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Biological equivalent dose informed flash radiation therapy

Abstract

A control circuit accesses information representing radiation dose deposition as a function of time for a particular patient as well as at least one biological parameter for that particular patient. The control circuit then determines a biological equivalent dose for the particular patient as a function of both the information representing radiation dose deposition as a function of time and the at least one biological parameter to provide a determined biological equivalent dose for the particular patient.

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References Cited

U.S. PATENT DOCUMENTS

Patent No.	Issued Date	Patentee Name	U.S. Cl.	CPC
7206377	12/2006	Svatos	378/65	A61N 5/103
7519150	12/2008	Romesberg, III	378/65	A61N 5/1031
7551717	12/2008	Tomé	378/65	A61B 5/6831
7831289	12/2009	Riker	378/65	G16H 70/20
7835493	12/2009	Keall	378/65	A61N 5/1042
9192782	12/2014	Grimm	N/A	A61N 5/1031
9289627	12/2015	Otto	N/A	A61N 5/1031
9387344	12/2015	Sgouros	N/A	A61B 6/5217
9387345	12/2015	Nord	N/A	A61N 5/1031
9486649	12/2015	Bert	N/A	A61N 5/107
9597529	12/2016	Lidestri	N/A	A61N 5/1043
9839793	12/2016	Harada	N/A	A61N 5/1075
10029121	12/2017	Li	N/A	A61B 5/055
10076673	12/2017	Ranganathan	N/A	A61N 5/1077
10166404	12/2018	Gemmel	N/A	A61N 5/1048
10252081	12/2018	Kauppinen	N/A	A61N 5/1042
10441819	12/2018	Sayeed	N/A	A61N 5/1071
10456600	12/2018	Owens	N/A	A61N 5/1081
10556124	12/2019	Zhang	N/A	A61N 5/1047
10556125	12/2019	Kuusela	N/A	A61N 5/1031
10688320	12/2019	Voronenko	N/A	A61N 5/1045
10702708	12/2019	Miettinen	N/A	A61N 5/1031
10828511	12/2019	Zhang	N/A	A61N 5/1031
10918886	12/2020	Smith	N/A	G16H 50/50
10940334	12/2020	Fishman	N/A	A61N 5/1083
11065471	12/2020	Wu	N/A	G16H 20/40
11090508	12/2020	Folkerts	N/A	A61N 5/1031
11103727	12/2020	Folkerts	N/A	G06T 7/0012
11167153	12/2020	Bokrantz	N/A	A61N 5/1031
11291861	12/2021	Cooley	N/A	A61N 5/1045
11517766	12/2021	Bzdusek	N/A	A61N 5/1039
11541252	12/2022	Folkerts	N/A	G16H 20/40

11559701	12/2022	Fan	N/A	A61N 5/1075
11590363	12/2022	Petaja	N/A	G16H 30/40
11612765	12/2022	Fallone	600/427	A61N 5/107
11642550	12/2022	Peltola	378/65	A61N 5/103
11648418	12/2022	Owens	378/65	A61N 5/1039
11654299	12/2022	Nord	378/65	A61N 5/103
11679273	12/2022	Kauppinen	600/1	A61N 5/1031
11786753	12/2022	Traneus	600/1	A61N 5/103
11786755	12/2022	Ohishi	600/1	A61N 5/1039
11813479	12/2022	Czeizler	N/A	A61N 5/103
11826560	12/2022	Zhang	N/A	A61N 5/1031
11839776	12/2022	Eriksson	N/A	A61N 5/1037
11938341	12/2023	Hirvonen	N/A	A61N 5/1031
11992703	12/2023	Lansonneur	N/A	A61N 5/1071
12059577	12/2023	Peltola	N/A	A61N 5/1036
12115386	12/2023	Voronenko	N/A	A61N 5/1043
12138472	12/2023	Labarbe	N/A	A61N 5/10
12138476	12/2023	Czeizler	N/A	A61N 5/1031
12138477	12/2023	Ropo	N/A	A61N 5/1048
12138478	12/2023	Peltola	N/A	A61N 5/1031
12138480	12/2023	Zhang	N/A	A61N 5/1031
12138483	12/2023	Czeizler	N/A	A61N 5/1031
12145001	12/2023	van den Heuvel	N/A	G16H 20/40
12157013	12/2023	Peltola	N/A	A61N 5/103
12168145	12/2023	Korhonen	N/A	A61N 5/1039
12179038	12/2023	Fredriksson	N/A	A61N 5/103
2009/0052623	12/2008	Tome	N/A	N/A
2016/0287903	12/2015	Sgouros	N/A	N/A
2021/0128946	12/2020	Smith	N/A	N/A
2021/0346719	12/2020	Folkerts	N/A	N/A
2021/0393982	12/2020	Lansonneur	N/A	N/A
2021/0402213	12/2020	Labarbe	N/A	N/A
2022/0296927	12/2021	Cooley	N/A	N/A
2022/0409927	12/2021	Petaja	N/A	N/A
2023/0115222	12/2022	Traneus	N/A	N/A

OTHER PUBLICATIONS

Krieger, Miriam et al.; A quantitative FLASH effectiveness model to reveal potentials and pitfalls of high dose rate proton therapy. Medical Physics, 49(3), 2026-2038, Dec. 2021. cited by applicant
Mazal, Alejandro et al.; FLASH and minibeam in radiation therapy: the effect of microstructures on time and space and their potential application to protontherapy. British Institute of Radiology, 2020;93:110720190807, 17 pages. cited by applicant

Petersson, Kristoffer et al., A Quantitative Analysis of the Role of Oxygen Tension in FLASH Radiation Therapy. International Journal of Radiation Oncology Biology Physics. 2020; 107(3):539-547. cited by applicant

Extended European Search Report from related European Patent Application No. 24180112.5 dated Aug. 26, 2024; 11 pages. cited by applicant

Background/Summary

TECHNICAL FIELD

(1) These teachings relate generally to treating a patient's planning target volume with energy pursuant to an energy-based treatment plan and more particularly to FLASH radiation therapy treatment plans.

BACKGROUND

(2) The use of energy to treat medical conditions comprises a known area of prior art endeavor. For example, radiation therapy comprises an important component of many treatment plans for reducing or eliminating unwanted tumors. Unfortunately, applied energy does not inherently discriminate between unwanted material and adjacent tissues, organs, or the like that are desired or even critical to continued survival of the patient. As a result, energy such as radiation is ordinarily applied in a carefully administered manner to at least attempt to restrict the energy to a given target volume. A so-called radiation treatment plan often serves in the foregoing regards.

(3) Many radiation treatment plans specify values for each of a variety of treatment-platform parameters during a single field or during each of a plurality of sequential fields. Treatment plans for radiation treatment sessions are often automatically generated through a so-called optimization process. As used herein, "optimization" will be understood to refer to improving a candidate treatment plan without necessarily ensuring that the optimized result is, in fact, the singular best solution. Such optimization often includes automatically adjusting one or more physical treatment parameters (often while observing one or more corresponding limits in these regards) and mathematically calculating a likely corresponding treatment result (such as a level of dosing) to identify a given set of treatment parameters that represent a good compromise between the desired therapeutic result and avoidance of undesired collateral effects.

(4) FLASH energy treatment is a known area of prior art endeavor, albeit a relatively new area of consideration. FLASH radiotherapy is a technique (at least for photon, electron, and proton treatments) that employs very brief, very high dose rates (utilizing large beam currents). FLASH treatments hold the promise of shortening treatment time to just one to three 1-second sessions while also potentially reducing side effects, perhaps considerably. For example, a significant sparing of normal tissue notwithstanding iso-effective tumor growth delay has been demonstrated through very brief irradiation at dose rates on the order of 40 Gy/s. This sparing of normal tissue has been dubbed the FLASH effect.

(5) Unfortunately, the applicant has determined that there is much unknown about FLASH treatments, including how to take into account one or more biological effects pertaining to a patient when prescribing a FLASH treatment, optimizing a FLASH treatment, quality checking a FLASH treatment plan, and/or administering a FLASH treatment plan.

Description

BRIEF DESCRIPTION OF THE DRAWINGS

(1) The above needs are at least partially met through provision of the biological equivalent dose informed FLASH radiation therapy described in the following detailed description, particularly when studied in conjunction with the drawings, wherein:

(2) FIG. 1 comprises a block diagram as configured in accordance with various embodiments of these teachings;

(3) FIG. 2 comprises a flow diagram as configured in accordance with various embodiments of these teachings;

(4) FIG. 3 comprises graphs as configured in accordance with various embodiments of these teachings; and

(5) FIG. 4 comprises a graph as configured in accordance with various embodiments of these teachings.

(6) Elements in the figures are illustrated for simplicity and clarity and have not necessarily been drawn to scale. For example, the dimensions and/or relative positioning of some of the elements in the figures may be exaggerated relative to other elements to help to improve understanding of various embodiments of the present teachings. Also, common but well-understood elements that are useful or necessary in a commercially feasible embodiment are often not depicted in order to facilitate a less obstructed view of these various embodiments of the present teachings. Certain actions and/or steps may be described or depicted in a particular order of occurrence while those skilled in the art will understand that such specificity with respect to sequence is not actually required. The terms and expressions used herein have the ordinary technical meaning as is accorded to such terms and expressions by persons skilled in the technical field as set forth above except where different specific meanings have otherwise been set forth herein. The word “or” when used herein shall be interpreted as having a disjunctive construction rather than a conjunctive construction unless otherwise specifically indicated.

DETAILED DESCRIPTION

(7) Generally speaking, pursuant to these various embodiments, a control circuit accesses information representing radiation dose deposition as a function of time for a particular patient as well as at least one biological parameter for that particular patient. The control circuit then determines a biological equivalent dose for the particular patient as a function of both the information representing radiation dose deposition as a function of time and the at least one biological parameter to provide a determined biological equivalent dose for the particular patient.

(8) By one approach, the foregoing accommodates determining a plurality of biological equivalent doses for the particular patient.

(9) By one approach, the aforementioned information representing radiation dose deposition as a function of time comprises both dose and dose rate information.

(10) By one approach, the aforementioned at least one biological parameter comprises at least one of: normal tissue complication probability information; oxygen level information; linear energy transfer information; an identified tissue; an identified organ; a specified portion of an identified organ; a temporal-based parameter; radiosensitivity of a given tissue/organ; patient image information; a tissue type; a blood oxygen level; a blood type; a medication; chemotherapy information; prior surgery information; immunotherapy information; prior medical treatment information; demographic information; biological age information; genetic marker information; a time of day; a tumor type; pH (potential hydrogen) balance information; temperature information; blood pressure information; blood panel information; and/or immunity health information.

(11) By one approach, these teachings will further accommodate any one or more of: generating a radiation treatment prescription as a function, at least in part, of the determined biological equivalent dose for the particular patient; optimizing a FLASH radiation therapy treatment as a function, at least in part, of the determined biological equivalent dose for the particular patient to provide an optimized FLASH radiation therapy treatment; quality evaluating an optimized FLASH radiation therapy treatment plan as a function, at least in part, of the determined biological equivalent dose for the particular patient; and/or administering radiation to the particular patient as a function of the aforementioned optimized FLASH radiation therapy treatment plan.

(12) So configured, these teachings can capture the joint effect of dose distributions, dose-rate distributions, and one or more items of relevant biological data into one metric that correlates to the effects observed at a biological level.

(13) These and other benefits may become clearer upon making a thorough review and study of the following detailed description. Referring now to the drawings, and in particular to FIG. 1, an illustrative apparatus **100** that is compatible with many of these teachings will first be presented.

(14) In this particular example, the enabling apparatus **100** includes a control circuit **101**. Being a “circuit,” the control circuit **101** therefore comprises structure that includes at least one (and typically many) electrically-conductive paths (such as paths comprised of a conductive metal such as copper or silver) that convey electricity in an ordered manner, which path(s) will also typically include corresponding electrical components (both passive (such as resistors and capacitors) and active (such as any of a variety of semiconductor-based devices) as appropriate) to permit the circuit to effect the control aspect of these teachings.

(15) Such a control circuit **101** can comprise a fixed-purpose hard-wired hardware platform (including but not limited to an application-specific integrated circuit (ASIC) (which is an integrated circuit that is customized by design for a particular use, rather than intended for general-purpose use), a field-programmable gate array (FPGA), and the like) or can comprise a partially or wholly-programmable hardware platform (including but not limited to microcontrollers, microprocessors, and the like). These architectural options for such structures are well known and understood in the art and require no further description here. This control circuit **101** is configured (for example, by using corresponding programming as will be well understood by those skilled in the art) to carry out one or more of the steps, actions, and/or functions described herein.

(16) The control circuit **101** operably couples to a memory **102**. This memory **102** may be integral to the control circuit **101** or can be physically discrete (in whole or in part) from the control circuit **101** as desired. This memory **102** can also be local with respect to the control circuit **101** (where, for example, both share a common circuit board, chassis, power supply, and/or housing) or can be partially or wholly remote with respect to the control circuit **101** (where, for example, the memory **102** is physically located in another facility, metropolitan area, or even country as compared to the control circuit **101**).

(17) In addition to information such as optimization information for a particular patient, information regarding a particular radiation treatment platform, information representing radiation dose deposition as a function of time for the particular patient, and at least one biological parameter for the particular patient as described herein, this memory **102** can serve, for example, to non-transitorily store the computer instructions that, when executed by the control circuit **101**, cause the control circuit **101** to behave as described herein. (As used herein, this reference to “non-transitorily” will be understood to refer to a non-ephemeral state for the stored contents (and hence excludes when the stored contents merely constitute signals or waves) rather than volatility of the storage media itself and hence includes both non-volatile memory (such as read-only memory (ROM) as well as volatile memory (such as a dynamic random access memory (DRAM).)

(18) By one optional approach the control circuit **101** also operably couples to a user interface **103**. This user interface **103** can comprise any of a variety of user-input mechanisms (such as, but not limited to, keyboards and keypads, cursor-control devices, touch-sensitive displays, speech-recognition interfaces, gesture-recognition interfaces, and so forth) and/or user-output mechanisms (such as, but not limited to, visual displays, audio transducers, printers, and so forth) to facilitate receiving information and/or instructions from a user and/or providing information to a user.

(19) If desired the control circuit **101** can also operably couple to a network interface (not shown). So configured the control circuit **101** can communicate with other elements (both within the apparatus **100** and external thereto) via the network interface. Network interfaces, including both wireless and non-wireless platforms, are well understood in the art and require no particular elaboration here.

(20) By one approach, a computed tomography apparatus **106** and/or other imaging apparatus **107** as are known in the art can source some or all of any desired patient-related imaging information.

(21) In this illustrative example the control circuit **101** is configured to ultimately output an

optimized energy-based treatment plan (such as, for example, an optimized radiation treatment plan **113**). This energy-based treatment plan typically comprises specified values for each of a variety of treatment-platform parameters during each of a plurality of sequential exposure fields. (It should be noted that these teachings are also applicable to a plan that makes use of only a single “field,” as when, for example, the gantry is constantly in motion and radiation is constantly applied.) In this case the energy-based treatment plan is generated through an optimization process, examples of which are provided further herein.

(22) By one approach the control circuit **101** can operably couple to an energy-based treatment platform **114** that is configured to deliver therapeutic energy **112** to a corresponding patient **104** having at least one treatment volume **105** and also one or more organs-at-risk (represented in FIG. **1** by a first through an Nth organ-at-risk **108** and **109**) in accordance with the optimized energy-based treatment plan **113**. These teachings are generally applicable for use with any of a wide variety of energy-based treatment platforms/apparatuses. In a typical application setting the energy-based treatment platform **114** will include an energy source such as a radiation source **115** of ionizing radiation **116**. By one approach this energy-based treatment platform **114** is particularly configured to deliver FLASH radiation therapy.

(23) By one approach this radiation source **115** can be selectively moved via a gantry along an arcuate pathway (where the pathway encompasses, at least to some extent, the patient themselves during administration of the treatment). The arcuate pathway may comprise a complete or nearly complete circle as desired. By one approach the control circuit **101** controls the movement of the radiation source **115** along that arcuate pathway, and may accordingly control when the radiation source **115** starts moving, stops moving, accelerates, de-accelerates, and/or a velocity at which the radiation source **115** travels along the arcuate pathway.

(24) As one illustrative example, the radiation source **115** can comprise, for example, a radio-frequency (RF) linear particle accelerator-based (linac-based) x-ray source. A linac is a type of particle accelerator that greatly increases the kinetic energy of charged subatomic particles or ions by subjecting the charged particles to a series of oscillating electric potentials along a linear beamline, which can be used to generate ionizing radiation (e.g., X-rays) **116** and high energy electrons.

(25) A typical energy-based treatment platform **114** may also include one or more support apparatuses **110** (such as a couch) to support the patient **104** during the treatment session, one or more patient fixation apparatuses **111**, a gantry or other movable mechanism to permit selective movement of the radiation source **115**, and one or more energy-shaping apparatuses (for example, beam-shaping apparatuses **117** such as jaws, multi-leaf collimators, and so forth) to provide selective energy shaping and/or energy modulation as desired.

(26) In a typical application setting, it is presumed herein that the patient support apparatus **110** is selectively controllable to move in any direction (i.e., any X, Y, or Z direction) during an energy-based treatment session by the control circuit **101**. As the foregoing elements and systems are well understood in the art, further elaboration in these regards is not provided here except where otherwise relevant to the description.

(27) Referring now to FIG. **2**, a process **200** that can be carried out, for example, in conjunction with the above-described application setting (and more particularly via the aforementioned control circuit **101**) will be described. Generally speaking, this process **200** can serve to facilitate generating an optimized radiation treatment plan **113** to thereby facilitate treating a particular patient with therapeutic radiation (and in particular FLASH radiation therapy) using a particular radiation treatment platform per that optimized radiation treatment plan.

(28) At block **201**, this process **200** provides for accessing information representing radiation dose deposition as a function of time for a particular patient. By one approach, this information comprises both dose and dose rate information.

(29) At block **202**, the control circuit **101** accesses at least one biological parameter for the

particular patient. These teachings are highly flexible in practice and will accommodate a wide variety of biological parameters. Examples of possibly beneficial biological parameters include, but are not limited to, any one or more of: normal tissue complication probability information (corresponding, for example, to an effect that is measured but which is not necessarily an input of the biological equivalent dose); oxygen level information (including, for example, a general oxygen level in a given tissue or anatomical structure, an oxygen level in the patient's blood, and/or even down to oxygen levels at a per-voxel or a sub-group-of-voxels level); linear energy transfer information; an identified tissue; an identified organ; a specified portion of an identified organ; a temporal-based parameter; radiosensitivity of a given tissue/organ (which information captures, for example, the alpha/beta ratio of a given organ); patient image information (including but not limited to imaging derived from computed tomography, magnetic resonance imaging, and/or positron emission tomography, as well as imaging that presents segmentation information regarding the patient); a tissue type; a blood oxygen level; a blood type; a medication (including one or more prescription or non-prescription medicines and/or supplements); chemotherapy information (including previous chemotherapy treatments, current chemotherapy treatments, and/or planned/scheduled chemotherapy treatments); prior surgery information; immunotherapy information; prior medical treatment information (including, for example, radio therapy, ultrasound, and so forth); demographic information (such as, but not limited to, age, sex, ethnicity, health factors (such as body mass index, autoimmune conditions, blood sugar levels, hemophilia, high blood pressure, and so forth), lifestyle factors (such as smoking and alcohol history and practices), stress factors, blood panel information (such as white blood cell count, cholesterol levels, and so forth), and even DNA information); biological age information; genetic marker information; a time of day; a tumor type; pH (potential hydrogen) balance information; temperature information; blood pressure information; blood panel information; and/or immunity health information.

(30) At block **203**, the control circuit **101** then determines a biological equivalent dose for the particular patient as a function of both the information representing radiation dose deposition as a function of time and the at least one biological parameter to thereby provide a determined biological equivalent dose for the particular patient. By one approach, this can comprise determining a plurality of biological equivalent doses for the particular patient to thereby provide a plurality of determined biological equivalent doses for the particular patient.

(31) By one approach, the foregoing comprises determining at least one biological equivalent dose on a per voxel basis (for example, for voxels within a patient target volume and/or within an organ-at-risk). The latter can be useful, for example, when the dose versus time information comprises a set of three-dimensional points within a patient image. By another approach, the foregoing may comprise determining at least one biological equivalent dose on a per patient structure basis.

(32) FIG. 3 presents an illustrative example in these regards, it being understood that the details of this example are intended to serve an illustrative purpose and are not intended to suggest any limitations with respect to these teachings.

(33) In this example, the dose **301** and dose-rate **302** distributions are combined into a biological equivalent dose (BED) **303** that correlates with Normal Tissue Complication Probability (NTCP) **304**. The relationship between dose (D) **301**, dose-rate (DR) **302** and Normal Tissue Complication Probability (NTCP) **304** is complex and not necessarily intuitive. It will be appreciated that combining the dose and dose rate into one distribution $BED=f(D,DR)$ allows better visualizations of their correlations with NTCP.

(34) As noted above, these teachings will accommodate using any of a variety of biological inputs. As but one illustrative example, the level of oxygen within a tissue can be used in conjunction with the dose-rate information to generate an oxygen enhancement ratio weighted biological dose.

(35) By one optional approach, referring to optional block **204**, these teachings will accommodate generating a radiation treatment prescription as a function, at least in part, of the above-described

biological equivalent dose for the particular patient. Those skilled in the art will understand that the radiation treatment prescription expresses the general dosing goals envisioned and intended by a prescribing oncologist. A radiation treatment prescription is accordingly silent regarding such things as treatment machine geometry, control points, treatment fields, and so forth. (To be clear, it will be understood that radiation treatment prescriptions are different from optimization objectives, the latter being very much specifically designed to reflect and accommodate the technical details and specifications of a particular radiation treatment platform, specific details regarding the patient's presentation, and/or other physical details pertaining to a particular application setting treatment machine geometry, control points, treatment fields, and so forth.)

(36) In lieu of the foregoing, or in combination therewith, by another optional approach and as expressed at optional block **205**, these teachings will accommodate optimizing a FLASH radiation therapy treatment plan as a function, at least in part, of the determined biological equivalent dose for the particular patient to provide an optimized FLASH radiation therapy treatment plan. By one approach, the foregoing can comprise calculating a plurality of biological equivalent dose values as part of the optimization loop. These different biological equivalent dose values may correspond to different patient structures, patient sub-structures, and/or voxels and may be included as part of a cost function that is iteratively reduced to a local (if not global) minimum. For example, a biological equivalent dose value AB can be iteratively generated until a seemingly most optimal value is found.

(37) When optimizing a FLASH plan, better performances may be expected when the analytical expression of the gradient, i.e., the derivative of the distribution to optimize with respect to the optimization variables, is known. Generally speaking, the variables to optimize include the beamlet intensities for Intensity Modulated Radiotherapy or the weights of the individual pencil beams for Intensity Modulated Proton Therapy. In both cases, the optimization variables can be expressed in Monitor Units (MU). For a model where the BED depends on the dose-rate DR and the dose D only, the gradient with respect to the optimization variables MU can be expressed as:

$$(38) \frac{\partial \text{BED}}{\partial \text{MU}} = \frac{\partial \text{BED}}{\partial D} \cdot \text{Math.} \frac{\partial D}{\partial \text{MU}} + \frac{\partial \text{BED}}{\partial \text{DR}} \cdot \text{Math.} \frac{\partial \text{DR}}{\partial \text{MU}} .$$

(39) For a model where the BED depends on the dose-rate DR, the dose D and any other biological metric X (e.g., the oxygen enhancement ratio, or the linear energy transfer), the gradient with respect to the optimization variables MU can be expressed as:

$$(40) \frac{\partial \text{BED}}{\partial \text{MU}} = \frac{\partial \text{BED}}{\partial D} \cdot \text{Math.} \frac{\partial D}{\partial \text{MU}} + \frac{\partial \text{BED}}{\partial \text{DR}} \cdot \text{Math.} \frac{\partial \text{DR}}{\partial \text{MU}} + \frac{\partial \text{BED}}{\partial X} \cdot \text{Math.} \frac{\partial X}{\partial \text{MU}} .$$

(41) These teachings will also accommodate allowing the BED to be a function of dose vs. delivery time curve, in which case, the impact on BED for a given dose delivery sequence could be evaluated with the aid of detailed treatment delivery modeling. Likewise, the chain rule still applies, and includes the proton pencil beam spot list (scan pattern) as a variable that can be manipulated by the optimizer to achieve a BED goal.

(42) In lieu of the foregoing, or in any permutation or combination therewith, at optional block **206** this process **200** will accommodate quality evaluating an optimized FLASH radiation therapy treatment plan as a function, at least in part, of the determined biological equivalent dose for the particular patient. Such an approach may be beneficial when, for example, the biological equivalent dose corresponds to a three-dimensional distribution and quality is evaluated, at least in part, using that distribution as an input.

(43) And again in lieu of the foregoing, or in any permutation or combination therewith, at optional block **207** these teachings will accommodate administering radiation to the particular patient as a function of the aforementioned optimized FLASH radiation therapy treatment plan using, for example, the above-described radiation treatment platform **114**.

(44) To assist the treatment planner, these teachings will also accommodate providing one or more corresponding evaluation tools. Examples include, but are not limited to, biological equivalent dose volume histograms, biological equivalent dose distribution displayed on top of computed tomography images, biological equivalent dose distribution displayed per field, biological

equivalent dose influence matrices, and so forth.

(45) By way of illustration, and referring to FIG. 4, a standard dose volume histogram (DVH) 401, 402 can be presented along with a biological equivalent dose volume histogram (BEDVH) 403, 404 for an organ at risk and a target volume, respectively. The target coverage is relatively unchanged, but the BED received by the organ-at-risk is significantly lower as compared with the physical dose. This interface facilitates clinical decision making with respect to the treatment modality (FLASH as versus conventional).

(46) It will be appreciated that, while existing FLASH radiation therapy treatment plan optimization techniques tend to focus only on improving physical dose rates, such existing approaches also tend to neglect consideration of any FLASH-based biological effects; the present teachings, however, successfully attend to both considerations.

(47) Those skilled in the art will recognize that a wide variety of modifications, alterations, and combinations can be made with respect to the above described embodiments without departing from the scope of the invention, and that such modifications, alterations, and combinations are to be viewed as being within the ambit of the inventive concept.

Claims

1. A method comprising: by a control circuit: accessing information representing a radiation dose deposition as a function of time for a particular patient; accessing at least one biological parameter for the particular patient; and determining a biological equivalent dose for the particular patient as a function of both the information representing a radiation dose deposition as a function of time for the particular patient and the at least one biological parameter for the particular patient to provide a determined biological equivalent dose for the particular patient.
2. The method of claim 1, wherein determining a biological equivalent dose for the particular patient comprises determining a plurality of biological equivalent doses for the particular patient as a function of both the information representing a radiation dose deposition as a function of time for the particular patient and the at least one biological parameter for the particular patient to provide a plurality of determined biological equivalent doses for the particular patient.
3. The method of claim 1, wherein the information representing a radiation dose deposition as a function of time for the particular patient comprises both dose information and dose rate information.
4. The method of claim 1, wherein the at least one biological parameter for the particular patient comprises normal tissue complication probability information.
5. The method of claim 1, wherein the at least one biological parameter for the particular patient comprises oxygen level information.
6. The method of claim 1, wherein the at least one biological parameter for the particular patient comprises linear energy transfer information.
7. The method of claim 1, wherein the at least one biological parameter for the particular patient comprises at least one of: an identified tissue; an identified organ; a specified portion of an identified organ; a temporal-based parameter; a radiosensitivity of a given tissue/organ; patient image information; a tissue type; a blood oxygen level; a blood type; a medication; chemotherapy information; prior surgery information; immunotherapy information; prior medical treatment information; demographic information; biological age information; genetic marker information; a time of day; a tumor type; pH (potential hydrogen) balance information; temperature information; blood pressure information; blood panel information; and immunity health information.
8. The method of claim 1, further comprising: optimizing a FLASH radiation therapy treatment plan as a function, at least in part, of the determined biological equivalent dose for the particular patient to provide an optimized FLASH radiation therapy treatment plan.
9. The method of claim 8, further comprising: administering radiation to the particular patient as a

function of the optimized FLASH radiation therapy treatment plan.

10. The method of claim 8, further comprising: quality evaluating the optimized FLASH radiation therapy treatment plan as a function, at least in part, of the determined biological equivalent dose for the particular patient.

11. The method of claim 1, further comprising: generating a radiation treatment prescription as a function, at least in part, of the determined biological equivalent dose for the particular patient.

12. An apparatus comprising: a memory having stored therein information representing a radiation dose deposition as a function of time for a particular patient and at least one biological parameter for the particular patient; and a control circuit operably coupled to the memory and configured to: access the information representing a radiation dose deposition as a function of time for a particular patient; access the at least one biological parameter for the particular patient; and determine a biological equivalent dose for the particular patient as a function of both the information representing a radiation dose deposition as a function of time for the particular patient and the at least one biological parameter for the particular patient to provide a determined biological equivalent dose for the particular patient.

13. The apparatus of claim 12, wherein the control circuit is configured to develop a biological equivalent dose for the particular patient by determining a plurality of biological equivalent doses for the particular patient as a function of both the information representing a radiation dose deposition as a function of time for the particular patient and the at least one biological parameter for the particular patient to thereby provide a plurality of determined biological equivalent doses for the particular patient.

14. The apparatus of claim 12, wherein the information representing a radiation dose deposition as a function of time for the particular patient comprises, at least in part, dose rate information.

15. The apparatus of claim 12, wherein the information representing a radiation dose deposition as a function of time for the particular patient comprises both dose information and dose rate information.

16. The apparatus of claim 12, wherein the at least one biological parameter for the particular patient comprises normal tissue complication probability information.

17. The apparatus of claim 12, wherein the at least one biological parameter for the particular patient comprises oxygen level information.

18. The apparatus of claim 12, wherein the at least one biological parameter for the particular patient comprises linear energy transfer information.

19. The apparatus of claim 12, wherein the at least one biological parameter for the particular patient comprises at least one of: an identified tissue; an identified organ; a specified portion of an identified organ; a temporal-based parameter; a radiosensitivity of a given tissue/organ; patient image information; a tissue type; a blood oxygen level; a blood type; a medication; chemotherapy information; prior surgery information; immunotherapy information; prior medical treatment information; demographic information; biological age information; genetic marker information; a time of day; a tumor type; pH (potential hydrogen) balance information; temperature information; blood pressure information; blood panel information; and immunity health information.

20. The apparatus of claim 12, wherein the control circuit is further configured to: optimize a FLASH radiation therapy treatment plan as a function, at least in part, of the determined biological equivalent dose for the particular patient to provide an optimized FLASH radiation therapy treatment plan.

21. The apparatus of claim 20, wherein the control circuit is further configured to: administer radiation to the particular patient as a function of the optimized FLASH radiation therapy treatment plan.

22. The apparatus of claim 20, wherein the control circuit is further configured to: quality evaluate the optimized FLASH radiation therapy treatment plan as a function, at least in part, of the determined biological equivalent dose for the particular patient.

23. The apparatus of claim 12, wherein the control circuit is further configured to: generate a radiation treatment prescription as a function, at least in part, of the determined biological equivalent dose for the particular patient.
