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(54) Title: APPARATUS FOR GENERATING MULTI-ENERGY X-RAY IMAGES AND METHODS OF USING THE SAME

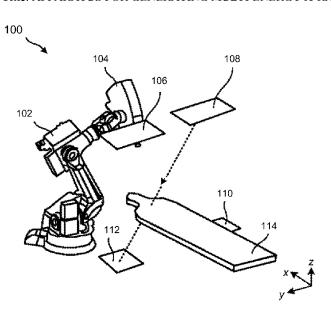
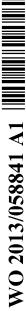


FIG. 1

(57) Abstract: Apparatus for generating multi-energy x-ray images and systems and methods for processing and using multi-energy x-ray images are provided for treating a target, e.g., a tumor, for image-guided radiation therapy. An image-guided radiation therapy system is provided including a treatment delivery system, a multi-energy imaging system, and a system processor. The treatment delivery system includes a radiation source configured to generate treatment radiation beams. The multi-energy imaging system is configured to generate a first set of image data of the patient using imaging radiation at a first energy level and a second set of image data of the patient using imaging radiation at a second energy level. The system processor is configured to process the first and second sets of image data to generate an enhanced image of the patient and to direct the treatment delivery system based on information obtained from the enhanced image.



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APPARATUS FOR GENERATING MULTI-ENERGY X-RAY IMAGES AND METHODS OF USING THE SAME

I. Field Of The Invention

[0001] This application relates to apparatus for generating multi-energy x-ray images and systems and methods for using multi-energy x-ray images in image-guided radiation therapy.

II. Background Of The Invention

[0002] Oncology is the branch of medicine directed to the study of the development, diagnosis, treatment, and prevention of tumors. A tumor is an abnormal growth of tissue serving no physiological function. A tumor may be malignant (cancerous) or benign. A malignant tumor may exhibit uncontrolled, progressive multiplication of cells and spread cancerous cells to other parts of the body (metastasizes) through blood vessels or the lymphatic system. A benign tumor does not metastasize, but can still be life-threatening if it impinges on critical body structures such as nerves, blood vessels, and organs.

[0003] Radiosurgery and radiotherapy are radiation treatment systems that use external radiation beams to treat tumors and other lesions by delivering a prescribed dose of radiation (e.g., x-rays, protons, or gamma rays) to a target volume (region of interest, or ROI) while minimizing radiation exposure to the surrounding tissue. The object of both radiosurgery and radiotherapy is the destruction of abnormal tissue while sparing healthy tissue and critical structures. Radiotherapy is characterized by a low radiation dose per treatment and many treatments (e.g., 30 to 45 days of treatment). Radiosurgery is characterized by a relatively high radiation dose to a tumor in one, or at most a few, treatments. In both radiotherapy and radiosurgery, the radiation dose is delivered to the tumor site from multiple angles. As the angle of each radiation beam is different, each beam passes through a different area of healthy tissue on its way to the tumor. As a result, the cumulative radiation dose at the tumor

is high, while the average radiation dose to the surrounding healthy tissue is low. Unless specified otherwise, radiosurgery and radiotherapy are used interchangeably in the present application.

[0004] Radiation treatment systems may be used together with an imaging system for image-guided radiation therapy (IGRT). The imaging system acquires in-treatment images, e.g., x-ray, ultrasound, CT, or PET, that may be used to for patient set up and in some instances (e.g., Accuray Incorporated's CyberKnife® Radiosurgery System) guide the radiation delivery procedure and track in-treatment target motion. Target motion tracking may be accomplished by correcting for differences in target position by acquiring and registering intratreatment images with reference images, known as digitally reconstructed radiographs (DRRs), rendered from a pre-treatment computed tomography (CT) scan, which may otherwise be known as the treatment planning image.

[0005] Previously-known IGRT systems use imaging systems that generate single energy x-ray images during treatment. Such systems suffer, however, from a variety of drawbacks. For example, x-ray attenuation characteristics are dependent on x-ray energy and thus a single energy x-ray image may have limited differentiation ability for certain materials. An x-ray image generated using a low x-ray energy (e.g., $\sim 50\text{-}100 \text{ kV}$) will display significant attenuation in soft tissue and radio-opaque objects such as skeletal structures, fiducials, and contrast agents. Conversely, an x-ray image generated using a high x-ray energy (e.g., $\sim 100 \text{ kV} - 6 \text{ MV}$, preferably 100-150 kV) will display less attenuation of soft tissue than a low x-ray energy image, but still significant attenuation in radio-opaque objects such as skeletal structures, contrast agents, and fiducials.

[0006] In view of the above-noted drawbacks of previously-known systems, it would be desirable to provide apparatus and methods for generating x-ray images with increased discrimination between soft tissue and radio-opaque objects.

[0007] It further would be desirable to provide systems and methods for processing and using such x-ray images in image-guided radiation therapy.

III. Summary Of The Invention

[0008] The present invention overcomes the drawbacks of previously-known imageguided radiation therapy (IGRT) systems by providing apparatus for generating multi-energy

x-ray images and systems and methods for processing and using multi-energy x-ray images in IGRT. The image-guided radiation therapy system of the present invention includes a treatment delivery system having a radiation source configured to generate treatment radiation beams and a collimator configured to collimate the treatment radiation beams, a multi-energy imaging system, and a system processor. The multi-energy imaging system is configured to generate a first set of image data of part or all of the patient using imaging radiation at a first energy level, e.g., an energy level within about 50-100 kV, and a second set of image data of part or all of the patient using imaging radiation at a second energy level, e.g., an energy level within about 100 kV – 6 MV, preferably 100-150 kV. The system processor is configured to process the first and second sets of image data to generate an enhanced image, e.g., a CT or tomosynthetic enhanced image, of part or all of the patient and to direct the treatment delivery system based on information obtained from the enhanced image.

[0009] The multi-energy imaging system may generate a set(s) of image data of part or all of the patient that may include the target, a skeletal structure within the patient, a soft tissue structure within the patient, a contrast agent within the patient, and/or fiducials implanted in the target/patient. The system processor may direct the treatment delivery system based on information obtained from the enhanced image, such as the position(s) of the target, the skeletal structure, the soft tissue structure, the contrast agent, and/or the fiducials. The system processor may direct the treatment delivery system to position the radiation source, position the patient, and/or enable or disable the treatment radiation beams based on such information. The treatment delivery system may include a robot arm, e.g., a six-degreeof-freedom robot arm, or a gantry to position the radiation source. The collimator of the treatment delivery system may be a fixed collimator or a variable aperture collimator having an aperture and may be a multileaf collimator. The system processor may direct the treatment delivery system to set the aperture based on information obtained from the enhanced image, to change the diameter of a circular field approximated by the variable aperture collimator, to move leaves of the multileaf collimator, to move the collimator, and/or to position the patient.

[0010] The set(s) of image data generated by the multi-energy imaging system may be volumetric image data, such as tomosynthetic image data or CT image data. In an alternative embodiment, first and second sets of image data may be first and second pluralities of two-

dimensional projection image data and the processor may be configured to generate a plurality of intermediate enhanced images from first and second pluralities of two-dimensional projection image data and to process the plurality of intermediate enhanced images to generate the enhanced image.

[0011] The multi-energy imaging system may include first and second x-ray sources configured to emit imaging radiation towards part or all of the patient and first and second x-ray detectors configured to receive the imaging radiation after at least a portion of the imaging radiation has passed through part or all of the patient. In a preferred embodiment, the first x-ray source emits the imaging radiation at the first energy level and the second x-ray source emits the imaging radiation at the second energy level. In one embodiment, the first x-ray source emits the imaging radiation at the first energy level and the second x-ray source substantially simultaneously emits the imaging radiation at the second energy level and the second x-ray source substantially simultaneously emits the imaging radiation at the first energy level and the second x-ray source substantially simultaneously emits the imaging radiation at the first energy level. In an alternative embodiment, the first and second x-ray sources emit radiation at the first energy level substantially simultaneously and, subsequently, the first and second x-ray sources emit radiation at the second energy level.

[0012]The multi-energy imaging system may include an x-ray source array having a plurality of emission spots. A first emission spot of the plurality of emission spots may be configured to emit imaging radiation at the first energy level, and optionally subsequently at the second energy level, and a second emission spot of the plurality of emission spots that may be configured to emit imaging radiation at the second energy level, and optionally subsequently at the first energy level. In one embodiment, the plurality of emission spots are configured to emit imaging radiation at the first energy level and to subsequently emit imaging radiation at the second energy level. The multi-energy imaging system may further include a second x-ray source array having a plurality of emission spots configured to emit imaging radiation in-phase or out of phase with the emission spots of the first x-ray source array. In one embodiment, the plurality of emission spots of the first x-ray source array are configured to emit imaging radiation at the first energy level, and optionally subsequently at the second energy level, and the plurality of emission spots of the second x-ray source array are configured to emit imaging radiation at the second energy level, and optionally subsequently at the first energy level. In another embodiment, the plurality of emission spots

of the first x-ray source array and the plurality of emission spots of the second x-ray source array are configured to emit imaging radiation at the first energy level and to subsequently emit imaging radiation at the second energy level.

[0013] In accordance with one aspect of the present invention, the multi-energy imaging system may include an x-ray source filter for generating multi-energy x-ray images. The x-ray source filter has first and second portions configured to filter the imaging radiation emitted from the first x-ray source. The first and second portions may comprise different materials, e.g., aluminum and copper, or may comprise the same material, but have different thicknesses. The x-ray source filter may be configured to move between a first position and a second position. At the first position, the first portion of the filter is adjacent to the first x-ray source such that imaging radiation emitted from the first x-ray source travels through the first portion and exits the first portion at the first energy level. At the second position, the second portion is adjacent to the first x-ray source such that imaging radiation emitted from the first x-ray source travels through the second portion and exits the second portion at the second energy level. Advantageously, the x-ray source filter may be used together with a conventional x-ray source for generating multi-energy x-ray images.

imaging system may include an x-ray detector filter disposed between the first x-ray source and the first x-ray detector or integrated within the first x-ray detector. The x-ray detector filter has first and second portions that may comprise different materials, e.g., aluminum and copper or copper and no material, or may comprise the same material, but having different thicknesses. Imaging radiation emitted from the first x-ray source travels through the first portion and exits the first portion at the first energy level and imaging radiation emitted from the first x-ray source travels through the second portion and exits the second portion at the second energy level. After traveling through the x-ray detector filter, the imaging radiation is received by the first x-ray detector. The x-ray source may be a broad energy x-ray source and the x-ray detector may be an energy-binning photon counting detector. Beneficially, the x-ray source filter may be used together with a conventional x-ray source and a conventional x-ray detector for generating multi-energy x-ray images. In one embodiment, the filter is configured to be selectively based on a projection of the target on a detector plane.

[0015] The present invention provides systems and methods for acquiring and processing multi-energy x-ray image data in image-guided radiation therapy. For acquisition, the system

may selectively image a material of interest, with the energy level(s) for imaging determined by a method that involves a phantom and calibration. The processing may enhance image definition selectively for materials of interest, e.g. bone, tissue, muscle, in interventional images by exploiting beam hardening effects using High Dynamic Range (HDR) x-ray imaging. The system processor of the IGRT system may be configured to determine an optimal energy level for imaging part or all of the patient, e.g., lung mass, using, for example, a phantom. The multi-energy imaging system generates the first set of image data of part or all of the patient using imaging radiation at the optimal energy level and the second set of image data of the target using imaging radiation at an adjusted energy level. The system processor of the IGRT system is configured to process the first and second sets of image data to generate an enhanced image that may be a High Dynamic Range (HDR) x-ray image. An HDR x-ray image includes more visual information, derived from multiple energy levels, than a single energy level image. An HDR x-ray image may further be combined with a single energy level to extract high definition images of selected materials.

[0016] The present invention further provides systems and methods for registering multienergy x-ray image data for image-guided radiation therapy. Preferably, the image-guided radiation therapy system further includes a diagnostic image processor configured to generate digitally reconstructed radiographs (DRRs) from treatment planning images of part or all of the patient. The system processor is configured to register the first and second sets of image data and/or the enhanced image with corresponding DRRs to obtain a registration result.

[0017] The systems and methods of the present invention may also be used to track the target(s), skeletal structure(s), soft tissue(s), contrast agent(s), fiducial(s), and patient movement and/or position during image-guided radiation therapy. For enhanced tracking, the IGRT system may use the system processor to process image data wherein the enhanced image comprises a weighted combination of the first and second sets of image data. The enhanced image may comprise a weighted combination in logarithmic space of the first and second sets of image data and/or a spatially varying weighted combination of the first and second sets of image data. The enhanced image may also be a weighted subtraction of the first and second sets of image data and/or a weighted subtraction in logarithmic space of the first and second sets of image data. The enhanced image may be an x-ray image displaying primarily soft tissue without radio-opaque objects such as skeletal structures, fiducials, and contrast agents. Alternatively, the enhanced image may be an x-ray image displaying

primarily radio-opaque objects without soft tissue. The system processor may be configured to track movement and position of the target(s), skeletal structure(s), soft tissue(s), contrast agent(s), and/or fiducial(s) and patient using the enhanced image. Additionally, the system processor may be configured to track movement and position of the target(s), skeletal structure(s), soft tissue(s), contrast agent(s), fiducial(s), and patient by registering the first and second sets of image data or the enhanced image with corresponding DRRs.

Advantageously, tracking the target with multi-energy x-rays images may provide accurate target location during IGRT for enhanced accuracy of treatment radiation delivery from the treatment delivery system to the target. Furthermore, multi-energy x-ray images may improve tumor visibility and offer the ability to track anatomical targets that were not clearly visible in conventional x-ray images.

[0018] In one embodiment, the image-guided radiation therapy system may use motion artifacts to track the target(s), skeletal structure(s), soft tissue(s), contrast agent(s), fiducial(s), and patient movement and/or position during image-guided radiation therapy. In this embodiment, the system processor may be configured to subtract an overlapping portion of the first set of image data from the second set of image data to generate the enhanced image having a motion-edge artifact image. The system processor then may track movement and position of the target(s), skeletal structure(s), soft tissue(s), contrast agent(s), fiducial(s), and patient using the motion-edge artifact image.

[0019] In another embodiment, the image-guided radiation therapy system is used to track the target(s), skeletal structure(s), soft tissue(s), contrast agent(s), fiducial(s), and patient movement and/or position during respiration. Preferably, the multi-energy imaging system is configured to generate the first set of image data at a position in the patient's respiratory cycle and to generate the second set of image data at the same position in a subsequent respiratory cycle. The image-guided radiation therapy system may include a non-x-ray based position sensing system configured to generate a respiratory motion model to predict target location.

[0020] In accordance with one aspect of the invention, a method for treating a target within a patient for image-guided radiation is provided. The method includes generating a first set of image data of part or all of the patient using imaging radiation at a first energy level and a second set of image data of part or all of the patient using imaging radiation at a second energy level, processing the first and second sets of image data to generate an

enhanced image, and directing a treatment delivery system based on information obtained from the enhanced image. Preferably, a machine-readable medium is provided that includes instructions that, when executed by a processor of an image-guided radiation therapy system, cause the processor to perform the method. Additionally, the IGRT system may execute a programmed routine configured to process a first set of image data of part or all of the patient generated using imaging radiation at a first energy level and a second set of image data of part or all of the patient generated using imaging radiation at a second energy level to generate an enhanced image, and to direct the treatment delivery system based on information obtained from the enhanced image using, for example, the system processor. The IGRT system may modify the radiation treatment by activating or deactivating the treatment beam based on the enhanced image using, for example, the system processor in conjunction with constraints on the allowable range of target motion defined when a treatment plan was created.

[0021] In accordance with another aspect of the present invention, a multi-energy imaging system is provided for, for example, medical applications or industrial applications such as cargo scanning. The multi-energy imaging system may include an x-ray source array configured to generate a first set of image data using imaging radiation at a first energy level and a second set of image data using imaging radiation at a second energy level, and a system processor configured to process the first and second sets of image data to generate an enhanced image. The multi-energy imaging system may further include a second x-ray source array positioned at a non-zero angle from the x-ray source array for stereoscopic imaging.

IV. Brief Description Of The Drawings

[0022] FIG. 1 illustrates a perspective view of a previously-known image-guided radiation therapy (IGRT) system.

[0023] FIG. 2 illustrates a perspective view of an IGRT system and a schematic diagram of a computer system in accordance with the principles of the present invention.

[0024] FIGS. 3A and 3B are, respectively, a side view and a plan view of an exemplary x-ray source filter for generating multi-energy x-ray images.

[0025] FIGS. 4A and 4B are, respectively, a side view and a plan view of an alternative x-ray source filter for generating multi-energy x-ray images.

- [0026] FIG. 5A is a side view of yet another x-ray source filter for generating multienergy x-ray images.
- [0027] FIG. 5B is a plan view of the x-ray source filter of FIG. 5A at a first position.
- [0028] FIG. 5C is a plan view of the x-ray source filter of FIG. 5A at a second position.
- [0029] FIGS. 6A and 6B are, respectively, an exploded view and an assembled view of an exemplary x-ray detector filter assembly for generating multi-energy x-ray images.
- [0030] FIG. 7 is a schematic view of an exemplary IGRT system for use with multienergy x-ray images.
- [0031] FIG. 8A and 8B are x-ray images generated by an exemplary real-time multienergy imaging system.
- [0032] FIGS. 8C and 8D are x-rays images processed by an exemplary system processor.
- [0033] FIG. 9 illustrates an exemplary method for processing multi-energy x-ray image data in IGRT.
- [0034] FIG. 10 is a plot illustrating change in x-ray spectra after traversal through a filter for obtaining an optimal energy level.
- [0035] FIG. 11 illustrates an exemplary method for registering multi-energy x-ray images in IGRT.
- [0036] FIG. 12 illustrates an exemplary method for target tracking using multi-energy x-ray images in IGRT.
- [0037] FIG. 13 illustrates an exemplary method for target tracking using a motion-edge artifact in IGRT.
- [0038] FIGS. 14A and 14B are illustrations of x-ray images of a target at first and second positions, respectively.

[0039] FIG. 14C is an illustration of a motion-edge artifact image generated after subtracting overlapping portions of the target at the first and second positions.

[0040] FIG. 15 illustrates a method for target tracking during respiration using multienergy x-ray images in IGRT.

V. <u>Detailed Description Of The Invention</u>

[0041] This disclosure relates to apparatus for generating multi-energy x-ray images and systems and methods for processing and using multi-energy x-ray images in image-guided radiation therapy. Advantageously, x-rays generated at an energy level within a low energy range (e.g., $\sim 50\text{-}100 \text{ kV}$) may be enhanced with x-rays generated at an energy level within a high x-ray energy range (e.g., $\sim 100 \text{ kV} - 6 \text{ MV}$, preferably 100-150 kV) to provide x-ray images having high definition for images of soft tissue, and radio-opaque objects such as skeletal structures, fiducials, and contrast agents. Moreover, x-rays generated within the low energy range may be enhanced with x-rays generated within the high x-ray energy range in a weighted manner to provide x-ray images without soft tissue or, alternatively, x-rays images without radio-opaque objects. The enhanced x-ray images may be used for enhanced target tracking and positioning for image-guided radiation therapy.

[0042] FIG. 1 illustrates a perspective view of a previously-known image-guided radiation therapy (IGRT) system 100. IGRT system 100 includes articulated robot arm 102, MV radiation source 104, x-ray sources 106 and 108, x-ray detectors 110 and 112, and treatment table 114. An example of IGRT system 100 is a CYBERKNIFE® Robotic Radiosurgery System available from Accuray, Incorporated, Sunnyvale, California.

[0043] In FIG. 1, MV radiation source 104 is configured to generate treatment radiation beams, e.g., x-ray photon, electron, or proton beams, at a target volume, e.g., a tumor, within a patient. MV radiation source 104 generally includes a linear accelerator (LINAC). MV radiation source 104 is mounted on the end of articulated robot arm 102 to provide multiple (e.g., 5 or more) degrees of freedom of motion in order to position MV radiation source 104 to irradiate tumorous tissue with highly-collimated beams delivered from many angles in an operating volume (e.g., sphere) around the patient. Treatment may involve beam paths with a single isocenter, multiple isocenters, or with a non-isocentric approach (e.g., the beams need only intersect with the targeted tumor mass and do not necessarily converge on a single point, or isocenter, within the target region). Treatment can be delivered in either a single session

(mono-fraction), in a small number of sessions (hypo-fractionation), or in a large number (30-40) of sessions (standard fractionation) as determined during treatment planning.

IGRT system 100 has a real-time imaging system that includes first x-ray source 106, second x-ray source 108, first x-ray detector 110, and second x-ray detector 112. First x-ray source 106 is paired with first x-ray detector 110 to establish a first "channel" of a stereoscopic x-ray imaging system, and second x-ray source 108 is paired with second x-ray detector 112 to establish a second "channel." During radiation treatment, first and second x-ray sources 106 and 108 emit single-energy x-ray imaging radiation that travels through a patient and treatment table 114 and is received by first and second x-ray detectors 110 and 112, respectively. The in-treatment, single-energy x-ray images are used to set up and guide the treatment radiation delivery procedure and track in-treatment target motion. The target is tracked by correcting for differences in target position in the in-treatment, single-energy x-ray images and registering them with reference images, known as digitally reconstructed radiographs (DRRs), rendered from a pre-treatment computed tomography (CT) scan.

[0045] Applicants have concluded that higher definition in-treatment x-ray images may be generated by providing apparatus for generating multi-energy x-ray images. Further, applicants have discovered systems and methods for processing and using multi-energy x-ray images in image-guided radiation therapy.

Apparatus For Generating Multi-Energy X-Ray Images

[0046] FIG. 2 illustrates a perspective view of image-guided radiation therapy (IGRT) system 200 and a schematic diagram of computer system 250 in accordance with the principles of the present invention. IGRT system 200 includes articulated robot arm 102, MV radiation source 104, x-ray sources 206 and 208, x-ray detectors 210 and 212, and treatment table 114. Articulated robot arm 102, MV radiation source 104, and treatment table 114 may be conventional and, thus, are not described in detail.

[0047] Advantageously, IGRT system 200 has a real-time imaging system configured to generate multi-energy x-ray images. The real-time imaging system includes first x-ray source 206, second x-ray source 208, first x-ray detector 210, and second x-ray detector 212. First x-ray source 206 is paired with first x-ray detector 210 to establish a first "channel" of a stereoscopic x-ray imaging system, and second x-ray source 208 is paired with second x-ray detector 212 to establish a second "channel." During radiation treatment, first and second x-

ray sources 206 and 208 emit imaging radiation that travels through part or all of a patient and treatment table 114 and is received by first and second x-ray detectors 210 and 212, respectively.

[0048] First and second x-ray sources 206 and 208 may include kV x-ray tube technology, x-ray source array technology, broad energy technology, or other suitable x-ray source technology. X-ray source array refers to a source of x-rays comprising a plurality of spatially distinct, electronically controlled x-ray emitters or emission spots (focal spots) that are addressable on at least one of an individual or groupwise basis such as an x-ray source array available from Triple Ring Technologies (Newark, California) or XinRay Systems (Research Triangle Park, North Carolina). First and second x-ray detectors 210 and 212 may be amorphous silicon detectors or other suitable detectors capable of producing high-quality two-dimensional x-ray images during an IGRT. X-ray sources 206 and 208 may be mounted in or near the ceiling of a treatment vault, while x-ray detectors 210 and 212 may be mounted in or near the floor of the treatment vault as illustrated, although the scope of the preferred embodiments is not limited thereto. It should be noted that although the real-time imaging system of IGRT system 200 is illustrated as having two x-ray sources and two x-ray detectors, IGRT system 200 is not limited thereto. For example, IGRT system 200 may have one, three, four or more x-ray sources and one, three, four or more x-ray detectors.

[0049] In accordance with one embodiment, first x-ray source 206 is configured to emit imaging radiation at an energy level within a low x-ray energy (e.g., \sim 50-100 kV). An x-ray generated from low energy radiation has significant attenuation in soft tissue and radio-opaque objects and the resulting image will display soft tissue in conjunction with radio-opaque objects. In this embodiment, second x-ray source 208 is configured to emit radiation at an energy level within a high x-ray energy range (e.g., \sim 100-150 kV). An x-ray generated from high energy radiation has greater attenuation in radio-opaque objects, but little attenuation in soft tissue, and the resulting image will display primarily radio-opaque objects without soft tissue.

[0050] After the emitted radiation travels through the patient, first x-ray detector 210 receives the low energy imaging radiation and second x-ray detector 212 receives the high energy imaging radiation. The received low and high energy imaging radiation then may be processed at computer system 250, described further below, to generate first and second sets of image data using suitable software and to produce real-time x-ray images having enhanced

definition for radio-opaque objects and soft tissue. In one embodiment, first and second x-ray detectors are energy-binning photon counting detectors.

[0051] In an alternative embodiment, first and second x-ray sources 206 and 208 may each alternate between low energy and high energy emission modes during respective periodic imaging cycles. In one embodiment, first and second x-ray sources 206 and 208 may be in-phase with each other (*i.e.*, both emitting at low energy, then both emitting at high energy, etc.), while for another embodiment, first and second x-ray sources 206 and 208 may be out of phase with each other (*i.e.*, one emitting at low energy while the other emits at high energy).

[0052] In accordance with another aspect of the present invention, the real-time imaging system of IGRT system 200 may be configured to provide multi-energy stereoscopic tomosynthesis images. X-ray tomosynthesis refers to the process of acquiring a number of two-dimensional x-ray projection images of a target volume using x-rays that are incident upon the target volume at a respective number of different angles, followed by the mathematical processing of the two-dimensional x-ray projection images to yield a set of one or more tomosynthetic reconstructed images representative of one or more respective slices of the target volume, wherein the number of x-ray projection images is less than that in a set that would be required for CT image reconstruction, and/or the number or range of incident radiation angles is less than would be used in a CT imaging procedure.

[0053] In this embodiment, first and second x-ray source-detector pairs (206/210, 208/212) are positioned to acquire tomosynthesis projection images over first and second non-overlapping projection angle ranges. First and second sets of tomosynthesis projection images of the target volume are acquired at distinct first and second x-ray energy levels, respectively (e.g., 80 kV and 140 kV), using the respective first and second x-ray tomosynthesis source-detector pairs.

[0054] The first and second sets of tomosynthesis projection images then are processed to generate respective first and second tomosynthesis reconstructed image sets of the target volume. Any of a variety of different tomosynthesis reconstruction algorithms may be used including, but not limited to, filtered backprojection (FBP), matrix inversion tomosynthesis (MITS), maximum likelihood expectation maximization (MLEM), and iterative ordered-subset convex (OSC) algorithms based on a maximum-likelihood models.

[0055] The first and second tomosynthesis reconstructed image sets then are processed in conjunction with each other on a locationwise basis (*e.g.*, voxelwise basis) within the target volume to generate a dual-energy processed image set. In one embodiment, the processing of the first and second tomosynthesis reconstructed image sets comprises registration (either by a known physical transformation between the imaging coordinate spaces or by image-based registration) and combination processing and/or other decomposition into soft-tissue and bone image components. Treatment radiation is delivered to the treatment target within the target volume based at least in part on the dual-energy processed image set.

[0056] Still referring to FIG. 2, computer system 250 is integrated with and/or coupled to IGRT system 200 using one or more buses, networks, or other communications systems 260, including wired and/or wireless communications systems, and being capable in conjunction therewith of implementing the methods of one or more of the preferred embodiments. Methods of image guided radiation treatment in accordance with one or more of the preferred embodiments may be implemented in machine-readable code (i.e., software or computer program product) and performed on computer systems such as, but not limited to, computer system 250. Computer system 250 includes central processing unit (CPU) 251 having microprocessor 252, random access memory 253, and nonvolatile memory 254 (e.g. electromechanical hard drive, solid state drive), display monitor 255, mouse 261, keyboard 263, and other I/O devices 256 capable of reading and writing data and instructions from machine-readable media 258 such as tape, compact disk (CD), digital versatile disk (DVD), blu-ray disk (BD), and so forth. In addition, computer system 250 may be connected via one or more buses, networks, or other communications systems 260 to other computers and devices, such as may exist on a network of such devices, e.g., Internet 259. Software to control the image guided radiation treatment steps described herein may be implemented as a program product and stored on a tangible storage device such as machine-readable medium 258, external nonvolatile memory device 262, cloud storage, or other tangible storage medium.

[0057] Referring now to FIGS. 3A and 3B, an x-ray source filter for generating multienergy x-ray images in accordance with the present invention is described. FIGS. 3A and 3B are, respectively, a side view and a plan view of x-ray source filter 300. X-ray source filter 300 includes motor 302, controller 304, shaft 306, and filter material 308. X-ray source filter

300 may be coupled to an x-ray source, e.g., x-ray source 106 of FIG. 1 as illustrated or x-ray sources 108, 206, 208, or any additional x-ray sources provided.

[0058] Motor 302 is configured to turn shaft 306 in response to commands from controller 304. Controller 304 communicates with computer system 250 (see FIG. 2) which determines desired motor characteristics, such as power and speed, using suitable software. Computer system 250 then transmits the desired motor characteristics to controller 304 using a wired or wireless communication link.

[0059] Shaft 306 is coupled to motor 302 and filter material 308 such that filter material 308 rotates when motor 302 turns shaft 306. Filter material 308 is made of at least two materials suitable for filtering x-ray radiation and is configured to filter imaging radiation emitted from x-ray source 106. Filter material 308 includes first filter material 310 and second filter material 312, wherein first filter material 310 comprises different material from second filter material 312. In a preferred embodiment, first filter material 310 includes aluminum (Al) and second filter material 312 includes copper (Cu).

[0060] In operation, x-ray source 106 emits imaging radiation at a first energy level. The x-rays travel through first filter material 310 which filters the imaging radiation such that low energy radiation exits first filter material 310. In response to information transferred from computer system 250, controller 304 commands motor 302 to turn shaft 306 which rotates filter material 308 such that second filter material 312 is positioned adjacent x-ray source 106. Then, x-ray source 106 emits imaging radiation at the same first energy level. The radiation travels through second filter material 312 which filters the imaging radiation such that high energy radiation exits second filter material 312. This process continues such that the energy levels of imaging radiation exiting filter material 308 alternate between low and high energies.

[0061] FIGS. 4A and 4B illustrate an alternative embodiment of an x-ray source filter for generating multi-energy x-ray images. X-ray source filter 400 includes motor 302, controller 304, shaft 306, and filter material 408. Motor 302, controller 304, and shaft 306 operate in the same manner as described with respect to FIGS. 3A and 3B. Filter material 408 is made of material suitable for filtering x-ray radiation, e.g., Al or Cu, and is configured to filter radiation emitted from x-ray source 106. Filter material 408 includes first filter thickness 410 and second filter thickness 412, wherein first filter thickness 410 is made of the same material

as second filter thickness 412, but has a different thickness. As such, imaging radiation emitted from x-ray source 106 has a first energy after traveling through first filter thickness 410 and has a second energy after traveling through second filter thickness 412.

[0062] Referring now to FIGS. 5A, 5B, and 5C, yet another alternative embodiment of an x-ray source filter for generating multi-energy x-ray images is described. FIG. 5A illustrates a side view of x-ray source filter 500 which includes motor 502, link 504, slot 506, and filter material 508. X-ray source filter 500 is coupled to an x-ray source, e.g., x-ray source 106 of FIG. 1 as illustrated or x-ray sources 108, 206, 208, or any additional x-ray sources provided.

[0063] Motor 502 is configured to slide link 504 along slot 506 in response to commands from controller 507. Controller 304 communicates with computer system 250 (see FIG. 2) which determines desired motor characteristics, such as power and speed, using suitable software. Computer system 250 then transmits the desired motor characteristics to controller 304 using a wired or wireless communication link.

[0064] Link 504 is coupled to motor 502 and filter material 508 such that filter material 508 translates linearly when motor 502 slides link 504 along slot 506. In one embodiment, filter material 508 is made of at least two materials suitable for filtering x-ray radiation and is configured to filter imaging radiation emitted from x-ray source 106. Filter material 508 includes first filter material 510 and second filter material 512, wherein first filter material 510 comprises different material from second filter material 512. In a preferred embodiment, first filter material 510 includes aluminum (Al) and second filter material 512 includes copper (Cu). In an alternative embodiment, first filter material 510 comprises the same material as second filter material 512, but has a different thickness.

[0065] In operation, x-ray source 106 emits imaging radiation at a first energy level when filter material 508 is in a first position as illustrated in FIGS. 5A and 5B. The imaging radiation travels through first filter material 510 which filters the imaging radiation such that low energy radiation exits first filter material 510. In response to information transferred from computer system 250, controller 507 commands motor 502 to slide link 504 along slot 506 which translates filter material 508 to a second position such that second filter material 512 is positioned adjacent x-ray source 106 as illustrated in FIG. 5C. Then, x-ray source 106 emits imaging radiation at the same first energy level. The radiation travels through second filter material 512 which filters the imaging radiation such that high energy radiation exits

second filter material 512. This process continues such that the energy levels of imaging radiation exiting filter material 508 alternate between low and high x-ray energies.

[0066] As a further alternative, an x-ray source filter may be disposed on a hinge, so that the filter may alternatingly be disposed or removed from the path of imaging radiation emitted by the x-ray source.

[0067] In FIGS. 3A-5C, although x-ray source filters 300, 400, and 500 illustratively are separate from x-ray source 106, the filters may be integrated into a common housing with an x-ray source without departing from the scope of the present invention.

[0068] Referring now to FIGS. 6A and 6B, an alternative embodiment is described in which a filter is selectively placed before the detector to filter imaging radiation that has already passed through the patient. In this embodiment, an x-ray detector filter for generating multi-energy x-ray images is described in which x-ray detector filter 600 includes first filter material 602 and second filter material 604. As seen in the exploded view of FIG. 6A and the assembled view of FIG. 6B, x-ray detector filter 600 may be disposed on an x-ray detector, e.g., x-ray detector 110 of FIG. 1 as illustrated or x-ray detector 112, 210, 212, or any additional x-ray detector provided. Alternatively, x-ray detector filter 600 may be integrated into a common housing with an x-ray detector.

[0069] X-ray detector filter 600 is made of material suitable for filtering x-ray radiation and is configured to filter imaging radiation, emitted from an x-ray source, that has passed through part or all of a patient. In one embodiment, first filter material 602 comprises different material from second filter material 604. In a preferred embodiment, first filter material 602 includes aluminum (Al) and second filter material 604 includes copper (Cu). In another preferred embodiment, first filter material 602 includes no material and second filter material 604 includes copper (Cu). In an alternative embodiment, first filter material 602 comprises the same material as second filter material 604, but has a different thickness.

[0070] In operation, an x-ray source, e.g., x-ray source 106 of FIG. 1, emits x-ray radiation 606 which travel through a patient to x-ray detector filter 600. X-ray radiation 606 passes through first filter material 602 or second filter material 604 of x-ray detector filter 600. First filter material 602 filters x-ray radiation 606 such that low energy x-ray radiation 608 exits first filter material 602 while second filter material 604 filters x-ray radiation 606 such that high energy x-ray radiation 610 exits second filter material 604. Low and high

energy x-ray radiations 608 and 610 are received by x-ray detector 110. X-ray detector 110 then transmits the received low and high energy x-ray radiation 608 and 610 to a suitable computer for processing the received radiation into first and second sets of image data and generating multi-energy x-ray images.

- [0071] X-ray detector filter 600 may take the form of a rectangular grid as illustrated, although many patterns are possible. For example, x-ray detector filter 600 may be patterned such that its columns alternate material or thickness. In a preferred embodiment, first and second filter materials 602 and 604 are each the size of a pixel on x-ray detector 110.
- [0072] FIG. 6B illustrates an assembled view of an exemplary pair of x-ray detector filters 600 and 600' according to one embodiment of the present invention. X-ray detector filter 600 is disposed on x-ray detector 110 and x-ray detector filter 600' is disposed on x-ray detector 112. X-ray detector filter 600' has a pattern complementary to the pattern of x-ray detector filter 600, for use in stereoscopic imaging. For example, the pattern of x-ray detector filter 600' may mirror the pattern of x-ray detector filter 600 as illustrated.
- [0073] In FIGS. 3A-6B, although filter materials 308, 408, 508, and x-ray detector filter 600 illustratively include two filter materials or two thicknesses, a greater number of materials and/or thicknesses, e.g., three, four, or more, may be employed to generate x-rays at a greater number of energies, e.g., three, four, or more, without departing from the scope of the present invention.

Systems and Methods for Processing and Using Multi-Energy X-Ray Images

- [0074] The present invention provides systems and methods for processing multi-energy x-ray images and using the same in image-guided radiation therapy (IGRT). Advantageously, x-rays generated at different energy levels may be processed to generate enhanced image data with positive/negative weight factors from the x-ray images. The enhanced x-ray images may be used for superior target tracking and positioning in IGRT.
- [0075] FIG. 7 is a schematic view of an exemplary IGRT system for use with multienergy x-ray images in accordance with the principles of the present invention. IGRT system 700 may include diagnostic imaging system 701, diagnostic image processor 702, treatment planning system 703, treatment planning library 703-A, system processor 704, memory 705, treatment and imaging system 706, and non-x-ray based position sensing system 710.

[0076] Diagnostic imaging system 701 is configured to generate treatment planning images of a region of interest within a patient. Diagnostic imaging system 701 may be a high precision volumetric imaging system such as a computed tomography (CT) system or a nuclear magnetic resonance imaging (MRI) system.

[0077] Images generated by diagnostic imaging system 701 may be processed to enhance image features by diagnostic image processor 702 using digital enhancement techniques known in the art. Diagnostic image processor 702 may process the images to render digitally reconstructed radiographs (DRRs) using techniques known in the art. The processed images from diagnostic image processor 702 may be stored in treatment planning library 703-A within treatment planning system 703. Treatment planning library 703-A may be any kind of digital storage medium such as, for example, magnetic or solid state media capable of storing digital x-ray images. Treatment planning system 703 may be configured to render 3-D diagnostic images and one or more treatment plans, which treatment plans may include the spatial relationship between a radiation treatment x-ray source and the region of interest during a prospective IGRT procedure. Treatment planning system 703 is coupled to system processor 704 which may be any type of general purpose or special purpose processing device capable of executing instructions and operating on image data and other data, and of commanding an IGRT system, such as treatment and imaging system 706. System processor 704 may include memory 705, which may be any type of memory capable of storing data and instructions for operating IGRT system 700. System processor 704 may be configured to process first and second sets of image data received from real-time multi-energy imaging system 709 to generate an enhanced image of part or all of the patient and to direct treatment delivery system 708 based on information obtained from the enhanced image, as described below.

[0078] Treatment and imaging system 706 includes controller 707 coupled to treatment delivery system 708 and real-time multi-energy imaging system 709. Controller 707 may be configured to coordinate the operations of treatment delivery system 708 and real-time multi-energy imaging system 709 in response to commands from system processor 704. Treatment delivery system 708 includes a radiation source configured to generate treatment radiation beams, e.g., x-ray, electron, or proton beams, based on the commands, e.g., a programmed routine, from system processor 704. In one embodiment, treatment delivery system 708 includes articulated robot arm 102, e.g., a six degree-of-freedom robot arm, MV radiation

source 104, and treatment table 114 of FIG. 1. In another embodiment, treatment delivery system includes a gantry configured to position the radiation source.

[0079] Treatment delivery system 708 may further include a collimator configured to collimate the treatment radiation beams. The collimator may be a fixed collimator, a variable aperture collimator having an aperture, or a multileaf collimator having leaves. In one embodiment, the variable aperture collimator approximates a circular field. System processor 704 may direct treatment delivery system 708 to set the aperture based on information obtained from the enhanced image, to change the diameter of the circular field, to move the leaves of the multileaf collimator, to move the variable aperture collimator, to move the multileaf collimator, and/or to position the patient.

[0080] System processor 704 may direct treatment delivery system 708 based on other information obtained from the enhanced image. Such information may include the position of the target within a patient, the position of a skeletal structure within the patient, the position of a soft tissue within the patient, and/or the position of fiducials within the patient and preferably within the target. System processor 704 may direct treatment delivery system 708 to position the radiation source, position the patient using, for example, treatment table 114, and/or enable or disable the treatment radiation beams based on the information.

[0081] Real-time multi-energy imaging system 709 is configured to generate two, three, four, or more sets of image data at two, three, four, or more energy levels for generating multi-energy x-ray images of a region of interest that includes the target, the skeletal structure, the soft tissue, and/or fiducials in real-time during an IGRT procedure. The sets of image data may be volumetric image data, such as tomosynthetic image data or CT image data, or a plurality of two-dimensional projection image data. Real-time multi-energy imaging system 709 may include apparatus described above with respect to FIGS. 2-6B for generating multi-energy x-ray images such as x-ray sources 106, 108, 206, 208, x-ray detectors 110, 112, 210, 212, x-ray source filters 300, 400, 500, and/or x-ray detector filter 600, or may include apparatus known to one of ordinary skill in the art for generating multi-energy x-ray images such as a sandwich-type x-ray detector. Real-time multi-energy x-ray images acquired from real-time multi-energy imaging system 709 may be processed by system processor 704 to enhance image features which may improve similarity measures between the pre-treatment and in-treatment images.

[0082] Real-time multi-energy imaging system 709 may include an x-ray source array having a plurality of emission spots. A first emission spot of the plurality of emission spots may be configured to emit imaging radiation at the first energy level, and optionally subsequently at the second energy level, and a second emission spot of the plurality of emission spots that may be configured to emit imaging radiation at the second energy level, and optionally subsequently at the first energy level. The plurality of emission spots may be configured to emit imaging radiation at the first energy level and to subsequently emit imaging radiation at the second energy level. The multi-energy imaging system may further include a second x-ray source array having a plurality of emission spots configured to emit imaging radiation in-phase or out of phase with the emission spots of the first x-ray source array. The plurality of emission spots of the first x-ray source array may be configured to emit imaging radiation at the first energy level, and optionally subsequently at the second energy level, and the plurality of emission spots of the second x-ray source array may be configured to emit imaging radiation at the second energy level, and optionally subsequently at the first energy level. The plurality of emission spots of the first x-ray source array and the plurality of emission spots of the second x-ray source array also may be configured to emit imaging radiation at the first energy level and to subsequently emit imaging radiation at the second energy level.

enhancement techniques known in the art. In a preferred embodiment, system processor 704 is configured to combine x-ray images generated at two or more energies to provide multi-energy x-ray images. For example, system processor 704 may combine x-rays generated within a low energy range (e.g., ~ 50-100 kV) with x-rays generated within a high x-ray energy range (e.g., ~ 100 kV – 6 MV, preferably 100-150 kV) to provide enhanced images of objects of interest, e.g., target, soft tissue, and radio-opaque objects such as skeletal structures, fiducials, and contrast agents. System processor 704 may process image data wherein the enhanced image comprises a weighted combination of the first and second sets of image data. The enhanced image may comprise a weighted combination in logarithmic space of the first and second sets of image data and/or a spatially varying weighted combination of the first and second sets of image data. The enhanced image may also be a weighted subtraction in logarithmic space of the first and second sets of image data. The enhanced image may also be a weighted subtraction in logarithmic space of the first and second sets of image data. The enhanced image may be an x-ray image displaying primarily soft tissue without radio-opaque objects

such as skeletal structures, fiducials, and contrast agents. Alternatively, the enhanced image may be an x-ray image displaying primarily radio-opaque objects without soft tissue.

[0084] FIGS. 8A and 8B are exemplary x-ray images that may be generated by real-time multi-energy imaging system 709 and FIGS. 8C and 8D are x-rays images that may be processed by system processor 704, available from http://en.wikibooks.org/wiki/Basic_Physics_of_Nuclear_Medicine/Dual-Energy_Absorptiometry. FIG. 8A is a chest x-ray generated at 56 kV, i.e., a low energy x-ray image. FIG. 8B is a chest x-ray generated at 120 kV, i.e., a high energy x-ray image, using a copper filter. FIG. 8C is a processed chest x-ray displaying soft tissue and having the skeletal structures subtracted out. FIG. 8D is a processed chest x-ray displaying skeletal structures and having the soft tissue subtracted out.

[0085] Referring back to FIG. 7, registration of the pre-treatment and in-treatment images may be performed by system processor 704 on image data sent to system processor 704 from treatment planning system 703 and real-time imaging system 709. The registration of the pre-treatment and in-treatment x-ray images may include calculation of in-plane translations, in-plane rotation and out-of-plane rotation, as is known in the art.

[0086] Non-x-ray based position sensing system 710 is configured to generate a respiratory motion model of a patient. Non-x-ray based position sensing system 710 senses position and/or movement of external marker(s) strategically affixed to the patient, and/or senses position and/or movement of the patient skin surface itself, using one or more methods that do not involve ionizing radiation, such as optically based or ultrasonically based methods. Non x-ray based position sensing system 710 may include external markers affixed in some manner to a patient's chest which move in response to respiration (other mechanisms for monitoring respiration may be used), and include a mono or stereoscopic xray imaging system which can precisely determine target location. System 710 correlates motion of the external markers with target motion, as determined from (for example) the mono or stereoscopic x-ray projections. Non x-ray based position sensing system 710 permits system processor 704 to monitor external marker motion, use the correlation model to precisely predict where the target will be located in real time (e.g., ~ 60 Hz), and direct the treatment beam to the target. As treatment of the moving target progresses, additional x-ray images may be obtained and used to verify and update the correlation model. In one

example, non-x-ray position sensing system 710 is similar to SYNCHRONY® Respiratory Tracking System available from Accuray Incorporated of Sunnyvale, California.

[0087] FIG. 9 illustrates a method for processing multi-energy x-ray image data in IGRT to enhance image quality in interventional images by exploiting beam hardening effects using High Dynamic Range (HDR) x-ray imaging. With reference to FIG. 7, method 900 may include determining an optimal energy level having the highest definition or differentiation ability for an object of interest, e.g., soft tissue, for imaging the object with real-time multi-energy imaging system 709 using a phantom with system processor 704, at step 910.

[0088] FIG. 10 is a plot illustrating a change in x-ray spectra after traversal through a filter for determining the optimal energy level. The plot shows count versus energy of x-rays that traversed aluminum plates having varied thickness. X-ray energy level 1000 is the determined optimal energy level for the selected object of interest.

[0089] Referring back to FIG. 9, at step 920, a first set of image data is generated using imaging radiation at the optimal energy level with real-time multi-energy imaging system 709. At step 930, the energy of the imaging radiation is adjusted with controller 707 and a second set of image data is generated using imaging radiation at the adjusted energy level with real-time multi-energy imaging system 709. Then, at step 940, the second set of image data generated at the adjusted energy level is processed with the first set of image data generated at the optimal energy level using system processor 704 to generate a processed HDR x-ray image. Advantageously, the processed HDR x-ray image is generated from a combination of x-rays generated over a wide energy spectrum.

[0090] FIG. 11 illustrates a method for registering multi-energy x-ray images in IGRT in accordance with one aspect of the present invention. With reference to FIG. 7, method 1100 may include, in a treatment planning phase, generating treatment planning images, e.g., a CT volume, with diagnostic imaging system 701 at step 1110. Then, at step 1120, the treatment planning images are pre-processed with diagnostic image processor 702 to generate digitally reconstructed radiographs (DRRs). In a treatment delivery phase, real-time multi-energy x-ray image data is generated with real-time multi-energy imaging system 709 at step 1130. For example, a first set of image data may be generated using imaging radiation at a first energy level and a second set of image data may be generated using imaging radiation at a second energy level. At step 1140, the real-time multi-energy x-ray image data is processed

with system processor 704 to extract real-time x-ray image features and to optionally scale the real-time multi-energy x-ray image data to correct for different imaging geometries between diagnostic imaging system 701 and real-time multi-energy imaging system 709. Processing may include combining image data from x-rays generated at different energy levels. At step 1150, the real-time multi-energy x-ray image data is registered with corresponding DRRs with system processor 704 to obtain a registration result. Optionally, at step 1160, the real-time multi-energy x-ray images are tracked against the DRRs with system processor 704 to determine onject of interest/patient movement and/or position. Beneficially, tracking the object of interest with multi-energy x-rays images can provide precise target location during IGRT for enhanced accuracy of treatment radiation emission from treatment delivery system 708 to the target.

[0091] FIG. 12 illustrates a method for tracking target(s), skeletal structure(s), soft tissue(s), contrast agent(s), fiducial(s), patient movement and/or position using multi-energy x-ray images in IGRT in accordance with another aspect of the present invention. With reference to FIG. 7, method 1200 may include, in a treatment delivery phase, generating realtime multi-energy x-ray image data with real-time multi-energy imaging system 709 at step 1210. For example, a first set of image data may be generated using imaging radiation at a first energy level and a second set of image data may be generated using imaging radiation at a second energy level. Then, at step 1220, the real-time multi-energy x-ray image data is processed with system processor 704 to combine image data from x-rays generated at different energy levels. In a preferred embodiment, system processor 704 processes the multi-energy x-ray image data to combine negative weight factor to subtract radio-opaque objects to generate primarily soft tissue x-ray images and to subtract soft tissues to generate primarily radio-opaque x-ray images. At step 1230, real-time soft-tissue x-ray images are registered with corresponding DRRs with system processor 704 to obtain a soft tissue registration result and real-time radio-opaque object x-ray images are registered with corresponding DRRs with system processor 704 to obtain a radio-opaque registration result. The soft tissue images and/or the radio-opaque object images may be tracked against the DRRs with system processor 704 to determine target/skeletal structure/soft tissue/fiducial/contrast agent/patient movement and/or position at steps 1240 and 1245.

[0092] Advantageously, tracking the target, e.g., a tumor, with soft tissue x-ray images may provide images in which the target is not obscured by overlapping bones, thereby

permitting precise determination of the target location during IGRT. In the case where a target is partially or completely blocked by skeletal structures using conventional x-ray images, the soft tissue x-ray images of the present invention may provide a clear target image, allowing for superior target localization and tracking. Moreover, skeletal x-ray images may be used for enhanced patient positioning, e.g., for initial spine alignment, and to track skeletal motion and detect patient shift. Because the techniques of the present invention provide enhanced target/patient movement and/or positioning, the accuracy of treatment radiation emission from treatment delivery system 708 to the target is expected to be superior to previously-known techniques.

[0093] FIG. 13 illustrates a method for target tracking using a motion-edge artifact in IGRT in accordance with yet another aspect of the present invention. With reference to FIG. 7, method 1300 may include, in a treatment delivery phase, generating a first set of image data with real-time multi-energy imaging system 709 at step 1310.

[0094] FIG. 14A is an illustration of an x-ray image of target 1400 at first position 1401 generated using the first set of image data.

[0095] Referring back to FIG. 13, at step 1320, a second set of image data is generated with multi-energy imaging system 709. In a preferred embodiment, the first set of image data is generated using radiation at a first energy level and the second set of image data is generated using radiation at a second energy level.

[0096] FIG. 14B is an illustration of an x-ray image of target 1400 at second position 1402 generated using second x-ray image data. Illustratively, target 1400 has moved along the Axis of Motion from first position 1401 to second position 1402. Additionally, the Axis of Motion of the target may be estimated from a 4-D CT scan or a 3-D CT inhale/exhale pair during treatment planning with treatment planning system 703.

[0097] Again, referring back to FIG. 13, at step 1330, the first and second sets of image data are processed with system processor 704 to perform weighted combination of an overlapping portion of the first and second sets of image data to generate a motion-edge artifact image. Motion artifacts are effects in imaging caused by movement of the target being imaged and are generally seen in images as blurring and/or streaking. The size of the motion artifact may be predicted from a 4-D CT scan because size mostly depends on the dimensions of the target and target velocity in a particular respiratory phase.

[0098] FIG. 14C is an illustration of the motion-edge artifact image generated after combining overlapping portions of the target at first position 1401 and second position 1402. The non-overlapping portions of the target at first and second positions 1401 and 1402 are motion-edge artifacts. Trailing edge artifact 1403 is the area in the x-ray image that was imaged at first position 1401, but not imaged at second position 1402. Leading edge artifact 1404 is the area in the x-ray image that was imaged at second position 1402, but not imaged at first position 1401.

[0099] Referring yet again back to FIG. 13, at step 1340, the movement and/or position of the target is tracked with system processor 704 using the motion-edge artifact image which may be inputted into suitable software. In a preferred embodiment, method 1300 is used to track the target, e.g., a tumor. In an alternative embodiment, method 1300 may be used to track non-target structures such as a rib or diaphragm. Advantageously, the methods of the present invention provide enhanced target localization using traditionally unwanted image artifacts as an additional constraint for target localization in IGRT.

[00100] FIG. 15 illustrates a method for target/soft tissue/skeletal structure/contrast agent/fiducial tracking during respiration using multi-energy x-ray images in IGRT in accordance with the present invention. With reference to FIG. 7, method 1500 may include, in a treatment delivery phase, generating a first set of image data at a position in the patient's respiratory cycle using imaging radiation at a first energy level with real-time multi-energy imaging system 709 at step 1510. At step 1520, a second set of image data is generated at the same position in the patient's subsequent respiratory cycle using imaging radiation at a second energy level with multi-energy imaging system 709. The first and second sets of image data are processed with system processor 704 to predict the target location at different positions in the respiratory cycle at step 1530. System processor 704 may communicate with non-x-ray based position sensing system 710 to enhance the target location prediction based on a respiratory motion model generated by non-x-ray based position sensing system 710. At step 1540, the movement and/or position of the object of interest and/or patient are tracked with system processor 704 using the first and second sets of image data and compared to the predicted target location.

[00101] In a preferred embodiment, the first energy level is a high energy level such that the first set of image data includes data having greater soft tissue attenuation and the second energy level is a low energy level such that the second set of image data includes data having

greater skeletal structure attenuation. Because soft tissue displacements may vary significantly between respiration cycles due to factors such as tissue hysteresis and variations in alveolar recruitment, skeletal structures may correlate well with the external markers of non-x-ray based position sensing system 710. As such, the first and second sets of image data may be processed to combine the data so that the soft tissue image data is the most recently acquired data and may be used for target tracking and to update the model from non-x-ray based position sensing system 710.

[00102] It will be apparent from the foregoing description to one skilled in the art that aspects of the present invention may be embodied, at least in part, in software. That is, referring to FIG. 7, the techniques may be carried out in a computer system or other data processing system in response to its processor, such as system processor 704, executing sequences of instructions contained in a memory, such as memory 705. In various embodiments, hardware circuitry may be used in combination with software instructions to implement the present invention. Thus, the techniques are not limited to any specific combination of hardware circuitry and software or to any particular source for the instructions executed by the data processing system. In addition, throughout this description, various functions and operations may be described as being performed by or caused by software code to simplify description. However, those skilled in the art will recognize what is meant by such expressions is that the functions result from execution of the code by a processor or controller, such as system processor 704 or controller 707.

[00103] A machine-readable medium can be used to store software and data which, when executed by a data processing system, causes the system to perform various methods of the present invention. This executable software and data may be stored in various places including, for example, memory 705 and treatment planning library 703-A or any other device that is capable of storing software programs and/or data.

[00104] A machine-readable medium includes any mechanism that provides (i.e., stores and/or transmits) information in a form accessible by a machine (e.g., a computer, network device, personal digital assistant, manufacturing tool, any device with a set of one or more processors, etc.). For example, a machine-readable medium includes recordable/non-recordable media (e.g., read only memory (ROM); random access memory (RAM); magnetic disk storage media; optical storage media; flash memory devices; etc.), as well as electrical,

optical, acoustical or other forms of propagated signals (e.g., carrier waves, infrared signals, digital signals, etc.).

[00105] While various illustrative embodiments of the invention are described above, it will be apparent to one skilled in the art that various changes and modifications may be made therein without departing from the invention. The appended claims are intended to cover all such changes and modifications that fall within the true scope of the invention.

WHAT IS CLAIMED:

1. An image-guided radiation therapy system for treating a target within a patient, the system comprising:

a treatment delivery system having a radiation source configured to generate treatment radiation beams and a collimator configured to collimate the treatment radiation beams;

a multi-energy imaging system configured to generate a first set of image data of part or all of the patient using imaging radiation at a first energy level and a second set of image data of part or all of the patient using imaging radiation at a second energy level; and

a system processor configured to process the first and second sets of image data to generate an enhanced image of part or all of the patient, the system processor further configured to direct the treatment delivery system based on information obtained from the enhanced image.

2. The image-guided radiation therapy system of claim 1, wherein part or all of the image data comprises the target; and

wherein the information obtained from the enhanced image comprises a position of the target.

3. The image-guided radiation therapy system of claim 1, wherein part or all of the image data comprises a skeletal structure within the patient; and

wherein the information obtained from the enhanced image comprises a position of the skeletal structure.

4. The image-guided radiation therapy system of claim 1, wherein part or all of the image data comprises a soft tissue structure within the patient; and

wherein the information obtained from the enhanced image comprises a position of the soft tissue.

5. The image-guided radiation therapy system of claim 1, wherein part or all of the image data comprises the target having fiducials implanted therein; and

wherein the information obtained from the enhanced image comprises a position of the fiducials.

6. The image-guided radiation therapy system of claim 1, wherein part or all of the image data comprises a contrast agent within the patient; and

wherein the information obtained from the enhanced image comprises a position of the contrast agent.

- 7. The image-guided radiation therapy system of claim 1, wherein the system processor is further configured to direct the treatment delivery system to position the radiation source based on information obtained from the enhanced image.
- 8. The image-guided radiation therapy system of claim 1, wherein the treatment delivery system comprises a robot arm configured to position the radiation source.
- 9. The image-guided radiation therapy system of claim 1, wherein the robot arm comprises a six degree-of-freedom robot arm.
- 10. The image-guided radiation therapy system of claim 1, wherein the treatment delivery system comprises a gantry configured to position the radiation source.
- 11. The image-guided radiation therapy system of claim 1, wherein the system processor is further configured to direct the treatment delivery system to position the patient based on information obtained from the enhanced image.
- 12. The image-guided radiation therapy system of claim 1, wherein the system processor is further configured to direct the treatment delivery system to enable or disable the treatment radiation beams based on information obtained from the enhanced image.
- 13. The image-guided radiation therapy system of claim 1, wherein the enhanced image comprises a weighted combination of the first and second sets of image data.

14. The image-guided radiation therapy system of claim 13, wherein the enhanced image comprises a weighted combination in logarithmic space of the first and second sets of image data.

- 15. The image-guided radiation therapy system of claim 13, wherein the enhanced image comprises a spatially-varying weighted combination of the first and second sets of image data.
- 16. The image-guided radiation therapy system of claim 13, wherein the enhanced image comprises a spatially-varying weighted combination in logarithmic space of the first and the second sets of image data.
- 17. The image-guided radiation therapy system of claim 1, wherein the enhanced image comprises a weighted subtraction in logarithmic space of the first and second sets of image data.
- 18. The image-guided radiation therapy system of claim 1, wherein the enhanced image comprises an x-ray image displaying primarily a radio-opaque object without soft tissue.
- 19. The image-guided radiation therapy system of claim 1, wherein the enhanced image comprises an x-ray image displaying primarily soft tissue without a radio-opaque object.
- 20. The image-guided radiation therapy system of claim 18, wherein the radioopaque object comprises a skeletal structure, a fiducial, a contrast agent, or any combination thereof.
- 21. The image-guided radiation therapy system of claim 1, wherein part or all of the image data comprises the target; and

wherein the system processor is configured to track movement and position of the target using the enhanced image.

22. The image-guided radiation therapy system of claim 1, wherein the collimator comprises a variable aperture collimator having an aperture; and

wherein the system processor is further configured to direct the treatment delivery system to set the aperture based on information obtained from the enhanced image.

- 23. The image-guided radiation therapy system of claim 22, wherein the variable aperture collimator approximates a circular field.
- 24. The image-guided radiation therapy system of claim 23, wherein the system processor is further configured to direct the treatment delivery system to change the diameter of the circular field, move the variable aperture collimator, position the patient, or any combination thereof.
- 25. The image-guided radiation therapy system of claim 22, wherein the variable aperture collimator comprises a multileaf collimator.
- 26. The image-guided radiation therapy system of claim 25, wherein the system processor is further configured to direct the treatment delivery system to move leaves of the multileaf collimator, move the multileaf collimator, position the patient, or any combination thereof.
- 27. The image-guided radiation therapy system of claim 1, wherein the treatment radiation beams comprise a photon beam.
- 28. The image-guided radiation therapy system of claim 1, wherein the treatment radiation beams comprise an electron beam.
- 29. The image-guided radiation therapy system of claim 1, wherein the treatment radiation beams comprise a proton beam.
- 30. The image-guided radiation therapy system of claim 1, wherein the multienergy imaging system further comprises a first x-ray source configured to emit imaging radiation towards part or all of the patient and a first x-ray detector configured to receive the

imaging radiation after at least a portion of the imaging radiation has passed through part or all of the patient.

- 31. The image-guided radiation therapy system of claim 30, further comprising a second x-ray source configured to emit imaging radiation at part or all of the patient and a second x-ray detector configured to receive the imaging radiation after at least a portion of the imaging radiation has passed through part or all of the patient.
- 32. The image-guided radiation therapy system of claim 31, wherein the first x-ray source emits the imaging radiation at the first energy level and the second x-ray source emits the imaging radiation at the second energy level.
- 33. The image-guided radiation therapy system of claim 31, wherein the first x-ray source emits the imaging radiation at the first energy level and the second x-ray source substantially simultaneously emits the imaging radiation at the second energy level; and

subsequently, the first x-ray source emits the imaging radiation at the second energy level and the second x-ray source substantially simultaneously emits the imaging radiation at the first energy level.

34. The image-guided radiation therapy system of claim 31, wherein the first x-ray source emits the imaging radiation at the first energy level and the second x-ray source substantially simultaneously emits the imaging radiation at the first energy level; and

subsequently, the first x-ray source emits the imaging radiation at the second energy level and the second x-ray source substantially simultaneously emits the imaging radiation at the second energy level.

35. The image-guided radiation therapy system of claim 30, wherein the multienergy imaging system further comprises an x-ray source filter having first and second portions configured to filter the imaging radiation emitted from the first x-ray source, the xray source filter configured to move between a first position and a second position,

wherein, at the first position, the first portion is adjacent to the first x-ray source such that imaging radiation emitted from the first x-ray source travels through the first portion and exits the first portion at the first energy level, and

wherein, at the second position, the second portion is adjacent to the first x-ray source such that imaging radiation emitted from the first x-ray source travels through the second portion and exits the second portion at the second energy level.

36. The image-guided radiation therapy system of claim 30, wherein the multienergy imaging system further comprises an x-ray detector filter disposed between the first xray source and the first x-ray detector, the x-ray detector filter having a first portion configured to filter imaging radiation and a second portion,

wherein imaging radiation emitted from the first x-ray source travels through the first portion and exits the first portion at the first energy level, and

wherein imaging radiation emitted from the first x-ray source travels through the second portion and exits the second portion at the second energy level.

- 37. The image-guided radiation therapy system of claim 30, wherein the first x-ray source comprises a broad energy x-ray source and the first x-ray detector comprises an energy-binning detector.
- 38. The image-guided radiation therapy system of claim 30, wherein the first x-ray source comprises a first x-ray source array having a plurality of emission spots.
- 39. The image-guided radiation therapy system of claim 38, wherein a first emission spot of the plurality of emission spots is configured to emit imaging radiation at the first energy level and a second emission spot of the plurality of emission spots is configured to emit imaging radiation at the second energy level.
- 40. The image-guided radiation therapy system of claim 39, wherein the first emission spot is configured to subsequently emit imaging radiation at the second energy level and the second emission spot is configured to subsequently emit imaging radiation at the first energy level.
- 41. The image-guided radiation therapy system of claim 38, wherein the plurality of emission spots are configured to emit imaging radiation at the first energy level and to subsequently emit imaging radiation at the second energy level.

42. The image-guided radiation therapy system of claim 38, further comprising a second x-ray source array having a plurality of emission spots and a second x-ray detector,

wherein the second x-ray source array is configured to emit imaging radiation at part or all of the patient, and

wherein the second x-ray detector is configured to receive the imaging radiation after at least a portion of the imaging radiation has passed through part or all of the patient.

- 43. The image-guided radiation therapy system of claim 42, wherein the plurality of emission spots of the first x-ray source array are configured to emit imaging radiation at the first energy level and the plurality of emission spots of the second x-ray source array are configured to emit imaging radiation at the second energy level.
- 44. The image-guided radiation therapy system of claim 43, wherein the plurality of emission spots of the first x-ray source array are configured to subsequently emit imaging radiation at the second energy level and the plurality of emission spots of the second x-ray source array are configured to subsequently emit imaging radiation at the first energy level.
- 45. The image-guided radiation therapy system of claim 42, wherein the plurality of emission spots of the first x-ray source array and the plurality of emission spots of the second x-ray source array are configured to emit imaging radiation at the first energy level and to subsequently emit imaging radiation at the second energy level.
- 46. The image-guided radiation therapy system of claim 1, wherein the system processor is configured to determine an optimal energy level for imaging part or all of the patient, and

wherein the first energy level is the determined optimal energy level and the second energy level is an adjusted energy level different from the optimal energy level.

- 47. The image-guided radiation therapy system of claim 46, wherein the enhanced image comprises a high dynamic range x-ray image.
- 48. The image-guided radiation therapy system of claim 1, further comprising a diagnostic image processor configured to generate digitally reconstructed radiographs from previously acquired treatment planning images of the patient.

49. The image-guided radiation therapy system of claim 1, wherein the system processor is configured to register the first and second sets of image data with a corresponding digitally reconstructed radiograph generated from previously acquired treatment planning images to obtain a registration result.

50. The image-guided radiation therapy system of claim 49, wherein part or all of the image data comprises the target; and

wherein the system processor is configured to track movement and position of the target using the first and second sets of image data against the corresponding digitally reconstructed radiographs.

- 51. The image-guided radiation therapy system of claim 1, wherein the system processor is configured to register the enhanced image with a corresponding digitally reconstructed radiograph generated from previously acquired treatment planning images to obtain a registration result.
- 52. The image-guided radiation therapy system of claim 51, wherein part or all of the image data comprises the target; and

wherein the system processor is configured to track movement and position of the target using the enhanced image against the corresponding digitally reconstructed radiographs.

53. The image-guided radiation therapy system of claim 1, wherein the enhanced image comprises a motion-edge artifact image, and

wherein the system processor is configured to subtract an overlapping portion of the first set of image data from the second set of image data to generate the motion-edge artifact image.

54. The image-guided radiation therapy system of claim 53, wherein part or all of the image data comprises the target; and

wherein the system processor is configured to track movement and position of the target using the motion-edge artifact image.

55. The image-guided radiation therapy system of claim 1, wherein the multienergy imaging system is configured to generate the first set of image data at a position in the patient's respiratory cycle and to generate the second set of image data at the same position in a subsequent respiratory cycle.

- 56. The image-guided radiation therapy system of claim 1, further comprising a non-x-ray based position sensing system configured to generate a respiratory motion model to predict target location.
- 57. The image-guided radiation therapy system of claim 1, wherein the first energy level is an energy level selected within the range of 50-100 kV and the second energy level is an energy level selected within the range of 100-150 kV.
- 58. The image-guided radiation therapy system of claim 1, wherein the first set of image data comprises first volumetric image data and the second set of image data comprises second volumetric image data.
- 59. The image-guided radiation therapy system of claim 58, wherein the first set of image data comprises first tomosynthetic image data and the second set of image data comprises second tomosynthetic image data.
- 60. The image-guided radiation therapy system of claim 59, wherein the enhanced image comprises a tomosynthetic image.
- 61. The image-guided radiation therapy system of claim 58, wherein the first set of image data comprises first computed tomography (CT) image data and the second set of image data comprises second CT image data.
- 62. The image-guided radiation therapy system of claim 61, wherein the enhanced image comprises a CT image.
- 63. The image-guided radiation therapy system of claim 1, wherein the first set of image data comprises a first plurality of two-dimensional projection image data and the

second set of image data comprises a second plurality of two-dimensional projection image data,

wherein the processor is further configured to generate a plurality of intermediate enhanced images from the first and second pluralities of two-dimensional projection image data,

and wherein the processor is configured to process the plurality of intermediate enhanced images to generate the enhanced image.

- 64. The image-guided radiation therapy system of claim 63, wherein the enhanced image comprises a tomosynthetic image.
- 65. An image-guided radiation therapy system for treating a target within a patient, the system comprising:

a treatment delivery system having a radiation source configured to generate treatment radiation beams and a collimator configured to collimate the treatment radiation beams;

a multi-energy imaging system configured to generate a first set of tomosynthetic image data of part or all of the patient using imaging radiation at a first energy level and a second set of tomosynthetic image data of part or all of the patient using imaging radiation at a second energy level; and

a system processor configured to process the first and second sets of tomosynthetic image data to generate an enhanced tomosynthesis image of part or all of the patient, the system processor further configured to direct the treatment delivery system based on information obtained from the enhanced tomosynthesis image.

66. A method for treating a target within a patient for image-guided radiation, the method comprising:

generating a first set of image data of part or all of the patient using imaging radiation at a first energy level and a second set of image data of part or all of the patient using imaging radiation at a second energy level;

processing the first and second sets of image data to generate an enhanced image; and directing a treatment delivery system based on information obtained from the enhanced image.

67. The method of claim 66, wherein the treatment delivery system comprises a radiation source configured to generate treatment radiation beams and a collimator configured to collimate the treatment radiation beams.

68. The method of claim 66, wherein part or all of the image data comprises the target; and

wherein the information obtained from the enhanced image comprises a position of the target.

69. The method of claim 66, wherein part or all of the image data comprises a skeletal structure within the patient; and

wherein the information obtained from the enhanced image comprises a position of the skeletal structure.

70. The method of claim 66, wherein part or all of the image data comprises a soft tissue structure within the patient; and

wherein the information obtained from the enhanced image comprises a position of the soft tissue.

71. The method of claim 66, wherein part or all of the image data comprises the target having fiducials implanted therein; and

wherein the information obtained from the enhanced image comprises a position of the fiducials.

72. The method of claim 66, wherein part or all of the image data comprises a contrast agent within the patient; and

wherein the information obtained from the enhanced image comprises a position of the contrast agent.

- 73. The method of claim 66, wherein directing the treatment delivery system comprises positioning a radiation source of the treatment delivery system.
- 74. The method of claim 66, wherein the treatment delivery system comprises a robot arm configured to position a radiation source of the treatment delivery system.

75. The method of claim 66, wherein the robot arm comprises a six degree-of-freedom robot arm.

- 76. The method of claim 66, wherein the treatment delivery system comprises a gantry configured to position a radiation source of the treatment delivery system.
- 77. The method of claim 66, wherein directing the treatment delivery system comprises positioning the patient.
- 78. The method of claim 67, wherein directing the treatment delivery system comprises enabling or disabling the treatment radiation beams.
- 79. The method of claim 66, wherein the enhanced image comprises a weighted combination of the first and second sets of image data.
- 80. The method of claim 79, wherein the enhanced image comprises a weighted combination in logarithmic space of the first and second sets of image data.
- 81. The method of claim 79, wherein the enhanced image comprises a spatially-varying weighted combination of the first and second sets of image data.
- 82. The method of claim 66, wherein the enhanced image comprises a weighted subtraction of the first and second sets of image data.
- 83. The method of claim 66, wherein the enhanced image comprises a weighted subtraction in logarithmic space of the first and second sets of image data.
- 84. The method of claim 66, wherein the enhanced image comprises an x-ray image displaying primarily a radio-opaque object without soft tissue.
- 85. The method of claim 66, wherein the enhanced image comprises an x-ray image displaying primarily soft tissue without a radio-opaque object.

86. The method of claim 84, wherein the radio-opaque object comprises a skeletal structure, a fiducial, a contrast agent, or any combination thereof.

- 87. The method of claim 66, wherein part or all of the image data comprises the target; the method further comprising tracking movement and position of the target using the enhanced image.
- 88. A machine-readable medium comprising instructions that, when executed by a processor of an image-guided radiation therapy system, cause the processor to perform steps comprising:

generating a first set of image data of part or all of the patient using imaging radiation at a first energy level and a second set of image data of part or all of the patient using imaging radiation at a second energy level;

processing the first and second sets of image data to generate an enhanced image; and directing a treatment delivery system based on information obtained from the enhanced image.

89. An image-guided radiation therapy system for treating a target within a patient, the image-guided radiation therapy system comprising a treatment delivery system, the improvement comprising:

a programmed routine configured to process a first set of image data of part or all of the patient generated using imaging radiation at a first energy level and a second set of image data of part or all of the patient generated using imaging radiation at a second energy level to generate an enhanced image, the programmed routine further configured to direct the treatment delivery system based on information obtained from the enhanced image.

90. A multi-energy imaging system comprising:

an x-ray source array configured to generate a first set of image data using imaging radiation at a first energy level and a second set of image data using imaging radiation at a second energy level; and

a system processor configured to process the first and second sets of image data to generate an enhanced image.

91. The multi-energy imaging system of claim 90, wherein the first set of image data comprises first volumetric image data and the second set of image data comprises second volumetric image data.

- 92. The multi-energy imaging system of claim 91, wherein the first set of image data comprises first tomosynthetic image data and the second set of image data comprises second tomosynthetic image data.
- 93. The multi-energy imaging system of claim 92, wherein the enhanced image comprises a tomosynthetic image.
- 94. The multi-energy imaging system of claim 90, wherein the first set of image data comprises a first plurality of two-dimensional projection image data and the second set of image data comprises a second plurality of two-dimensional projection image data,

wherein the processor is further configured to generate a plurality of intermediate enhanced images from the first and second pluralities of two-dimensional projection image data,

and wherein the processor is configured to process the plurality of intermediate enhanced images to generate the enhanced image.

95. The multi-energy imaging system of claim 90, further comprising an additional x-ray source array positioned at a non-zero angle from the x-ray source array.

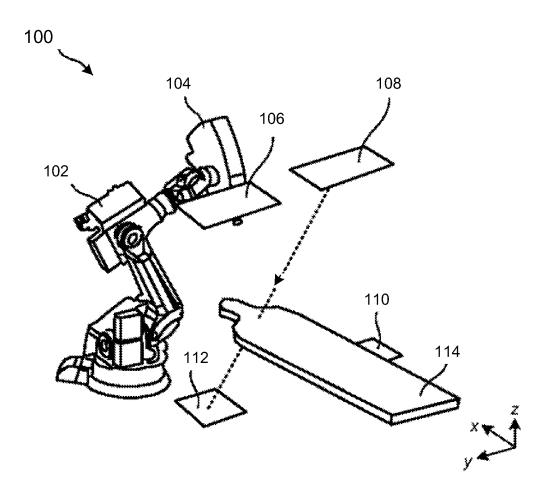
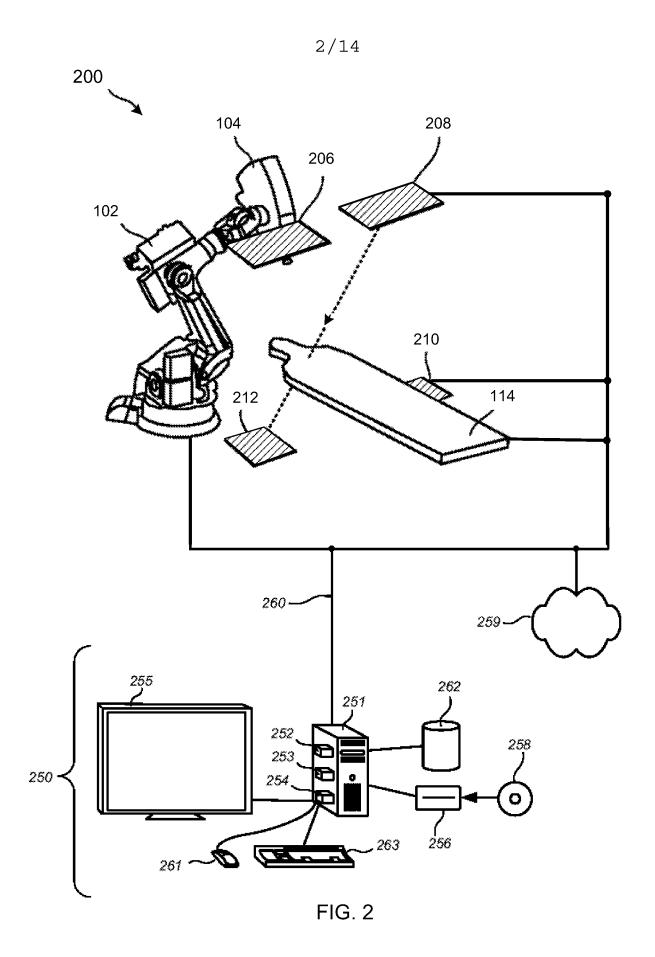


FIG. 1



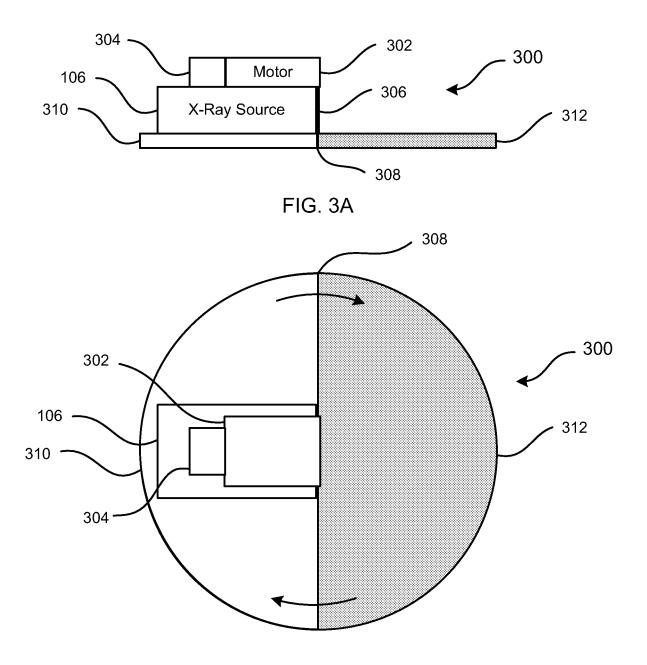


FIG. 3B

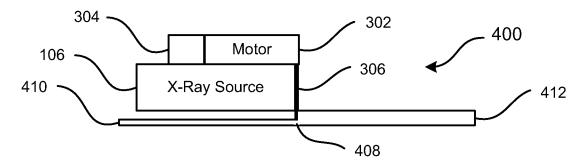


FIG. 4A

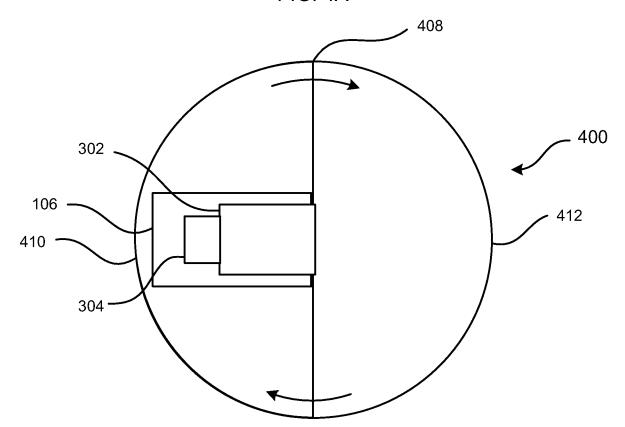
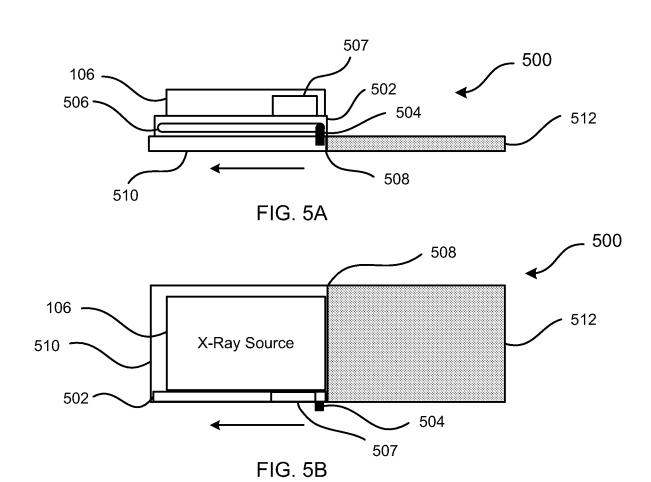
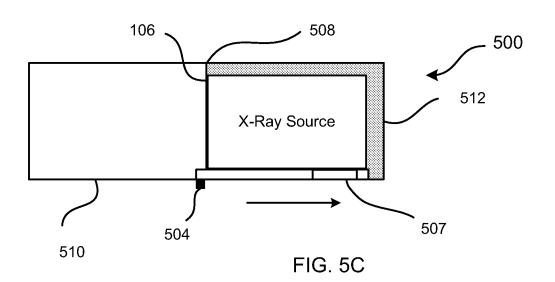


FIG. 4B







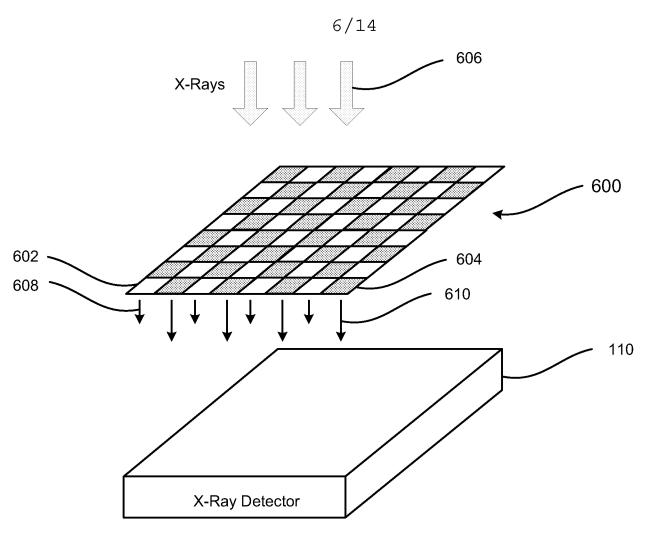


FIG. 6A

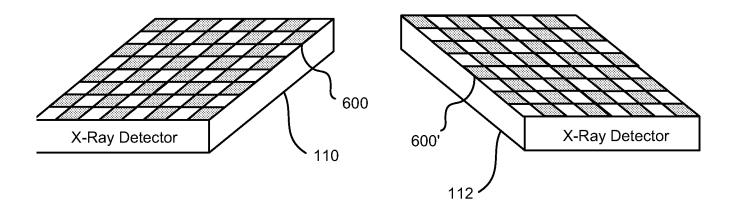


FIG. 6B

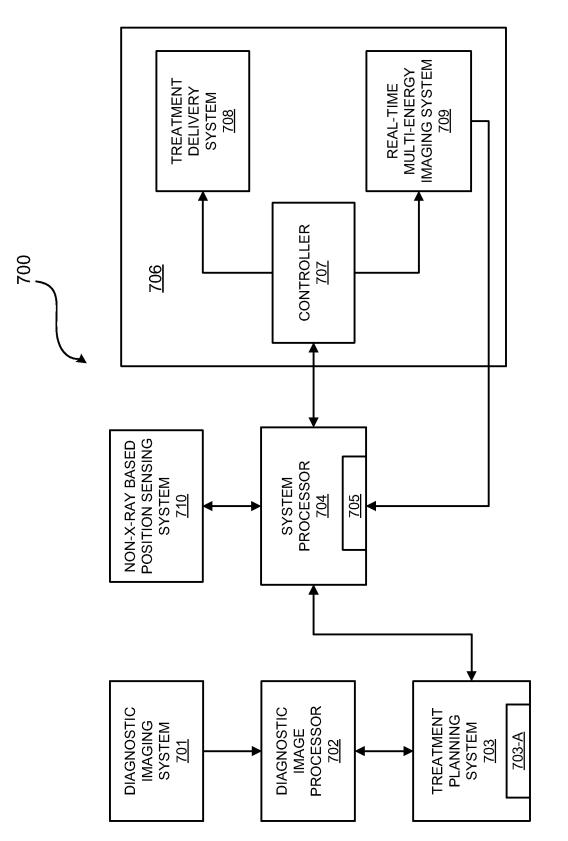


FIG. 7

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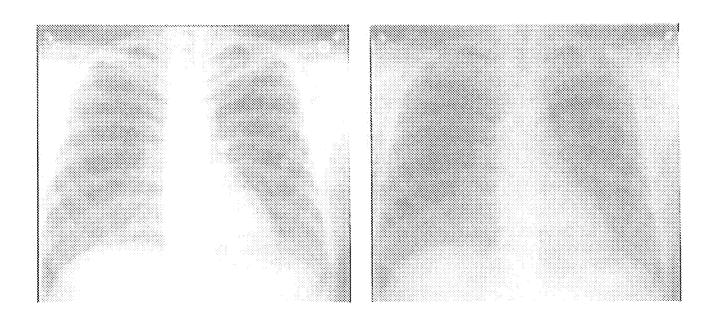


FIG. 8A FIG. 8B

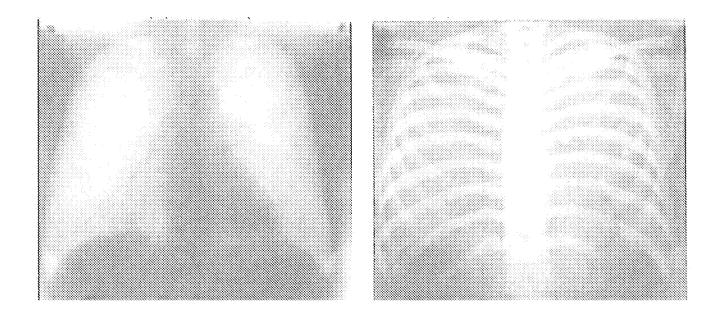


FIG. 8C FIG. 8D

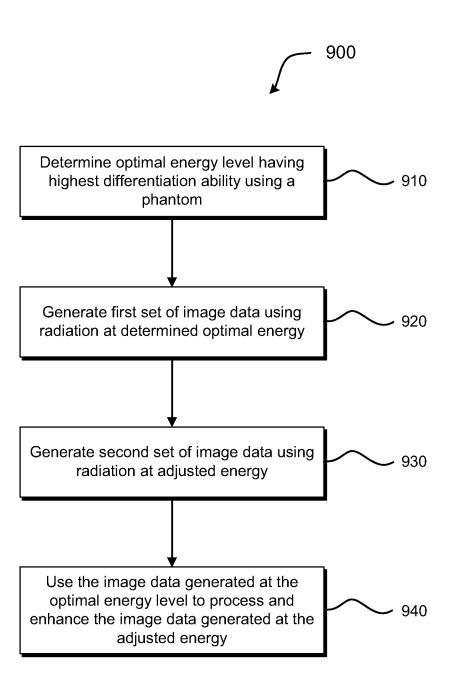


FIG. 9

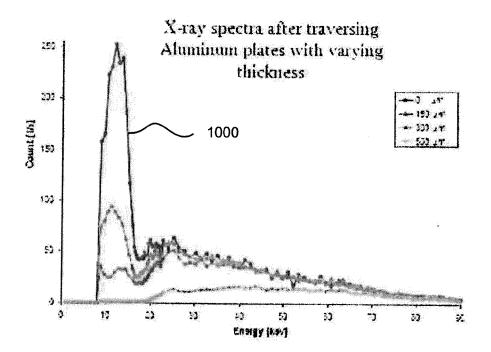


FIG. 10

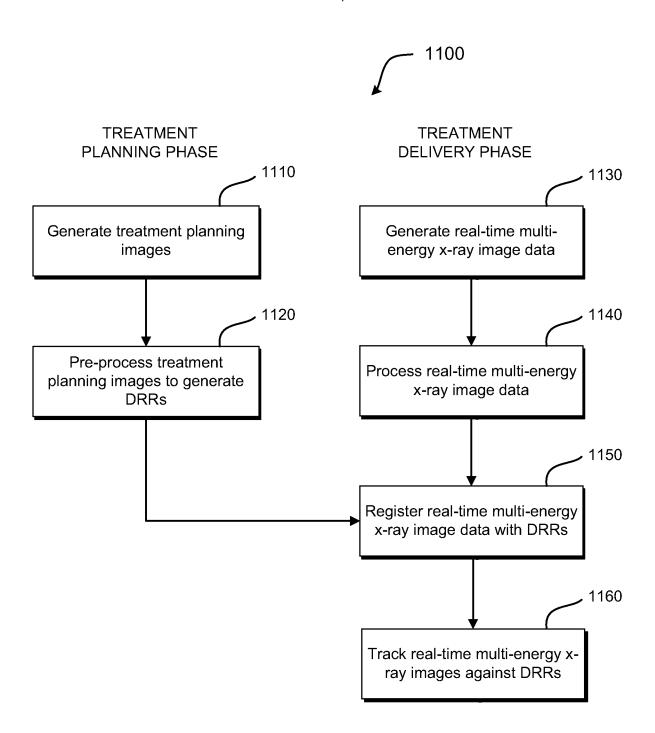


FIG. 11

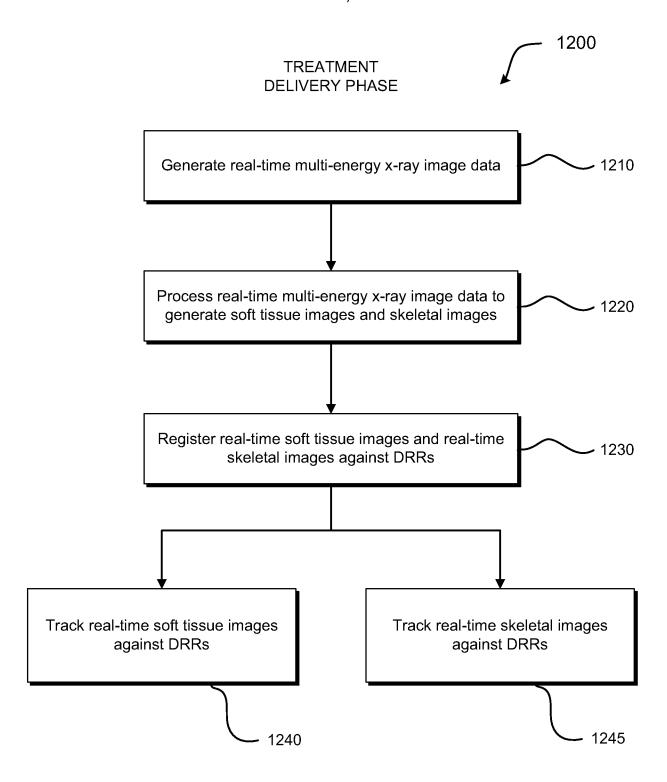


FIG. 12

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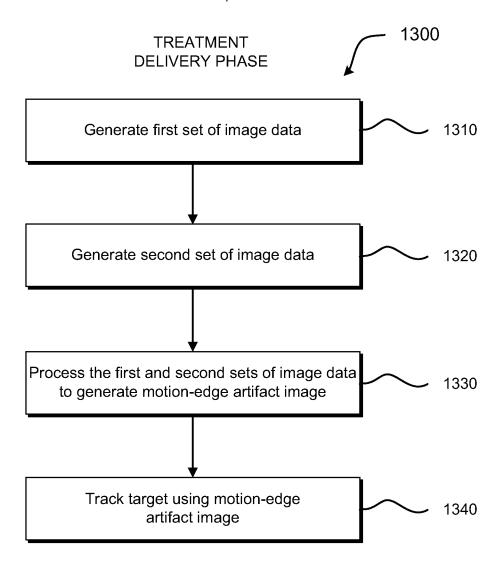
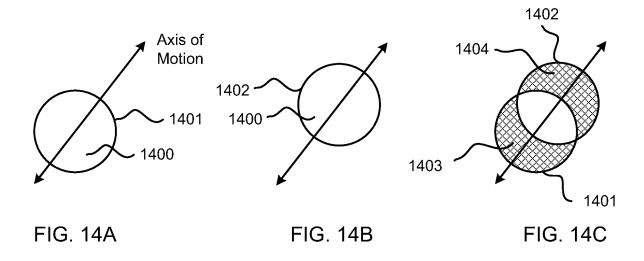


FIG. 13



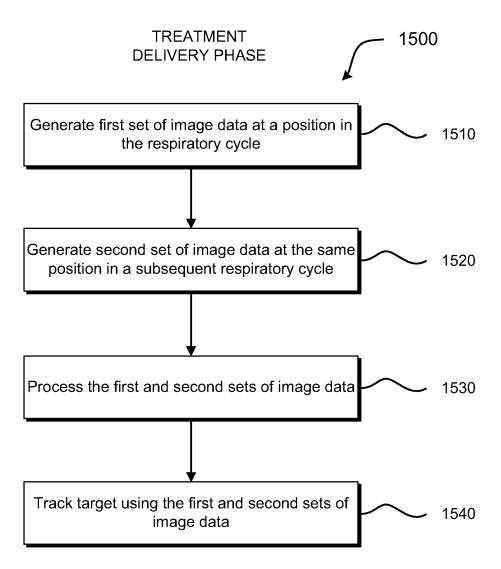


FIG. 15

International application No PCT/US2012/044513

A. CLASSIFICATION OF SUBJECT MATTER INV. A61N5/10 A61B6/00

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

A61N A61B

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

EPO-Internal, WPI Data

Category*	Citation of document, with indication, where appropriate, of the r	elevant passages	Relevant to claim No.
X	US 2009/274269 A1 (FOLAND ANDRE	w DEAN [US]	1,30,37,
	ET AL) 5 November 2009 (2009-11		88-90
Υ	abstract; figures 1-3	,	2-20,
	paragraphs [0004] - [0012], [0	020] -	27-29,
	[0038], [0043] - [0057], [006	4]	57,
			66-86,
			91,94,95
Χ	US 2009/283682 A1 (STAR-LACK JO	SH [US] ET	1,30-34,
	AL) 19 November 2009 (2009-11-1	9)	37,46,
.,			47,88-90
Y	abstract; figures 1, 3A-B, 10,	11	2-20,
	paragraphs [0006] - [0009], [0 [0034], [0044] - [0046]	025] -	27-29, 35,36,
	[0054], [0044] - [0040]		57,
			66-86,
			91,94,95
		,	
		-/	
	her documents are listed in the continuation of Box C.	X See patent family annex.	
* Special o	pategories of oited documents :	"T" later document published after the interr	
	ent defining the general state of the art which is not considered of particular relevance	date and not in conflict with the applica the principle or theory underlying the in	
"E" earlier : filing c	application or patent but published on or after the international	"X" document of particular relevance; the cla	
"L" docume	ent which may throw doubts on priority claim(s) or which is	considered novel or cannot be conside step when the document is taken alone	
	o establish the publication date of another citation or other al reason (as specified)	"Y" document of particular relevance; the cla considered to involve an inventive step	
"O" docum mean:	ent referring to an oral disclosure, use, exhibition or other s	combined with one or more other such being obvious to a person skilled in the	documents, such combination
	ent published prior to the international filing date but later than iority date claimed	"&" document member of the same patent for	amily
Date of the	actual completion of the international search	Date of mailing of the international seam	ch report
1	4 February 2013	22/02/2013	
Name and I	mailing address of the ISA/	Authorized officer	
	European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040,		

International application No
PCT/US2012/044513

(Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT				
ategory*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.		
(US 6 683 934 B1 (ZHAO JIANGUO [US] ET AL) 27 January 2004 (2004-01-27)	2-6, 13-20, 35,36, 57, 68-72, 80-86		
	abstract; figures 1-7B column 1, line 23 - column 2, line 6 column 2, line 22 - column 6, line 64			
'	US 2009/003528 A1 (RAMRAJ SANKARALINGAM [US] ET AL) 1 January 2009 (2009-01-01)	2-20, 27-29, 57, 66-86, 91,94,95		
	abstract; figures 1-6 paragraphs [0015] - [0034], [0036] - [0051]			
Y	US 2009/122953 A1 (IMAI YASUHIRO [JP]) 14 May 2009 (2009-05-14)	2-6, 13-20, 68-72, 80-86		
	abstract; figures 1-5 paragraphs [0004] - [0006], [0029] - [0063]			
X	US 2011/210261 A1 (MAURER JR CALVIN R [US]) 1 September 2011 (2011-09-01)	1,21, 30-34, 37-46, 48-52, 55,56, 88-90		
Y	abstract; figures 1-2D, 15-16, paragraphs [0006] - [0008], [0011] - [0012], [0017] - [0023] paragraphs [0045] - [5390]	2-20, 27-29, 35,36, 53,54, 57, 66-86, 91,94,95		
X Y	US 2003/048868 A1 (BAILEY ERIC M [US] ET AL) 13 March 2003 (2003-03-13) abstract; figures 1-3	1,30-34, 46,88-90 2-20,		
-	paragraphs [0003] - [0004], [0021] - [0025], [0037] - [0046]	27-29, 35,36, 57, 66-86, 91,94,95		
	- <i>/</i>			

International application No
PCT/US2012/044513

		PCT/US2012/044513				
C(Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT						
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.				
Υ	US 2002/191751 A1 (BOGATU IOAN NICULAE [US] ET AL) 19 December 2002 (2002-12-19) abstract; figures 2-3 table 1 paragraphs [0009] - [0016], [0031] - [0038]	35,36				
Y	US 2010/111388 A1 (SEPPI EDWARD J [US] ET AL) 6 May 2010 (2010-05-06) abstract; figures 1, 4A-G paragraphs [0023] - [0027], [0043] - [0058]	35,36				
Y	US 2010/119035 A1 (KARCH THOMAS [DE]) 13 May 2010 (2010-05-13) abstract; figures 1-11 paragraphs [0007] - [0020], [0048] - [0063]	35,36				
X	US 2006/182326 A1 (SCHILDKRAUT JAY S [US] ET AL) 17 August 2006 (2006-08-17)	1,21,30, 31, 48-56, 87-90				
Y	abstract; figures 1-4 paragraphs [0009] - [0010], [0018] - [0025], [0038] - [0045], [0059] - [0060]	53,54				
Υ	US 2010/080354 A1 (FU DONGSHAN [US] ET AL) 1 April 2010 (2010-04-01) paragraphs [0016] - [0026], [0031] - [0034]; figures 1-4, 7	53,54				
Y	US 2006/002615 A1 (FU DONGSHAN [US] ET AL) 5 January 2006 (2006-01-05) abstract; figures 1A-B, 2, 4 paragraphs [0052] - [0067]	53,54				

International application No. PCT/US2012/044513

INTERNATIONAL SEARCH REPORT

Box No. II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)
This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
Claims Nos.: because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box No. III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
see additional sheet
As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fees, this Authority did not invite payment of additional fees.
3. X As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.: 1-21, 27-57, 66-91, 94, 95
4. No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
The additional search fees were accompanied by the applicant's protest and, where applicable, the payment of a protest fee. The additional search fees were accompanied by the applicant's protest but the applicable protest fee was not paid within the time limit specified in the invitation. X No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. claims: 1-20, 27-29, 57, 66-86, 88-91, 94, 95

Image-guided radiation therapy system and method. A contribution to the art might reside in the special technical features of the weighted combination of the first and second sets of image data, solving the problem of optimising an enhanced patient image.

2. claims: 22-26

Image-guided radiation therapy system.
A contribution to the art might reside in the features of a variable aperture collimator, solving the problem of ensuring an accurate radiation beam shape.

3. claims: 30-47

Image-guided radiation therapy system. A contribution to the art might reside in the structural or functional features of the X-ray sources, solving the problem of providing an improved dual-energy X-ray imaging system.

4. claims: 21, 48-56, 87

Image-guided radiation therapy system. A contribution to the art might reside in the features of registering image data with a digital radiograph and corresponding target tracking, thereby solving the problem of enabling real-time adjustment of the radiation treatment.

5. claims: 58-65, 92, 93

Image-guided radiation therapy system. A contribution to the art might reside in the features of using tomosynthetic volumetric image data, solving the problem of providing a radiation therapy system with improved multi-energy volumetric imaging system.

Information on patent family members

International application No PCT/US2012/044513

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
US 2009274269 A	A1 05-11-2009	US 2009274269 A1 WO 2009135202 A1	05-11-2009 05-11-2009
	A1 19-11-2009	NONE	
US 6683934 E	31 27-01-2004	NONE	
	A1 01-01-2009	US 2009003528 A1 US 2012265060 A1	01-01-2009 18-10-2012
US 2009122953 /	A1 14-05-2009	CN 101433464 A JP 2009118887 A US 2009122953 A1	20-05-2009 04-06-2009 14-05-2009
US 2011210261 /	A1 01-09-2011	EP 2539020 A1 US 2011210261 A1 US 2011211665 A1 WO 2011106433 A1	02-01-2013 01-09-2011 01-09-2011 01-09-2011
US 2003048868 /	13-03-2003	NONE	
US 2002191751 /	19-12-2002	NONE	
US 2010111388 /	A1 06-05-2010	US 2005084073 A1 US 2010111388 A1 WO 2005037074 A2	28-04-2005
US 2010119035 /	13-05-2010	DE 102008056891 A1 US 2010119035 A1	27-05-2010 13-05-2010
US 2006182326 /	A1 17-08-2006	CN 101237907 A EP 1896132 A1 JP 2008528102 A US 2006182326 A1 WO 2006078386 A2	06-08-2008 12-03-2008 31-07-2008 17-08-2006 27-07-2006
US 2010080354 /	A1 01-04-2010	US 2010080354 A1 WO 2010039404 A1	01-04-2010 08-04-2010
US 2006002615 /	A1 05-01-2006	EP 1779298 A2 US 2006002615 A1 US 2009091567 A1 WO 2006011928 A2	02-05-2007 05-01-2006 09-04-2009 02-02-2006