Introduction

Single photon emission computed tomography (SPECT) is a nuclear medicine imaging modality where two-dimensional projections are acquired with a gamma camera and the projections are used for reconstruction of a three-dimensional image volume. The projections are distorted by several factors, including attenuation and scattering of gamma ray, collimator structure, data acquisition, reconstruction method, and organ motions. The collimator in SPECT is a crucial component of the imaging chain that controls the noise, resolution, and sensitivity of the final functional image. Specialized collimators are used for a specific imaging task, for example the fan beam collimator for brain imaging, cone beam collimator for posterior portion of the thalamus, slit slat collimator for radionuclide imaging with high-energy gamma ray emission, and parallel-hole collimators that routinely use in SPECT and planar imaging. The pinhole and multi-pinhole collimators used for research, preclinical study, small animal, and focal uptake nuclear medicine imaging. The main objective of this research is to review the pinholes and multi-pinholes that recently have been used in clinical and preclinical nuclear medicine imaging.

Pinhole Collimator

Pinhole collimators have a single hole that drilled into the sheet with high atomic number material. Lead and...
tungsten generally used for fabricating the pinhole collimator. This collimator generates magnified images.\textsuperscript{[13]} Smaller pinhole diameter leads to an improvement the spatial resolution, but also a loss in sensitivity.\textsuperscript{[14]} Pinhole collimators are widely used for SPECT imaging of small organ or small region of interest such as the thyroid,\textsuperscript{[15]} the parathyroid glands,\textsuperscript{[16,17]} the breast,\textsuperscript{[18-22]} the knee joints and shoulder,\textsuperscript{[23,24]} or physiological imaging for the small animals.\textsuperscript{[25-28]} Pinhole collimator is used to obtain a superior spatial resolution compared to conventional parallel-hole collimator [Figure 1].\textsuperscript{[30,31]} Pinhole collimator provides a smaller field of view (FOV) in comparison with a parallel-hole collimator, so these collimators are suited to image the small organs of the body or focal uptake.\textsuperscript{[32]} In the pinhole collimator, penetration and scattering of gamma radiation at the edge of the aperture, also the attenuation coefficient of the aperture material have a significant effect on the spatial resolution of the system.\textsuperscript{[33,34]} Small pinhole opening angle makes reduce penetration and short object to collimator distance improve sensitivity and spatial resolution.\textsuperscript{[35]} The sensitivity of a parallel-hole collimator is almost constant as a function of distance from the collimator; however, the sensitivity of a pinhole collimator depends on the inverse square of the distance from the pinhole aperture. It also increases as the square of the pinhole diameter with simultaneous linearly loss in spatial resolution. The magnification factor of a pinhole is determined by the ratio of the pinhole-to-detector distance relative to the pinhole-to-source distance.\textsuperscript{[36]} With increased magnification factor, the detector intrinsic resolution has decreased effect on the total system spatial resolution. Therefore, a pinhole collimator is the most effective for imaging a small object placed close to the pinhole aperture.\textsuperscript{[37]}

**Multi-Pinhole Collimator**

Multi-pinhole collimator is used for superior spatial resolution trade off sensitivity compared to pinhole collimator. Since the single pinhole collimator suffers from the low detection efficiency, it has been investigated to increase the sensitivity and FOV with preserved spatial resolution. In multi-pinhole collimator, multiple projections through the different pinholes provide relatively efficient coverage of the detector area while multiplexing effect can make a problem for image reconstruction. If the projections from different apertures overlap, then the multiplexing artifacts occurred. The primary limitation in multi-pinhole collimator design includes the overlap of the projections from different pinholes.\textsuperscript{[38,39]} Irregular apertures configuration could suppress this artifact.\textsuperscript{[40]} In some cases, perpendicular lead septa were employed to acquire nonoverlapping projections.\textsuperscript{[41,42]} Multi-pinhole collimation was used in the early days of nuclear cardiology to provide multiple angle views of the myocardium.\textsuperscript{[43,44]} With advances in iterative image reconstruction, multi-pinhole collimators have made a come-back in nuclear cardiology.\textsuperscript{[45]} The new model assumes that the multi-pinhole collimator and the detector with spherical layer shape can optimal resolution and sensitivity.\textsuperscript{[46]} Multi-pinhole SPECT provides for sub-millimeter resolution imaging of the radio molecule distribution in small laboratory animals.\textsuperscript{[47]}

**Pinhole Geometry**

The most important reasons of error in quantitative pinhole SPECT are the photon penetrating throughout the edge of the pinhole aperture, photon scattered by the object, photon scattered by the collimator material, and attenuation of photons within the object. The magnitudes of these components highly depend on collimator geometry.\textsuperscript{[48]} Two types of pinhole collimators commonly used in SPECT imaging, channel-edge, and knife-edge shaped [Figure 2]. The channel-edge pinholes were designed in order to reduce the penetration of gamma rays through the edge of the pinhole aperture caused by the large acceptance angle of knife-edge pinhole.\textsuperscript{[49,50]} Compared to the channel-edge, the penetration, object scatter, and collimator scatter for knife-edge are higher than channel-edge.\textsuperscript{[48]} At perpendicular incidence of the photons, the channel-edge has lower penetration and scatter fractions rather than knife-edge.\textsuperscript{[51]} The advantage from channel-edge pinhole of lower penetration and scatter contributions significantly diminishes at nonperpendicular incident. Furthermore, the channel-edge pinholes have a lower

![Figure 1: The images of $^{99m}$Tc-sestamibi from thyroide at 15 min in a patient with hyperparathyroidism acquired with (a) pinhole and (b) parallel-hole collimators](image1)

![Figure 2: Two types of the pinhole collimators that routinely used in pinhole single photon emission computed tomography imaging: (a) Knife-edge collimator and (b) channel-edge pinhole collimator](image2)
sensitivity than knife-edge pinholes. By increasing the pinhole diameter, the penetration and the scattered photon approximately decreased by the linear function for the knife-edge micro-pinhole. For the knife edge micro-pinhole, platinum and gold are superior rather than tungsten and lead with regard to pinhole scatter while platinum performs slightly better than gold. Song et al. showed the penetration and the scatter fraction for pinhole collimators with a tungsten aperture in channel shape is preferable over knife edge and the optimal range of channel height have an average from 0.3 to 0.6 mm. I imaging is challenging because the primary photon is emitted at 364 keV, and the penetration through the pinhole aperture is significant, which will degrade SPECT image resolution. The tungsten pinhole with channel edge shapes has an advantage over lead for high resolution Imaging due to reduced photon penetration near the pinhole aperture. Briefly, the purpose of designing a channel height collimator is to reduce photon penetration near the pinhole aperture.

**Reconstruction Algorithms for Pinhole Single Photon Emission Computed Tomography**

The most reconstruction algorithms used in pinhole imaging are the filtered back projection (FBP) method proposed by Feldkamp, Davis, and Kress (FDK). Ramp filter routinely used during the reconstruction. If a FBP method was used for pinhole SPECT combined with mechanical shift correction, the doughnut type artifact could correct. The pinhole SPECT data with a circular orbit acquisition do not satisfy the data sufficiency condition, so pinhole SPECT restricted by artifacts, mainly axial blurring in the noncentral slices. The iterative reconstruction algorithm can reduce these artifacts in comparison with FBP. The expectation maximization (EM) algorithm could improve the quality of pinhole SPECT, but the long reconstruction times make this method less practical in clinical usage in comparison with FDK algorithm. The maximum likelihood EM (ML-EM) reconstruction algorithm can use in the pinhole SPECT. This method improves image quality compared with FBP, but the major drawback of ML-EM is increased noise along the iteration. Postreconstruction filters are used to remove noise, but these filters reduce the spatial resolution, contrast and have a tendency to blur image edge. The noise created by ML-EM can also reduce using Bayesiam method. The median root prior (MPR) is a new Bayesian reconstruction algorithm which has mainly used in positron emission tomography. MPR is adapted for pinhole SPECT and can reduce the noise generated from ML-EM algorithm. The ordered subsets EM (OS-EM) algorithm is a type of iterative reconstruction algorithm, and its function depends on the number of subset and iteration. The processing time with one iteration and eight subsets is only 2.5 times longer compared with FDK. This algorithm significantly improves the reconstructed image quality, also suppresses the streak artifacts in comparison with FBP. The primary attraction of OS-EM versus FBP is the absence of noise amplification. The insufficient projection data acquired by a single circular pinhole orbit are the mainly cause of blurring. A pinhole SPECT by two circular orbits and a three-dimensional OS-EM can provide complete projection data, furthermore this method disappears the blurring edge in the reconstructed image. This imaging method can improve axial blurring and resolution uniformity in pinhole SPECT imaging and have a potential to become the standard tool for quantification of physiological functions in small animal laboratory.

The main goal of SPECT in preclinical molecular imaging by multi-pinhole collimator is to measure the concentration radiolabeled biomolecules in vivo, so the quantification is important. Accuracy of the reconstructed image is not sufficient due to the partial volume effect. Anatomy-based reconstruction using micro-computed tomography information has ability to decrease the partial volume effect and improve the quantifying accuracy in multi-pinhole SPECT.

**The New Configuration of Pinhole and Multi-Pinhole Collimator**

A blended collimator uses the two different collimators, pinhole and multi-pinhole collimator, installed on each detector in the SPECT imaging system. The specific reconstruction software applied to create a single image that combined from the two collimators data acquisitions, so a high quality image can be achieved. Most multi-pinhole collimators prevent the multiplexing...
effect by using an additional shielding or by spacing far apart the pinholes, but additional shielding has the drawback of increase weight, designing complexity and cost, also spacing far apart the pinholes causes the noneffective use of detector. This is due to the circular projections from pinholes on the detector. New pinhole collimator geometry, the lofthole collimator, has a circular aperture and a rectangular entrance/exit opening, so it has a rectangular projection on the detector. The lofthole collimator irradiates precisely the surface of the square detector [Figure 4]. This has the benefit that no extra shielding is necessary in case of a multi-pinhole collimator and that the amount of penetrating photons is lower than with an equivalent pinhole. The lofthole collimator has less penetration than the knife-edge pinhole collimator. Multi-lofthole collimator has high detector coverage without additional shielding. The most advantage of this collimator is the lower amount of penetration compared to multi-pinhole collimator [Figure 5]. The drawback of the pinhole and multi-pinhole collimators includes cone beam imaging geometry, so that does not provide a complete data as the detector rotates around the object. For reducing the data-insufficiency artifacts, the SPECT imaging needs the small cone angle in the axial direction. A multi skew-slit collimator has a single vertical slit and several horizontal slits. The slits prepare separate projections with small cone angle and small magnification in the axial direction, which makes less severe data-insufficiency artifacts; also partly larger cone angle and magnification in the transaxial direction provide a better detector area usage comparison with multi-pinhole collimator [Figure 6]. Thus, multi skew-slit collimator provided an appropriate transaxial resolution, detector area usage, and minimum multiplexing effect.

**Conclusion**

Collimator is a very important component in SPECT as critical equipment in sensitivity and resolution of the system. Many types of collimators are used for nuclear medicine imaging. Pinhole collimators are used for SPECT imaging of small animals or small organs like thyroid. Pinhole collimator often provides a superior resolution-sensitivity tradeoff for radionuclide imaging compared to SPECT with parallel-hole collimator. Multi-pinhole collimator has been invented to increase the FOV, sensitivity, and detection efficiency without degrading spatial resolution. Channel-edge and knife-edge pinhole collimators are the routinely collimators that used in pinhole SPECT imaging. The magnitude of penetration and scatter for channel-edge or Knife-edge pinhole collimators depends on the collimator material and the energy of radionuclide. The type of reconstruction algorithms in pinhole SPECT imaging is very important because the quality of the final image is relevant to this item. The most reconstruction algorithm that used in pinhole SPECT is the FDK FBP algorithm. The interested reconstruction algorithm for utilizing

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**Figure 4:** Schematic of (a) a pinhole collimator and (b) a lofthole collimator with a circular aperture and a rectangular entrance/exit opening. This type of collimator creates the rectangular projection on the detector.

**Figure 5:** Projections from (a) a multi-pinhole collimator that do not fully coverage the detector, (b) multiplexing effect in a multi-pinhole collimator, and (c) projections created from a multi lofthole collimator without the multiplexing effect.
in pinhole SPECT imaging is OS-EM algorithm. This algorithm significantly improves reconstructed image quality, also suppresses streak artifacts in comparison with FBP. To eliminate the multiplexing effect in multi-pinhole collimators, the additional shielding or spacing the pinholes has been used but these adjustment methods themselves have some disadvantages. The new collimators such as skew slit and lothole collimators produce noncircular projections; therefore, provide high detector coverage, spatial resolution, and sensitivity without multiplexing effect.

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