140 keV and a spatial resolution of 59 µm. 369

370 7 Event separation

The results shown in Figure 3 are taken from a Monte Carlo simulation de-371 signed to investigate the influence of the re-absorbed fluorescence on the spa-372 tial resolution of the detector. A 'block' of scintillator, CsI(Tl) has been sim-373 ulated with dimensions of 1 cm \times 1 cm \times t μ m, where t is the scintillator 374 thickness (here shown at 70 μ m, 350 μ m and 700 μ m). A set flux of incident 375 photons is entered into the system along the normal to the imaging plane 376 using a random distribution in the xy plane. Each incident photon to the sim-377 ulation is generated with a randomly generated interaction distance (based 378 on the interaction length) that will either fall within or outside of the block 379 of scintillator. Photons that do not interact within the scintillator are lost 380 from the system. Photons that interact within the scintillator, using standard 381 Monte Carlo techniques, will generate a fluorescence X-ray as appropriate 382 based on the probabilities discussed in Section 2. If a fluorescence X-ray is 383 emitted, then this photon will travel in a randomly generated direction away 384 from the production position and this is then tracked until interaction or exit 385 from the scintillator structure (as with the primary photons). All locations of 386 interactions are recorded and the distance travelled in the xy-plane (the plane 387 of the detector pixels) is then calculated and this is shown in the distribu-388 tions in Figure 3. The incident photons are able to interact at any position 380 in the scintillator as determined by probability of interaction defined by the 390 interaction length and the scintillator dimensions. This 'realistic' positioning 391 of the initial interaction positions ensures the emitted fluorescence photons 392 will pass through the scintillator having been generated across the scintillator 393 with no bias. If one considers the distance between the primary and secondary 394 interaction positions from the simulated results shown in Figure 3, it is clear 395 that negligible signal will be detected outside a millimetre in radius from the 396 primary interaction location. 397

With a spatial resolution of the order of 0.5 millimetres and above, one is forced to consider the majority of primary and secondary events as 'single' events. In this way, the sum of the energy deposited in the primary and secondary interactions is recorded for this 'single' event, albeit with a bias on the positioning of the interaction position from this combination of the original interaction and the re-absorbed fluorescence X-ray, further degrading the spatial resolution.

⁴⁰⁵ Despite the apparent limits on the spatial resolution, the grouping of the ⁴⁰⁶ energy deposited from primary and secondary interactions leads to a high ⁴⁰⁷ spectral resolution (as high a resolution as 6% FWHM at 140 keV and 3% ⁴⁰⁸ FWHM at 662 keV [37]). The presence of the secondary interactions has a ⁴⁰⁹ major impact on not only the ability to achieve a higher spatial resolution, ⁴¹⁰ but also a major impact on the spectral resolution of the detector.



Fig. 3. The interaction distance for the fluorescence X-ray produced in CsI, simulated here for scintillator thicknesses of 70 μ m, 350 μ m and 700 μ m with a constant incident flux. The results are presented in the xy plane only, as this is the plane in which an image would be taken. As the distance approaches 1 mm, it is seen that the number of counts has rapidly decreased.

411 8 Scintillator-coupled EM-CCD

In order to demonstrate the spatial and spectral capabilities of a gamma cam-412 era designed for high spatial resolution applications, a 70 µm columnar CsI(Tl) 413 scintillator layer has been coupled to an e2v CCD97 [38] through a fibre-optic 414 plate (approximately 3 mm thick). The CCD97 has been operated here with a 415 pixel readout rate of 1 MHz for demonstration purposes but can be operated 416 at up to 15 MHz with appropriate drive electronics. The increased amplifier 417 noise introduced through increasing the readout rate has a negligible impact 418 on the image output from an EM-CCD detector as the multiplication gain can 419 simply be increased to reduce the effective readout noise to the sub-electron 420 level [25]. 421

422 8.1 Photon-counting

The standard operation of a CCD-scintillator imaging system, such as those used in dental imaging, involves the integration of signal over a pre-defined period of time. During this time period, any light generated in the scintillator that is collected by the CCD will be summed to create a single contrast image.
Any spread of light in the scintillator away from the initial interaction location
will be summed in the image, degrading the spatial resolution by adding a
secondary (wider) Gaussian profile in the Edge Spread Function (ESF), thus
causing the rapid drop in MTF at low spatial frequencies, Figure 4(a).

An alternative method of operation involves the capture of many images over 431 much shorter integration times such that each image contains a number of indi-432 vidual interactions, with each interaction separated by several pixels from the 433 next. Through the analysis of a batch of images of this type, one can improve 434 the imaging capabilities of the camera system dramatically. This 'photon-435 counting' technique is not straightforward and there are many factors that 436 must be overcome, such as the impact of the depth of the primary interaction 437 in the scintillator on the spread of the event observed by the detector [3]. Once 438 these challenges have been considered, the batch of images can be analysed 439 using a variety of methods on an event by event basis (Section 6). 440

A simple tungsten edge was placed against the scintillating layer to act as a 441 mask to the incident X-ray photons, in this case emitted from a solid ²⁴¹Am 442 source (59.5 keV) with an activity of 18.4 kBq at the time of the experiment. 443 The 0.5 mm thick tungsten edge prevents approximately 95% [39] of the inci-444 dent 59.5 keV photons from reaching the scintillator. The transmitted photons 445 produce a small reduction in contrast between the covered and uncovered ar-446 eas of the CCD97. The active area of the ²⁴¹Am source is in the form of a 447 disc of diameter 5 mm. A thin layer of silver foil is present in the source 448 holder, contributing a small amount of K α X-ray fluorescence with an energy 449 of 22 keV to the lower energy peaks in the spectrum. With the source placed 450 at a distance of 4 cm from the tungsten edge, the geometry of the system 451 broadens the spatial resolution that is measured, giving a spreading of the 452 incident photons 'under the tungsten edge' across a width of approximately 453 6 ± 2 microns (depending on the exact alignment) on the back surface of the 454 scintillator due to the angular incidence of the X-rays. Further improvements 455 in the measured spatial resolution would be expected if a columnar beam of 456 incident photons was supplied to the system, but the arrangement used here 457 allows a proof of concept study to be undertaken in which the results can be 458 compared with that of the 'low resolution gamma cameras' discussed earlier. 459 K series tungsten fluorescence is of too high an energy to be generated from 460 the 59.5 keV incident photons (the K absorption edge for Tungsten has an 461 energy over 59.5 keV) and the L series fluorescence (12.6 keV and below) will 462 be removed during thresholding of the images if present. 463

Following the extraction of the individual events from the images using a technique described in [4] where it is possible to adjust the event selection criteria to fit the detector environment and application, it is possible to centroid the events such that a new 'integrated' image can be created from the individual



(b) Modulation Transfer Function (MTF).

Fig. 4. Fits taken to experimental data at 60 keV in integrating and photon-counting modes. (a) The LSF is not formed from a single Gaussian profile but is instead formed from the sum of two independently characterised profiles. (b) The resulting MTF, taken as the discrete Fourier transform of the LSF, showing the 'knee' in the curve due to the two components of the LSF. [2]

centroid locations. Through the use of this process, one can remove the effectof the spread in the light generated in the scintillator.

470 8.2 Spatial resolution

Experimental results presented in [2] detail the spatial capabilities of such a 471 detector, as summarised in Table 1 and Figure 4. It must be noted that the line 472 spread function (LSF) is not formed from the standard Gaussian-like profile. 473 The complication in the ESF and LSF profile introduced was proposed to be 474 due to the addition of a secondary Gaussian component to the LSF (such that 475 profile can be fitted with the sum of two Gaussian profiles, both centred at 476 the same position) due to re-absorption of the internal fluorescence X-rays 477 [2]. A narrow profile represents the standard detector LSF with a broader 478 component found at a lower count rate. 479

Imaging method	FWHM of LSF
Integration (standard)	80 µm
Photon-counting	$31~\mu{ m m}$

Table 1

Spatial resolution measurements for the scintillator-coupled EM-CCD in integrating and photon-counting modes [2].

480 8.3 Spectral resolution

With the approximately linear relationship between the light output of the 481 scintillator and the incident photon energy, several methods are available to 482 produce an energy spectrum of events. The use of the photon-counting mode 483 is essential for this purpose; in the integrating mode all energy information is 484 lost. The most simple method involves the use of the peak intensity of each 485 event [31]. This does not, however, make allowances for the variation in the 486 event profiles with depth of interaction in the scintillator, hence producing sub-487 standard results. This method can be improved through the summation of the 488 central pixel of the events with surrounding pixels, now including the edges of 489 the Gaussian profiles [2]. Unfortunately, for every extra pixel included in this 490 summation, additional noise is also included. Using more complex methods, 491 such as the use of Scale-Selection as discussed in [3.5,36], one is able to produce 492 more detailed information about each event, such as the inference of the depth 493 of interaction in the scintillator from the spread of the photons in the event 494 profile. 495

⁴⁹⁶ With the requirements on detectors to enable high-resolution imaging, the ⁴⁹⁷ detector noise is a limiting factor unless an EM-CCD is used to reduce the ef-

fective readout noise to the sub-electron level. In reducing the effective readout 498 noise, one must use a gain process to increase the number of signal electrons. 499 This gain process introduces an approximate increase in the noise level on the 500 signal of $\sqrt{2}$ (as discussed in Section 5.2). Taking an ²⁴¹Am event in 70 µm of 501 CsI(Tl), as implemented in [2], a peak signal (in the central pixel of an event 502 hitting the centre of a pixel) of approximately 40 photons was measured when 503 considering the 30 keV components that dominate the spectrum. If a more ef-504 ficient coupling mechanism is used between the scintillator and detector then 505 it is possible to increase the number of photons detected, whilst if a less ef-506 ficient coupling mechanism is used then fewer photons may be recorded. If a 507 fibre-optic taper is used, one must include the associated losses involved. The 508 directly neighbouring pixels were found to have 5-25 photons each (dependent 509 on photon hit position in the pixel) in the previous study [2]. Using simulated 510 Gaussian profiles one is able to determine a 'best-case' energy resolution from 511 an approximation to the light spread, multiplication noise components and 512 intrinsic scintillator resolution. The results presented in Figure 5 are based on 513 the experimental results with respect to the form of the event profiles achieved 514 with the CsI(Tl) detector system detailed in this paper and literature values 515 are used for the comparisons to NaI(Tl) and LaBr₃ and for the incorporation 516 of the fibre-optic tapers [8,40]. These predicted results compare well with the 517 experimental system detailed in Section 8, taking measurements from the ex-518 perimental spectra shown in Figure 7, and demonstrate that there are indeed 519 limitations on the energy resolution achievable with such systems formed from 520 scintillator-coupled EM-CCDs. In comparison to the specific experimental de-521 tector system detailed in this study and the spectra recorded for Am^{241} and 522 Co^{57} , it can be seen in Figure 5 that the performance is verging on the intrinsic 523 performance limits achievable with such coupling. 524

⁵²⁵ 9 Simulating a scintillator-based gamma-camera

In order to facilitate a better understanding of the camera system, several 526 new simulations have been produced, allowing the demonstration and expla-527 nation of the effects noted in previous experimental studies and investigation 528 of their impact. The energy deposition and localisation in the scintillator has 529 been simulated, making use of the interaction lengths of each photon energy 530 in the scintillator (CsI in this case). Using a three dimensional scintillation 531 body, random directional generation for each X-ray photon following interac-532 tion in the scintillator allows the position of the event profile in the plane of 533 the EM-CCD to be modelled. The interaction chain for the simulation is based 534 on Figure 6, detailing the emission and re-absorption (or loss) of fluorescence 535 photons. From the experimental results of [2] it was suggested that the 'two 536 Gaussian' profile of the LSF was caused by the re-absorbed fluorescence com-537



Fig. 5. Expected best-case energy resolution for the experimental camera system (CsI:Tl with a 1:1 fibre-optic taper) over a range of energies. Literature values for different scintillators [8] have been incorporated to predict the performance of similar systems following a change in scintillator or fibre-optic taper. The experimental results taken from Figure 7 for Am^{241} and Co^{57} are shown for comparison with the theoretical calculations and demonstrate that the detector detailed in this study is verging on the intrinsic limits for the coupling system used.

ponent, resulting in the curvature at the edges of the ESF (Figure 4(a)). These 538 simulations aim to demonstrate the effect of the imaging of the fluorescence 539 X-rays through a test-case simulation of the "perfect detector'. The "perfect 540 detector" is assumed to have an MTF of one across all spatial frequencies and 541 a scintillator in which the visible photons generated in the scintillator do not 542 spread out as they pass towards the EMCCD. Therefore any degradation in 543 resolution (or degradation of the ESF from a simple step function) is caused 544 by the detection of the X-ray fluorescence. Also of interest here is the impact 545 of the interaction chain (as detailed in Figure 6) on the spectral performance. 546

547 9.1 Spectral performance

Two spectra, measured using the experimental set-up detailed in Section 8.1 using a CCD97 and CsI(Tl) scintillator, for Am^{241} (18.4 kBq) and Co^{57} (70 kBq) are shown in Figure 7 (data points, [4]). The CCD97 was operated in inverted mode at -20±5°C (to suppress dark current) at a readout rate of 1 MHz (0.3 seconds per frame), with approximately 10⁵ frames analysed for Am^{241} and 1.7×10⁴ frames analysed for Co^{57} (a large number of frames was required due to the low activity of the source). A higher frame rate and a source with a



Fig. 6. Flowchart of the interactions incorporated into the simulation process for irradiation of CsI by 59.5 keV gamma-rays.

higher activity could be easily implemented into the system with appropriate electronics and source availability; the increased readout noise from the increased readout speed can be counteracted by an increase in the gain applied with the EM-CCD to remain at an effective readout noise of 1 electron rms. The total number of interactions recorded per frame is the only limit on the activity at any readout speed, such that events must not overlap (no pileup) for the centroiding of individual events to be possible.

The spectrum achieved using Am²⁴¹ shows how the peak at approximately 562 30 keV dominates over the full-energy peak at 59.5 keV. The peak at approx-563 imately 30 keV is formed from many components, namely the re-absorbed 564 fluorescence (K α and K β from both Cs and I), the related escape peaks and 565 fluorescence from the silver foil in the ²⁴¹Am source. With the limits on the 566 spectral resolution incurred from the intrinsic scintillator resolution and the 567 multiplication gain noise, one cannot separate out the re-absorbed fluorescence 568 events (located away from the primary photon interaction position) from the 569 escape events (at the primary photon interaction position). It is only possi-570 ble, therefore, to exclude 'all events' around 30 keV in order to remove the 571 re-absorbed fluorescence and therefore to reduce the overall number of counts 572 dramatically by removing the escape events. In this respect, it is therefore 573 essential to determine the requirements of the imaging system and consider a 574 spatial resolution versus count rate trade-off. In the case of the Am²⁴¹ spec-575 trum, there is an element of separation between the peaks at 30 keV and 576 the full-energy peak and therefore one can improve the resolution in the case 577 where this is the priority over the number of counts measured (see Section 11). 578

The simulated results shown in Figure 7 take the simulation of events dis-579 cussed in Section 9 and incorporate the energy resolutions shown in Figure 5, 580 measured from the experimental results; each event generated in the simula-581 tion has a "recorded energy" randomly generated across a Gaussian profile. 582 The simulations include full-energy events, $K\alpha$ and $K\beta$ fluorescence from the 583 caesium and iodine in the scintillator, the related escape events and $K\alpha$ fluo-584 rescence from silver (in the ²⁴¹Am case) and tungsten (in the ⁵⁷Co case), with 585 the quantity of external fluorescence (silver or tungsten as appropriate) set as 586 a free parameter to be fitted with the experimental data due to the complex 587 geometries and uncertainties involved in quantising the number of fluorescent 588 X-rays generated and subsequently detected. 589

Taking the case of the ²⁴¹Am spectrum, Figure 7(a), the appropriate structure can be clearly seen in the simulated results as normalised to the intensity of the 60 keV peak. However, the simulated peak at approximately 30 keV is seen to be higher than that in the experimental results, with a dip in the simulated results at approximately 30-50 keV. This discrepancy between the simulated and experimental results is thought to be due to the methods used to calculate the energy of each event in the experimental case. The overestimation in the



(a) Experimental spectra for Am^{241} (data points) and the simulated spectrum (solid line).



(b) Experimental spectra for Co^{57} (data points) and the simulated spectrum (solid line).

Fig. 7. (a) The spectrum achieved experimentally from the Am^{241} source (data points, [4]) shows the components at the energies specified in the flowchart from Figure 6. A cut-off at approximately 40-50 keV allows the removal of some of the reabsorbed fluorescence, but this removes the bulk of the escape events also and therefore dramatically reduces the number of counts. (b) The experimental spectrum obtained using Co^{57} (data points, [4]) can be explained following a similar analysis as shown in Figure 6 but for Co^{57} . The peak at approximately 60 keV is thought to be the detection of $K\alpha$ fluorescence from the tungsten edge.

⁵⁹⁷ peak at approximately 30 keV is approximately equal to underestimation of ⁵⁹⁸ the spectrum around 40 keV. Although the simulation accounts for directly ⁵⁹⁹ coincident events, interactions in close proximity that are not counted as single ⁶⁰⁰ events lead to an increased energy being attributed to some events, Figure 8. ⁶⁰¹ It is thought that the dip in the simulated spectrum can be accounted for in ⁶⁰² this way.

In order to demonstrate the further complications that occur when using a 603 higher energy gamma source, the 122 keV photons from a Co^{57} were imaged 604 and a similar spectrum produced, Figure 7(b). A similar analysis to that de-605 tailed in the flowchart in Figure 6 can be used to determine the multiple 606 energies that will be detected from the 122 keV source, including fluorescence 607 from the tungsten edge at approximately 60 keV and the related escape peaks 608 (noting the spectrum achieved from the incident 59.5 keV photons from the 609 previous testing with the Am²⁴¹ source). In decreasing energy, the full-energy 610 peak can be seen at 122 keV, but this is not fully resolvable from the escape 611 peak at approximately 90 keV. The fluorescence of the tungsten edge can 612 be seen at approximately 60 keV. The large peak at approximately 30 keV is 613 formed from several components, namely the reabsorbed fluorescence from the 614 122 keV incident X-rays and the reabsorbed fluorescence (and corresponding 615 escape peaks) from the tungsten fluorescence. The low energy peak at 14 keV 616 is due to emission from the Co^{57} source at this energy (although much of 617 this has been removed through thresholding). Taking these energies, it is pos-618 sible to separate the spectrum into the related components, Figure 7(b). In 619 this case, the spatial coincidence of fluorescence, escape and primary events 620 is greatly complicated by the inclusion of the fluorescence from the tungsten 621 edge and the corresponding K-shell fluorescence of the scintillator. 622

The simulation has again been run for the case of the 57 Co and the results shown in Figure 7(b). As detailed for the 241 Am case, the intensity of the lower energy peak is overestimated in the simulation, followed by a similar dip around 40 keV. This is thought to be for the same reasons as detailed in the 241 Am case, as outlined in Figure 8.

In the case of higher energy sources, therefore, the improvement of the spa-628 tial resolution through the removal of fluorescence events must be considered 629 as somewhat more complex. Through the removal of low energy events (for 630 example, below 50 keV), one can remove the re-absorbed fluorescence and 631 leave a higher number of full-energy and escape events than in the Am^{241} 632 case detailed above. However, the detection of many escape events (due to the 633 potential increased overlap between primary energy peak and escape peak) 634 can leave a minor degradation of the spatial resolution from the range of the 635 electrons emitted of a few micrometers for each interaction. 636

 $_{637}$ The energy resolution measured for the main peak at 60 keV from the Am²⁴¹



(c) Semi-coincident events.

Fig. 8. The "energy" of each event is assumed to be proportional to the summed signal over a fixed area window surrounding the peak in intensity of each event. (a) Two events have occurred across the pixilated structure of the CCD such that no signal from the second event is summed with the first. The "energy" recorded is that of the first event only. (b) Two events occur in very close proximity, such that the spread of signal from each event creates one "single" event that is recorded with the summed energy (60 keV here). (c) Two events occur a short distance apart such that a fraction of the signal from the second event is included in the "energy" recorded for the first event. The amount of "extra energy" included with the first event will vary with the distance between the events and therefore some of the low energy events will be moved to higher energies in the spectrum across a continuum from 30 keV up to 60 keV, from (a) to (b).

source in the experimental results shown is 33% (20 keV at 60 keV), converging 638 on the limits imposed by the multiplication gain process and intrinsic energy 639 resolution of the scintillator as demonstrated in Figure 5. This result, whilst 640 not reaching the energy resolutions found in the literature for PMT-based 641 gamma cameras (Section 3), the results are consistent with those reported for 642 EM-CCD based camera systems (Section 6). The energy resolution could be 643 improved through the more efficient coupling of the scintillator to the EM-644 CCD or through the use of a scintillator with a higher light output. 645

646 9.2 Positional accuracy

A second simulation has been used to look at the errors involved in the cen-647 troiding process when using a 'high spatial-resolution' gamma-camera. Cen-648 troiding must be used to provide sub-pixel locations for the event profiles. The 649 sub-pixel locations can be used to create an image that no longer includes the 650 spread of light in the scintillator (as would be found in an integrated image). 651 It is this removal of the light spread that provides the dramatic improvements 652 in spatial resolution detailed in Section 8.2. The noise on the signal does, how-653 ever, affect the centroiding process. The noise components of most importance 654 here are the noise on the signal input to the EM-CCD, the noise on the gain 655 process and the readout noise of the device. 656

The simulation produced for this study creates a Gaussian profile with param-657 eters based on the experimental data of [2]. The central point of the Gaussian 658 profile (FWHM of 38 µm and peak signal of 80 electrons at 60 keV, scaling 659 linearly with energy) is positioned in the two extremes of the pixel location: in 660 the centre and in the corner of the pixels. Edge effects are apparent in the cen-661 troiding process of this noiseless signal due to the use of the 'centre of mass' 662 approach, as one would expect, where the calculated centroid positions are 663 biased towards the centre of the pixel. These effects can be corrected for using 664 the η -algorithm [41]. One is then able to simulate the noise on the signal and 665 add readout noise (in this case 10 electrons rms) to simulate the noise sources 666 for an 'equivalent non-EM CCD' running at approximately 1-2 MHz (effec-667 tively the CCD97 with multiplication gain of one to allow direct comparison 668 with the same pixel sizes). A second option is to apply gain and subsequently 669 add the noise component from the gain process and the readout noise to sim-670 ulate the noise sources for an EM-CCD. The results from these tests with 671 varying energy are shown in Figure 9, demonstrating the 'best case' centroid-672 ing performance if no other degrading factors relating to the scintillator are 673 considered. 674

The centroid error, as shown in Figure 9, refers to the difference between the input Gaussian central location and that calculated from the centroid



(b) Corner of the pixel.

Fig. 9. A comparison of the expected optimal centroiding accuracy using a simple 3×3 "centre of mass" for a standard non-EM CCD as discussed previously (10 electrons rms readout noise) and an EM-CCD (with a gain of 10) from simulations of a representative Gaussian profile (FWHM of 38 µm and peak signal of 80 electrons at 60 keV, scaling linearly with energy) placed in the centre (a) and corner (b) of a pixel. The excess noise from the gain process, when coupled with the reduction in the effective readout noise, has minimal impact and the 'best case' resolution improves beyond that of the standard CCD. The effects are emphasised for the corner of the pixel where the signal does not 'peak' in one pixel, but instead is shared over four pixels in a 2×2 grid, with subsequent reductions in signal for the pixel.

algorithm ("centre of mass" on a 3×3 pixel area around the pixel of peak signal) following the inclusion of the specified noise sources. Centroiding errors are provided across a range of energies relevant to this study where a 60 keV event is said to peak at approximately 80 photons for events in which the signal is centred around the centre of a pixel.

It is clear from Figure 9 that the EM-CCD offers improved performance over 682 the equivalent CCD in all cases, shown here for a Gaussian with FWHM of 683 $38 \ \mu m$ on a pixel size of 16 μm for both the CCD and EM-CCD. The effects 684 are more pronounced at lower energies as the signal-to-noise ratio is reduced in 685 comparison to the higher energy cases. When the event signal is centred over 686 the corner of a pixel, the peak signal, to a first approximation, may be consid-687 ered as almost one quarter of that in the pixel-centre case, hence producing 688 a large reduction in the signal-to-noise ratio and therefore offering very poor 689 spatial centroiding performance when using a standard CCD. The EM-CCD, 690 however, performs consistently well over all energies and event locations in the 691 pixel. 692

The simulated optimal centroiding accuracy data discussed above show that 693 the noise on the gain process has little effect on the accuracy of the cen-694 troiding process, despite the added problems caused to the spectral resolution 695 discussed in Section 8.3. If operating at the high-readout speeds required for 696 many photon-counting applications, the readout noise of the standard CCD 697 could be expected to be higher than the 10 electrons rms used in the simu-698 lations here and hence the low-noise performance of the EM-CCD becomes 699 essential to maintain the centroiding accuracy. 700

⁷⁰¹ 10 The impact of internal X-ray fluorescence on spatial resolution

Previous experimental studies have detailed the observation of reabsorbed 702 fluorescence events detected outside the area of imaging area [2,4,13,14]. Our 703 simulations are consistent with hypothesis that the two Gaussians seen in the 704 LSF, as demonstrated in Figure 4(a), result from the initial interaction events 705 (sharper profile) and the re-absorbed fluorescence events (broader profile). The 706 impact of the broader profile is dramatic and can be seen in the MTF curves 707 shown in Figure 4(b) as the sharp drop in MTF at low spatial frequencies. If 708 one was to use lower energy X-rays (with energies below the ionisation energy 709 of the components of the scintillator) one would expect the MTF to be greatly 710 improved, following the form of the MTF shown here at higher frequencies, 711 but now across all frequencies. 712

713 10.1 Simulating the impact of re-absorbed fluorescence on the edge spread 714 function

In order to separate out the impact of re-absorbed fluorescence on the spa-715 tial resolution of a gamma camera, all aspects of the detector itself must be 716 removed. To this end, a simulation of a camera system with a perfect Edge 717 Spread Function (ESF), a step-function from zero to one, was produced to 718 match the detector used in the previously mentioned experimental programme 719 [2]. The detector is formed from a 1 cm square CsI scintillator of thickness 720 70 µm. For the purposes of this simulation, all 'detector' aspects of the resolu-721 tion are to be removed, hence the interactions of photons in the scintillator are 722 assumed to be point-like and depth of interaction effects and light spread are 723 not included in the resulting ESF. In this way, one can separate out the purely 724 fluorescence-induced effects. It is therefore possible to simulate the equivalent 725 set-up to that used experimentally to determine the ESF and LSF, but here 726 with the 'detector' components removed. 727

The experimental set-up has been simulated through the provision of incident 728 photons of 60 keV to the simulation across a set area only (to simulate an 729 effective tungsten edge). The interactions in the scintillator are tracked and 730 the location of photon interactions in the scintillator are recorded. The re-731 sults produced show a clear curvature of the ESF. For the 'perfect detector' 732 as simulated here, one would expect a sharp step-function, however, the ESF 733 is anything but sharply defined. The curvature is consistent with that seen 734 experimentally and shows clearly the influence that the re-absorbed internal 735 fluorescence has on the edge of an object being imaged, Figure 10. The recog-736 nisable curvature to the ESF is seen in the simulated results in which only the 737 re-absorbed fluorescence is included; no other aspects that might degrade the 738 detector resolution have been included. This curvature to the 'perfect' detector 739 response is responsible for the secondary, broader Gaussian component of the 740 LSF seen in the experimental results of Figure 4(a) and can be seen to stretch 741 across several hundred micrometers as implied in Figure 3 in which the distri-742 bution of re-absorbed fluorescence in the xy plane is displayed. This therefore 743 demonstrates that the secondary Gaussian in the LSF can be caused by the 744 re-absorption of the fluorescence X-rays. In the experimental data, the 'cur-745 vature' displayed here is convoluted with the detector response (the central, 746 more narrow, Gaussian shown in the experimental LSF, Figure 4(a)). 747

748 10.2 Re-absorption distance and the energy resolution

Referring back to Figure 3, one can see that for this example of using a 70 μ m thick CsI(Tl) layer with an incident photon energy of 60 keV, the initial event



Fig. 10. Top: A simulated image of an edge for a detector system with a perfect (step-function) ESF if internal fluorescence were to be ignored. Recorded events are not limited to the right hand half of the image upon which the photons were incident to the system, showing the influence of the re-absorbed internal X-ray fluorescence (in this case for 60 keV incident photons in a 70 μ m thick CsI:Tl layer). Bottom: Without re-absorbed fluorescence one would expect a perfect step-function, with the roll-over of the signal demonstrating the presence of signal from the re-absorbed fluorescence, as displayed in the inset ESF from experimental data.

and the re-absorbed fluorescence X-ray will be included in the same 'event' if 751 the detector spatial resolution is of the order of a few hundred micrometres 752 or more. Not only is this the case, but if a much larger pixel size is used with 753 individual scintillator modules, one can also capture additional visible photons 754 from reflections occurring at the edges of the scintillation objects and it then 755 becomes possible to achieve energy resolutions of the order of the limits of 756 the scintillation process itself. With this greatly improved energy resolution, 757 however, comes a significant reduction in the spatial capabilities of the device 758 as the effective binning of the signal increases the size of each imaging 'pixel'. 759

Fig. 11. Results from the same experimental campaign as detailed in Section 8. An energy threshold of 45 keV was used to remove many'low energy' events and hence remove a portion of the fluorescent and escape events. The improved resolution is shown by the improved MTF across all spatial frequencies, but particularly at high spatial frequencies (a factor of 2 improvement at 20 lp/mm).

⁷⁶⁰ 11 Reducing the impact of the internal X-ray fluorescence

It has been detailed in [2] that through the use of energy discrimination (using 761 the calculated energy of a profile through methods as described in Section 8.3) 762 one can remove the many of the re-absorbed fluorescence events. This process 763 does, however, come at a cost to the effective detection efficiency, as a large 764 number of primary interaction events will also be removed, leaving only ap-765 proximately 10% of events for which no X-ray fluorescence occurs when imag-766 ing at 60 keV. At 60 keV, for a ²⁴¹Am source, the spatial resolution (FWHM 767 of the LSF) can be improved from $31 \ \mu m$ to $25 \ \mu m$ [2], although this improve-768 ment in FWHM is mostly due to the removal of the escape electrons which 769 are responsible for 'events' recorded a few micrometers away from the initial 770 interaction positions. The main improvement from the energy discrimination 771 process is instead that of the reduction in the intensity of the broader Gaus-772 sian component of the LSF, that due to the fluorescence, and hence a much 773 improved MTF and greatly improved imaging performance, Figure 11. 774

Fig. 12. Simplified summary of the most appropriate gamma camera detector choice.

775 12 Related detector developments

Over recent years, the development of hybrid pixel detectors has brought the 776 possibility of high spatial resolution detectors which are capable of also pro-777 viding a high energy resolution. The bonding of CdTe and CdZnTe detectors 778 to CMOS readout chips has enabled a combined performance to be achieved, 779 albeit on the small area scale as one loses the option to apply optics to expand 780 the imaging area (such as a lens system or through fibre-optic tapering). Re-781 sults reported so far include an intrinsic spatial resolution of 75 μ m at 122 keV 782 in studies detailed in [20] using a CdTe pixel detector $(14 \times 14 \times 1 \text{ mm}^2)$ with 783 256×256 square pixels and a 55 µm pitch. This pixel detector is coupled to 784 a CMOS single photon counting integrated circuit from the Medipix2 series. 785 In similar early developments using a CdZnTe sensor, a FWHM (at room 786 temperature) at 122 keV of 2.5% has been reported [42], although no spatial 787 results are presented. 788

789 13 Conclusions

The choice of detector for X-ray imaging, as with all applications, must be carefully considered, as the experimental and simulated data presented here have shown. The most appropriate choice of detector can be briefly summarised as shown in Figure 12.

If the spatial resolution of a system is limited by an external factor such as the 794 collimators used in medical imaging, then the scintillator-coupled EM-CCD is 795 the optimal detector choice. The dramatic improvement in spatial resolution 796 over more traditional X-ray imaging systems has negligible impact on the 797 images taken due to the limits on the resolution set by the collimator. A lower 798 spatial resolution system, one with a better energy resolution, should be the 799 imaging system of choice. The spectral performance of gamma-cameras based 800 on the PMT (Section 3) cannot be reached by current EM-CCD based systems 801 in which the limit with CsI(Tl) has been shown to be approximately 20-30% 802 using americium-241 with a 1:1 fibre-optic plate. Using scintillators with a 803 higher light output would improve the spectral resolution but the increase in 804 noise from the multiplication gain process cannot yet be overcome. 805

If one is not limited spatially by external sources, then the scintillator-coupled 806 EM-CCD should be seriously considered. Imaging for synchrotron-based appli-807 cations where high resolution, high sensitivity and a high signal-to-noise ratio 808 are required may benefit greatly from the improved spatial resolution when 809 low fluxes are required to be measured to a high positional accuracy, although 810 care would be required in appropriately shielding the detector from direct de-811 tection to prevent damage to the EM-CCD and the electronics. For example, 812 in macromolecular crystallography, the high spatial resolution will bring the 813 ability to better resolve closely lying distributions of peaks whilst improv-814 ing the signal-to-noise ratio for low intensity peaks. In systems in which the 815 integration imaging mode is predominantly used, the improvements through 816 photon-counting with such a detector allow the basic use of energy discrimina-817 tion that would not otherwise be possible. However, the energy resolution of a 818 high-resolution gamma-camera such as this is limited by the processes through 819 which the high spatial resolution is achieved. By using only a small proportion 820 of the visible photons emitted by the scintillator for each event (to keep the 821 spatial information in the form of the Gaussian-like event profile), the noise on 822 the detected signal is comparatively high and is increased further still through 823 the gain process of the EM-CCD (required to keep the effective readout noise 824 low at the high frame-rates required for photon-counting imaging). 825

In systems in which the energy of the incident X-rays is lower than the K binding energy of the scintillator elements (less than approximately 30 keV for CsI), the problems associated with internal ionisation are no longer present and the scintillator-coupled EM-CCD is expected to provide further improvements in spatial resolution and MTF.

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