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Lee et al.(10) **Pub. No.: US 2018/0218508 A1**(43) **Pub. Date: Aug. 2, 2018**(54) **PEN-TYPE MEDICAL FLUORESCENT
IMAGING DEVICE AND SYSTEM FOR
ALIGNING MULTIPLE FLUORESCENT
IMAGES USING THE SAME***G06T 7/174* (2006.01)*A61B 5/00* (2006.01)*A61K 49/00* (2006.01)(52) **U.S. CL.**CPC *G06T 7/37* (2017.01); *G06T 7/12*(2017.01); *G06T 7/174* (2017.01); *A61B**5/0071* (2013.01); *A61B 5/7425* (2013.01);*A61B 2562/0233* (2013.01); *G06T 2207/10064*(2013.01); *G06T 2207/10016* (2013.01); *G06T**2207/30096* (2013.01); *G06T 2207/30101*(2013.01); *A61B 2560/0418* (2013.01); *A61K**49/0034* (2013.01)(71) Applicant: **National Cancer Center**, Gyeonggi-do
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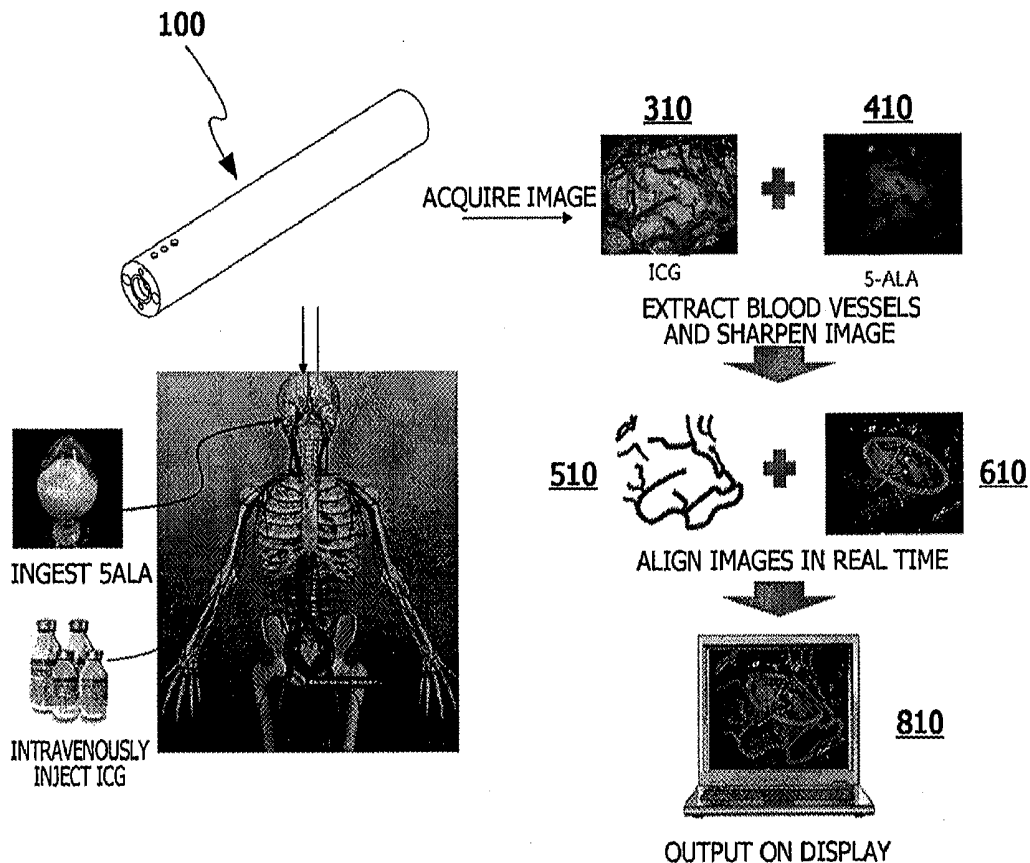
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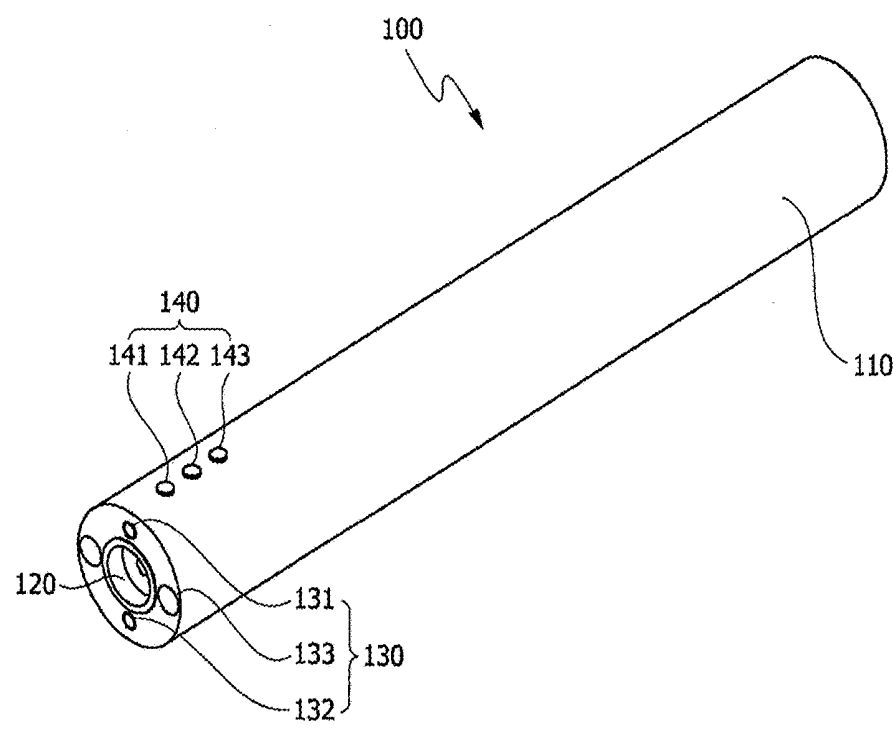
Aug. 3, 2015 (KR) 10-2015-0109516

Publication Classification(51) **Int. Cl.***G06T 7/37* (2006.01)*G06T 7/12* (2006.01)(57) **ABSTRACT**

The present invention relates to a pen-type medical fluorescent imaging device comprising: a probe which is elongated in the longitudinal direction and has an image capture unit at one end; a plurality of light source units surrounding the image capture unit; and a control unit for controlling the light source units, wherein the light source units comprise: a first light source for emitting light of a first wavelength range so that a blood vessel is marked by a first fluorescent material; and a second light source for emitting light of a second wavelength range so that a glioma is marked by a second fluorescent material, wherein the first and second light sources are selectively controlled by the control unit.

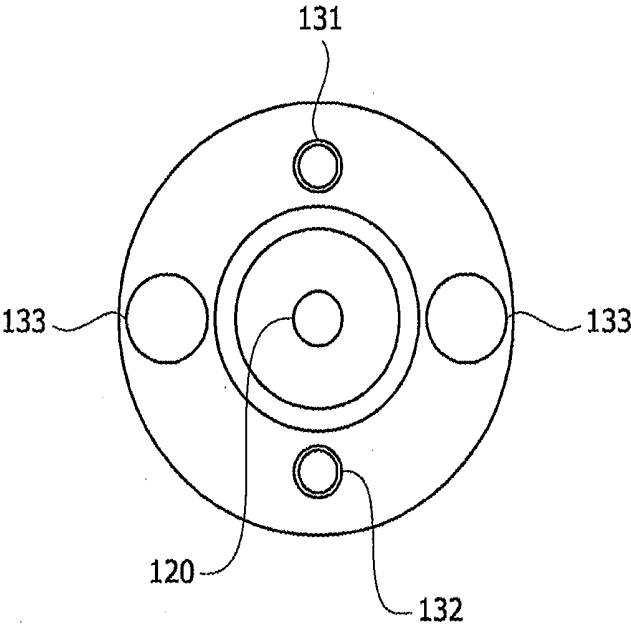


[Fig. 1]

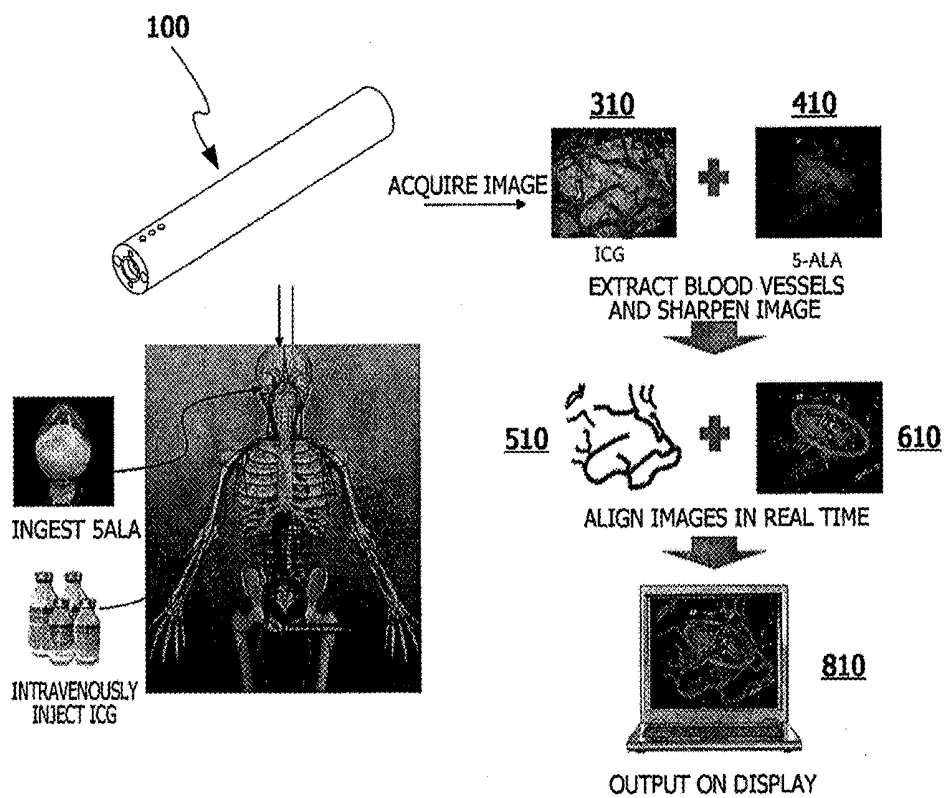


[Fig. 2]

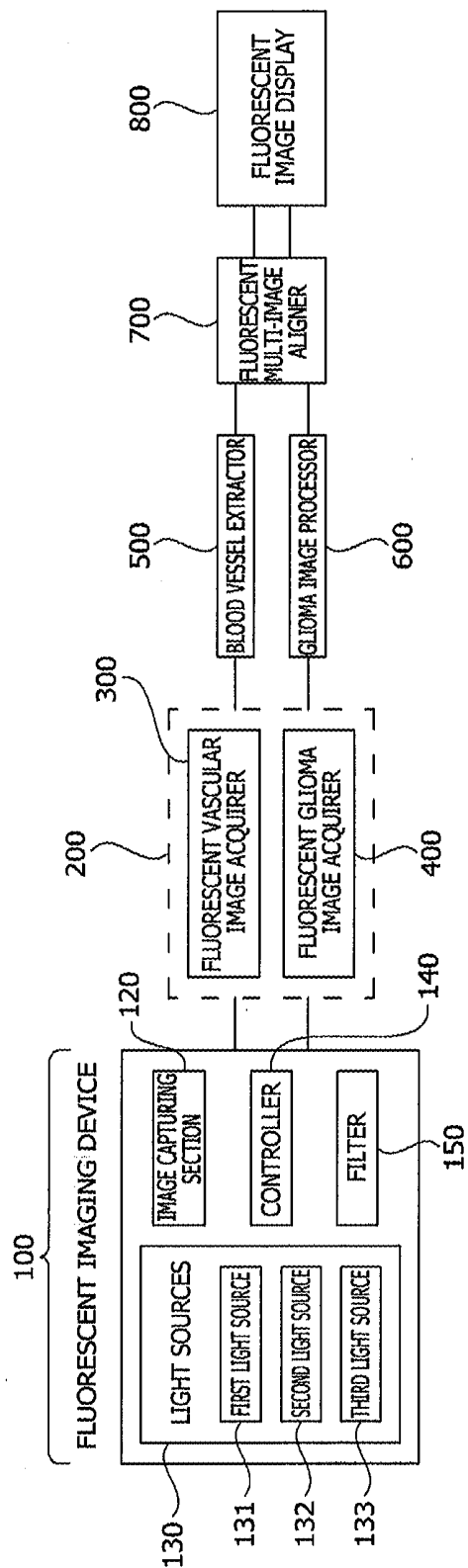
130



[Fig. 3]



[Fig. 4]



**PEN-TYPE MEDICAL FLUORESCENT
IMAGING DEVICE AND SYSTEM FOR
ALIGNING MULTIPLE FLUORESCENT
IMAGES USING THE SAME**

BACKGROUND

1. Field of the Invention

[0001] The present invention relates to a medical fluorescent imaging device and a system for aligning multiple fluorescent images using the same.

2. Discussion of Related Art

[0002] In general, various types of video equipment are being used to obtain biomedical imaging information at biological and medical sites. In such video equipment, in comparison to other imaging techniques, a biomedical imaging method employing light is widely being used due to its convenience of providing real-time information to an observer or a surgeon.

[0003] In addition, the rapid development of molecular imaging technology lately has made it possible to diagnose diseases and detect lesions. In particular, nuclear medicine imaging and magnetic resonance imaging (MRI) technology is attracting attention as examination systems that are very useful for diagnosing diseases. MRI technology is a method of obtaining anatomical, physiological, and biochemical information images of a body by using a phenomenon in which the spinning of hydrogen atoms is relaxed. MRI technology is an excellent diagnostic imaging technology that is not invasive towards body organs of a person or an animal and makes it possible to capture images thereof in real time.

[0004] However, in the diagnosis of a malignant glioma that is highly infiltrative, there are problems with such MRI in that it is difficult to determine the boundary between a malignant glioma and normal tissue.

[0005] During surgical operation on a tumor, it is necessary to check the positions and connections of cerebral blood vessels as well as the tumor for the safety of the patient. In particular, to increase the survival rate of patients and prevent recurrences of diseases, it is very important to completely remove a malignant tumor or a tumor and minimize damage to normal tissue by minimizing the removed parts.

[0006] Relevant to this, 5-aminolevulinic acid (5-ALA), which is a fluorescent luminescent material for marking tumors and blood vessels, is injected into the human body and then fluoresces due to generation of protoporphyrin IX (PpIX) derived from a material transformation process caused by metabolism in the body. With such a reaction process, 5-ALA serves as a target fluorescence marker targeting only cancer cells.

[0007] Also, indocyanine green (ICG), which is a fluorescent luminescent material for marking blood vessels, marks blood vessels and lymph nodes while circulating through the blood vessels and the lymph nodes.

[0008] In other words, 5-ALA is used to cause cancer cells to fluoresce, and ICG is used to cause blood vessels to fluoresce.

[0009] However, current surgical fluorescent imaging equipment for surgical microscopes may display only one type of fluorescent image due to one the presence of type of

fluorescent luminescent material, and thus displays only ICG fluorescent images. Also, since fluorescent imaging equipment is very large, there is an inconvenience in terms of using fluorescent imaging equipment only for surgery.

[0010] Consequently, an apparatus for accurately detecting and showing the position of a glioma and the positions of blood vessels at the same time is required for a patient's safety and ease of surgery.

SUMMARY OF THE INVENTION

[0011] The present invention is directed to a fluorescent imaging device capable of detecting and showing the positions of blood vessels and the position of a glioma at the same time for a patient's safety and ease of surgery.

[0012] The present invention is also directed to a system for aligning multiple fluorescent images which makes it possible for an operator to accurately and easily check the positions and connections of a tumor and blood vessels by acquiring multiple fluorescent images in which the positions of the blood vessels and the boundary of a glioma are marked to luminesce due to multiple fluorescent materials, and by aligning and displaying the acquired multiple fluorescent images in real time.

[0013] According to an aspect of the present invention, a pen-type medical fluorescent imaging device is provided including: a probe configured to extend in a longitudinal direction and include an image capturing section at one end; a plurality of light sources configured to be arranged to surround the image capturing section; and a controller configured to control the light sources. The light sources include a first light source configured to emit light of a first wavelength range so that blood vessels are marked by a first fluorescent material, and a second light source configured to emit light of a second wavelength range so that a glioma is marked by a second fluorescent material. The first and second light sources are selectively controlled by the controller.

[0014] According to another aspect of the present invention, a system for aligning multiple fluorescent images is provided, the system including: the pen-type medical fluorescent imaging device; a fluorescent multi-image acquirer configured to acquire an actual image of a target part, a fluorescent vascular image in which blood vessels are marked by the first fluorescent material, and a fluorescent glioma image in which a glioma is marked by the second fluorescent material; a blood vessel extractor configured to extract a vascular shape from the acquired fluorescent vascular image; a glioma image processor configured to process the acquired fluorescent glioma image so that a boundary of the glioma becomes distinct; a fluorescent multi-image aligner configured to align the extracted fluorescent vascular image and fluorescent glioma image in real time; and a fluorescent image display configured to display a fluorescent multi-image in which the fluorescent vascular image and the fluorescent glioma image are aligned.

BRIEF DESCRIPTION OF THE DRAWINGS

[0015] The above description and other objects, features and advantages of the present invention will become more apparent to those of ordinary skill in the art by describing in detail exemplary embodiments thereof with reference to the accompanying drawings, in which:

[0016] FIG. 1 is a perspective view of a pen-type medical fluorescent imaging device according to the present invention;

[0017] FIG. 2 is a front view of the pen-type medical fluorescent imaging device according to the present invention;

[0018] FIG. 3 is a conceptual diagram for facilitating understanding of the overall operation of a system for aligning multiple fluorescent images according to the present invention; and

[0019] FIG. 4 is a block diagram of the system for aligning multiple fluorescent images according to the present invention.

DETAILED DESCRIPTION OF EXEMPLARY EMBODIMENTS

[0020] The present invention relates to a pen-type medical fluorescent imaging device capable of detecting and showing the position of a glioma and the positions of blood vessels at the same time for a patient's safety and ease of surgery.

[0021] Here, a pen type denotes a small pen shape that may be conveniently and freely used with a hand like a pen, which is a writing instrument. According to the present invention, a probe 110 may be formed in a pen shape.

[0022] The pen-type medical fluorescent imaging device according to the present invention includes: a probe that extends in a longitudinal direction and includes an image capturing section at one end; a plurality of light sources that are arranged to surround the image capturing section; and a controller that controls the light sources. The light sources include a first light source that emits light of a first wavelength range so that blood vessels are marked by a first fluorescent material, and a second light source that emits light of a second wavelength range so that a glioma is marked by a second fluorescent material. The first and second light sources are selectively controlled by the controller.

[0023] Meanwhile, the light sources may include filters. The first light source may include a first filter, and the second light source may include a second filter.

[0024] Here, the first fluorescent material may include indocyanine green (ICG), and the second fluorescent material may include one of 5-aminolevulinic acid (5-ALA) and protoporphyrin IX (PpIX) converted from 5-ALA. The light sources may include a third light source that emits white light.

[0025] Also, the first wavelength range may be 750 nm to 800 nm, and the second wavelength range may be 350 nm to 450 nm. The probe is formed of a flexible material.

[0026] In addition, the present invention relates to a system for aligning multiple fluorescent images. The system includes: the pen-type medical fluorescent imaging device; a fluorescent multi-image acquirer that acquires an actual image of a target part, a fluorescent vascular image in which blood vessels are marked by the first fluorescent material, and a fluorescent glioma image in which a glioma is marked by the second fluorescent material; a blood vessel extractor that extracts a vascular shape from the acquired fluorescent vascular image; a glioma image processor that processes the acquired fluorescent glioma image so that the boundary of the glioma becomes distinct; a fluorescent multi-image aligner that aligns the extracted fluorescent vascular image and the fluorescent glioma image in real time; and a fluo-

rescent image display that displays a fluorescent multi-image in which the fluorescent vascular image and the fluorescent glioma image are aligned.

[0027] Here, the fluorescent multi-image acquirer includes a fluorescent vascular shape image acquirer that acquires the fluorescent vascular image in which the positions of the blood vessels are marked by the first fluorescent material, and a fluorescent glioma image acquirer that acquires the fluorescent glioma image in which the position of the glioma is marked by the second fluorescent material.

[0028] In addition, the fluorescent multi-image acquirer may acquire transformation parameters corresponding to the highest similarity by measuring pixel value-based similarities of a moving image with respect to the actual image, and match the moving image to the actual image by performing two-dimensional (2D) centered similarity registration based on the acquired transformation parameters so that coordinate errors between the actual image and the moving image are minimized, and the moving image may be one of the fluorescent vascular image and the fluorescent glioma image.

[0029] The fluorescent image display may display at least one of the actual image, the fluorescent vascular image, the fluorescent glioma image, and the fluorescent multi-image, according to a selection of an operator or a user. The fluorescent image display may set individual opacity values of the actual image, the fluorescent vascular image, the fluorescent glioma image, and the fluorescent multi-image to simultaneously display a plurality of images overlapping with each other, and may adjust the opacity values of the individual images to display only a specific image as necessary.

[0030] The pen-type medical fluorescent imaging device may include a transmitter capable of transmitting the actual image, the fluorescent vascular image, and the fluorescent glioma image captured by the image capturing section to the fluorescent multi-image acquirer through wireless communication.

[0031] Hereinafter, exemplary embodiments of the present invention will be described in detail with reference to the accompanying drawings. Terms and words used in the present specification and claims are not to be construed to have general or dictionary meanings, but are to be construed to be meanings and concepts meeting the technical spirit of the present invention based on a principle that the present inventors may appropriately define the concepts of terms to describe their inventions in the best modes.

[0032] Therefore, exemplary embodiments described in this specification and configurations illustrated in the drawings are merely most preferable embodiments, but do not represent all of the technical spirit of the present invention. Consequently, the present invention should be construed as including all changes, equivalents, and substitutions of the embodiments at the time of the filing of this application.

[0033] FIG. 1 is a perspective view of a pen-type medical fluorescent imaging device according to the present invention, FIG. 2 is a front view of the pen-type medical fluorescent imaging device according to the present invention, FIG. 3 is a conceptual diagram for facilitating understanding of overall operation of a system for aligning multiple fluorescent images according to the present invention, and FIG. 4 is a block diagram of the system for aligning multiple fluorescent images according to the present invention. A pen-type medical fluorescent imaging device and a system

for aligning multiple fluorescent images according to the present invention will be described below with reference to FIGS. 1 to 4 and exemplary embodiments.

[0034] The present invention provides a pen-type medical fluorescent imaging device **100** capable of detecting and showing the positions of blood vessels and the position of a glioma at the same time for a patient's safety and ease of surgery.

[0035] As shown in FIGS. 1 and 2, the medical fluorescent imaging device **100** of the present invention is formed as a pen type, and includes the probe **110** extending in a longitudinal direction and an image capturing section **120** and light sources **130** at one end of the probe **110**.

[0036] Here, a pen type denotes a small pen shape that may be conveniently and freely used with a hand like a pen, which is a writing instrument. According to the present invention, the probe **110** may be formed in a pen shape.

[0037] The probe **110** may include the image capturing section **120** and the light sources **130** at the end, and the light sources **130** may be plural in number to surround the image capturing section **120**. Here, the light sources **130** may include a first light source **131** and a second light source **132**. The first light source **131** may emit light of a first wavelength range so that blood vessels are marked by a first fluorescent material, and the second light source **132** may emit light of a second wavelength range so that a glioma is marked by a second fluorescent material. For reference, the light sources **130** may be light-emitting diode (LED) lamps, and may preferably be near-infrared LED lamps. The number of LED lamps that emit light may be adjusted according to the size and shape of a target part.

[0038] The image capturing section **120** of the present invention captures an image of a target part, and may be a general camera for capturing an actual image of a target part, a fluorescent vascular image in which blood vessels are marked by the first fluorescent material, and a fluorescent glioma image in which a glioma is marked by the second fluorescent material.

[0039] In particular, a digital camera or a near infrared ray camera capable of capturing and outputting or storing an image in real time may be used as the image capturing section **120**. For example, a charge coupled device (CCD) or complementary metal-oxide semiconductor (CMOS) camera may be used, and a CCD or CMOS camera having a resolution of megapixel level or greater may preferably be used. Such a camera with megapixel level resolution or greater may be used to acquire a vascular and/or glioma image at high resolution and to observe blood vessels in real time (it is possible to acquire images of 30 frames or more per second). Therefore, convenience for doctors and patients is improved, and it is possible to rapidly and easily observe blood vessels and/or a glioma during a medical examination and surgery of a patient.

[0040] In particular, according to the present invention, the light sources **130** and the image capturing section **120** are installed at a front end of the probe **110** such that loss of light and image quality may be reduced.

[0041] In addition, the fluorescent imaging device **100** of the present invention may include a controller **140** that controls the light sources **130**. The first light source **131** and the second light source **132** may be selectively controlled by the controller **140** to emit light. The controller **140** may be a conventional switch, and may be supplied with current from a power supply and may turn on or off the light sources

130. Meanwhile, the controller **140** that controls the first light source **131** and the second light source **132** may be a first switch **141** and a second switch **142**.

[0042] In the present invention, ICG, 5-ALA, and the like, which are near-infrared fluorescent materials, may be used as fluorescent materials. The first fluorescent material may be ICG, and the second fluorescent material may be 5-ALA.

[0043] In particular, ICG, which is efficient in terms of detecting the positions of a patient's blood vessels and lymph nodes, is a fluorescent material that has been used since 1957 and has few side effects. When injected into a patient, ICG is combined with plasma protein, and when light in a range of 750 nm to 800 nm is transmitted, ICG radiates fluorescent light having a fluorescence peak value of 845 nm.

[0044] 5-ALA, which is efficient in terms of detecting the position of a glioma, has a fluorescence peak value of 635 nm and radiates fluorescent light when light of about 400 nm is transmitted to PpIX converted in cells (it is also possible to use radiated light of about 405 nm). Using this property, it is possible to distinguish between normal tissue and a malignant glioma. The use of 5-ALA increases a success rate of complete removal of a malignant glioma about 1.4 times, and reduces a malignant glioma which is not removed to $\frac{1}{6}$ of its size, and is thus effective in terms of preventing recurrence of the malignant glioma.

[0045] Meanwhile, the medical fluorescent imaging device **100** of the present invention may emit light of various wavelengths through the light sources **130**, and a laser, an LED, and the like may be used as light sources. The light sources **130** may include a filter **150**, which filters an excitation wavelength band at which multiple fluorescent materials distributed in a target body such as a human body are excited and filters a light-emitting wavelength band at which light is emitted from the excited multiple fluorescent materials. The filter **150** may be provided to filter an excitation wavelength band and a light-emitting wavelength band that are different from each other.

[0046] More specifically, the first light source **131** may include a first filter, and the second light source **132** may include a second filter.

[0047] In particular, after ICG is intravenously injected into a patient and 5-ALA is ingested by the patient, light in a range of 750 nm to 800 nm may be transmitted to a treatment area of the patient through the first light source of the medical fluorescent imaging device **100** of the present invention. Then, ICG combined with plasma protein in the target body may radiate fluorescent light having a peak value of 845 nm.

[0048] Also, when the second light source **132** transmits light of 400 nm to the treatment area including PpIX converted from 5-ALA through metabolism in cells, fluorescent light having a peak value of 635 nm may be radiated by PpIX distributed to a glioma.

[0049] In other words, the first wavelength range may be 750 nm to 800 nm, and the second wavelength range may be 350 nm to 450 nm.

[0050] In addition, the medical fluorescent imaging device **100** of the present invention may include a third light source **133** that may radiate white light. The controller **140** for controlling the third light source **133** may be a third switch **143**.

[0051] More specifically, the third light source 133 may be used in diagnosis of a patient by a doctor, and medical personnel may use the third light source 133 as a substitute for a medical pen light.

[0052] Here, a pen light is used by a doctor to diagnose a patient. As a light formed in a pen shape, a pen light enables a user to conveniently carry it around or turn it on in the dark to identify objects.

[0053] The medical fluorescent imaging device 100 of the present invention may include a battery for power supply that is installed so as to be replaceable in the probe 110. In addition, a clip (not shown) for fixing the medical fluorescent imaging device 100 to a doctor's gown may be included.

[0054] The medical fluorescent imaging device 100 having such a configuration including the third light source 133 may be used in a doctor's office, a ward, etc. to emit light on dim areas of a patient's body such as the mouth, nose, ears, etc., that natural light, general lighting, etc. cannot reach and to examine an affected area or used to emit light to an eye and check a pupillary reflex.

[0055] Meanwhile, the probe 110 of the present invention may be formed of a flexible material. In this case, a user may change the position of a total cross section irradiated by the light sources 130 as necessary, and may easily emit light to a part to be examined and capture an image of the part, such that the convenience and accuracy of work are improved. The probe 110 may be formed of a metal or a synthetic resin such as polypropylene (PP), polyethylene (PE), nylon, or the like, and a bellow pipe may be formed as a particular aspect.

[0056] Here, any flexible material fulfilling such a purpose may be used for the probe 110.

[0057] The present invention relates to a system for aligning multiple fluorescent images which makes it possible for an operator to accurately and easily check the positions and connections of a tumor and blood vessels by acquiring multiple fluorescent images in which the positions of the blood vessels and the boundary of a glioma are marked to luminesce due to multiple fluorescent materials, and by aligning and displaying the acquired multiple fluorescent images in real time.

[0058] More specifically, referring to FIGS. 3 and 4, the system for aligning multiple fluorescent images according to the present invention includes the pen-type medical fluorescent imaging device 100, a fluorescent multi-image acquirer 200 that acquires an actual image of a target part, a fluorescent vascular image 310 in which blood vessels are marked by the first fluorescent material, and a fluorescent glioma image 410 in which a glioma is marked by the second fluorescent material, a blood vessel extractor 500 that extracts vascular shapes from the acquired fluorescent vascular image 310, a glioma image processor 600 that processes the acquired fluorescent glioma image 410 so that the boundary of the glioma becomes distinct, a fluorescent multi-image aligner 700 that aligns an extracted vascular image 510 of the blood vessel extractor 500 and a processed glioma image 610 in real time, and a fluorescent image display 800 that displays a fluorescent multi-image in which the extracted vascular image 510 and the processed glioma image 610 are aligned.

[0059] Here, the fluorescent multi-image acquirer 200 includes a fluorescent vascular image acquirer 300 that acquires the fluorescent vascular image 310 in which the positions of blood vessels are marked by the first fluorescent

material, and a fluorescent glioma image acquirer 400 that acquires the fluorescent glioma image 410 in which the position of a glioma is marked by the second fluorescent material.

[0060] Meanwhile, the medical fluorescent imaging device 100 additionally includes a transmitter capable of transmitting the actual image, the vascular image, and the glioma image captured by the image capturing section 120 from the medical fluorescent imaging device 100 to the fluorescent multi-image acquirer 200 through wireless communication.

[0061] The transmitter may transmit images captured by the image capturing section 120 as well as the fluorescent multi-image acquirer 200 to devices capable of monitoring the images such as a computer, a smart phone, and the like. Doctors may check images captured in real time as required by using their smart phones. In this case, images may be transmitted through wireless Internet, Bluetooth, or the like.

[0062] The fluorescent multi-image aligner 700 acquires transformation parameters corresponding to the highest similarity by measuring pixel value-based similarities of moving images with respect to a fixed image to perform 2D centered similarity registration, and matches moving images to the fixed image by using normalized cross-correlation (NCC).

[0063] A fixed image denotes the actual image, and moving images denote the acquired fluorescent vascular image 310 and the acquired fluorescent glioma image 410.

[0064] Meanwhile, the present invention may display one or more of the actual image, the fluorescent vascular image 310, and the fluorescent glioma image 410 captured by the image capturing section 120 and a fluorescent multi-image 810 according to a selection of an operator or a user. The display 800 may simultaneously display a plurality of images to overlap with each other by separately setting the opacity of the actual image, the fluorescent vascular image 310, the fluorescent glioma image 410, and the fluorescent multi-image 810, and may display only specific images as necessary by adjusting an opacity value of each image.

[0065] A method of aligning multiple fluorescent images by using the system for aligning multiple fluorescent images including the pen-type medical fluorescent imaging device 100 according to the present invention will be described in detail below with reference to accompanying drawings.

[0066] First, ICG is intravenously injected into a patient, and 5-ALA is ingested by the patient. After 5-ALA is ingested, the time for a sufficient amount of PpIX to accumulate in tumor tissue is preferably necessary before excited light is emitted to excite PpIX. Specifically, it may take four to eight hours.

[0067] Subsequently, when light in a range of 750 nm to 800 nm is transmitted to a treatment area of the patient through the first light source 131 of the medical fluorescent imaging device 100, ICG combined with plasma protein in the target body may radiate fluorescent light having a peak value of 845 nm. The fluorescent vascular image acquirer 300 may acquire the fluorescent vascular image 310 in which blood vessels radiate fluorescent light of 845 nm due to ICG combined with plasma protein.

[0068] Also, when the second light source 132 transmits light of about 400 nm to the treatment area including PpIX converted from 5-ALA through metabolism in cells, it is possible to acquire the fluorescent glioma image 410 in

which a glioma radiates fluorescent light having a peak value of 635 nm due to PpIX distributed to the glioma.

[0069] Then, vascular positions are marked with fluorescence due to ICG as shown in the fluorescent vascular image 310 of FIG. 3, and also the position of the glioma is marked with fluorescence due to PpIX converted from 5-ALA as shown in the fluorescent glioma image 410.

[0070] Here, the blood vessel extractor 500 may extract vascular shapes from the fluorescent vascular image 310, and the vascular shapes may be extracted from the ICG fluorescent vascular image 310 by using histogram equalization and Otsu thresholding, which is an automatic binarization technique using a histogram.

[0071] The glioma image processor 600 may process the fluorescent glioma image 410 so that the position and the boundary of the glioma become more distinct.

[0072] For reference, the extracted vascular image 510, excluding the blood vessels marked with fluorescence, may be converted into a grey color and then used for image alignment based on feature points (edges, line parts, or the like). Also, the processed glioma image 610, excluding the glioma, may be converted into a grey color and then used for image alignment.

[0073] The fluorescent multi-image aligner 700 may perform image matching on even a fluorescent image that shows morphology slightly different from the color values of the actual image, and may align and match, as a functional image, two or three such images to each other to show different characteristics based on wavelength range. Therefore, it is possible for the operator to see a fluorescent image in which the positions of a glioma and blood vessels are distinct.

[0074] The fluorescent multi-image aligner 700 performs 2D centered similarity registration to minimize coordinate errors between the actual image and an infrared (IR) image (the extracted vascular image 510 or the processed glioma image 610). As shown in FIG. 3, the fluorescent multi-image aligner 700 acquires transformation parameters corresponding to the highest similarity by measuring a pixel value-based similarity of a moving image (the vascular shape image or the fluorescent glioma image) with respect to a fixed image (the actual image) to perform 2D centered similarity registration, and matches the moving image to the fixed image by using NCC.

[0075] The fluorescent multi-image aligner 700 measures a pixel value-based similarity of the fluorescent vascular image 310 with respect to the actual image and continuously changes the fluorescent vascular image 310 until a preset condition is satisfied. When the preset condition is not satisfied within the maximum number of interpolations, transformation parameters corresponding to a highest similarity are acquired.

[0076] A total of six parameters may be acquired through similarity registration: a scale factor, a radian-based angle, x and y values of a center position (x, y) of a moving image after transformation, and translated x and y values (x', y').

[0077] The six transformation parameters acquired through similarity registration are used to transform a vascular image to coincide with the actual image using Equation 1. In other words, similarity conversion parameters for rotation, translation, and uniform scaling of an ICG image (the fluorescent vascular image 310) are calculated with respect to the actual image that is a color image, and the calculated parameters are applied to the ICG image for

resampling. Then, an image matching section matches the ICG image to the actual image.

$$\begin{bmatrix} x' \\ y' \end{bmatrix} = \begin{bmatrix} \lambda & 0 \\ 0 & \lambda \end{bmatrix} \begin{bmatrix} \cos\theta & -\sin\theta \\ \sin\theta & \cos\theta \end{bmatrix} \begin{bmatrix} x - C_x \\ y - C_y \end{bmatrix} + \begin{bmatrix} T_x + C_x \\ T_y + C_y \end{bmatrix} \quad [\text{Equation 1}]$$

[0078] In Equation 1, λ is a scale factor, θ is a rotation angle, (C_x, C_y) are values of a rotation center position, and (T_x, T_y) are values of translated elements.

[0079] After a differential image is generated, similarity is calculated by using NCC as shown in Equation 2. Since the images have different colors, no pixel value-based image is generated. Rather, differential values of the images are used, and then a normalized correlation value is used to perform image matching.

$$R(x, y) = \frac{\sum_{x', y'} (T'(x', y') - I'(x + x', y + y'))}{\sqrt{\sum_{x', y'} T'(x', y')^2} \sqrt{\sum_{x', y'} I'(x + x', y + y')^2}} \quad [\text{Equation 2}]$$

[0080] In Equation 2, $R(x, y)$ denotes a calculated similarity value, T denotes a target image, and I denotes an original image.

[0081] The fluorescent vascular image 310 and the fluorescent glioma image 410 are aligned by the fluorescent multi-image aligner 700 so that different sets of data are transformed into one coordinate system.

[0082] The fluorescent image display 800 displays, to the operator, an image in which both the positions of blood vessels and the boundary of the glioma aligned by the fluorescent multi-image aligner 700 are distinctly marked. The fluorescent image display 800 displays at least one of the actual image of the treatment area, the fluorescent vascular image 310 in which blood vessels are marked to fluoresce due to ICG, the fluorescent glioma image 410 in which the boundary of a glioma is marked to fluoresce based on 5-ALA, and a fluorescent multi-image in which the fluorescent vascular image 310 and the fluorescent glioma image 410 are aligned according to a selection of the operator or a user.

[0083] Also, the fluorescent image display 800 may set an opacity value of each image and simultaneously display a plurality of images overlapping with each other, or may adjust the opacity value of each image and display only a specific image as necessary.

[0084] Due to such operations, it is possible to align, as one image, the fluorescent vascular image 310 in which the positions of blood vessels are marked due to ICG and the fluorescent glioma image 410 in which the boundary of the glioma is marked based on 5-ALA, and it is possible to provide the operator with an image in which both the positions of blood vessels and the boundary of the glioma are distinctly marked.

[0085] Since the pen-type medical fluorescent imaging device of the present invention includes both a first light source and a second light source for showing the positions of both a glioma and blood vessels, it is possible to selectively know the positions of the blood vessels and the glioma.

[0086] Since the pen-type medical fluorescent imaging device of the present invention is miniaturized in the form of a probe, it can be easily held by an operator and carried around. Also, it is possible to reduce loss of light and image quality by installing light sources and an image capturing section at a front end.

[0087] The system for aligning multiple fluorescent images of the present invention makes it possible for an operator to accurately and easily check the positions and connections of a tumor and blood vessels by acquiring multiple fluorescent images in which the positions of the blood vessels and the boundary of a glioma are marked to luminesce due to multiple fluorescent materials having different properties in a target part, and by aligning and displaying the acquired multiple fluorescent images in real time.

[0088] It will be apparent to those skilled in the art that various modifications can be made to the above-described exemplary embodiments of the present invention without departing from the spirit or scope of the invention. Thus, it is intended that the present invention covers all such modifications provided they fall within the scope of the appended claims and their equivalents.

DESCRIPTION OF REFERENCE NUMERALS

- [0089] 100: FLUORESCENT IMAGING DEVICE
- [0090] 120: IMAGE CAPTURING SECTION
- [0091] 130: LIGHT SOURCES
- [0092] 131: FIRST LIGHT SOURCE
- [0093] 132: SECOND LIGHT SOURCE
- [0094] 133: THIRD LIGHT SOURCE
- [0095] 140: CONTROLLER
- [0096] 150: FILTER
- [0097] 300: FLUORESCENT VASCULAR IMAGE ACQUIRER
- [0098] 400: FLUORESCENT GLIOMA IMAGE ACQUIRER
- [0099] 500: BLOOD VESSEL EXTRACTOR
- [0100] 600: GLIOMA IMAGE PROCESSOR
- [0101] 700: FLUORESCENT MULTI-IMAGE ALIGNER
- [0102] 800: FLUORESCENT IMAGE DISPLAY

What is claimed is:

1. A pen-type medical fluorescent imaging device comprising:

a probe configured to extend in a longitudinal direction and include an image capturing section at one end;
 a plurality of light sources configured to be arranged to surround the image capturing section; and
 a controller configured to control the light sources, wherein the light sources include a first light source configured to emit light of a first wavelength range so that blood vessels are marked by a first fluorescent material, and a second light source configured to emit light of a second wavelength range so that a glioma is marked by a second fluorescent material, and the first and second light sources are selectively controlled by the controller.

2. The pen-type medical fluorescent imaging device of claim 1, wherein the light sources include filters, wherein the first light source includes a first filter, and the second light source includes a second filter.

3. The pen-type medical fluorescent imaging device of claim 1, wherein the first fluorescent material includes indocyanine green (ICG), and

the second fluorescent material includes one of 5-aminolevulinic acid (5-ALA) and protoporphyrin IX (PpIX) converted from 5-ALA.

4. The pen-type medical fluorescent imaging device of claim 1, wherein the light sources further include a third light source configured to emit white light.

5. The pen-type medical fluorescent imaging device of claim 1, wherein the first wavelength range is 750 nm to 800 nm, and

the second wavelength range is 350 nm to 450 nm.

6. The pen-type medical fluorescent imaging device of claim 1, wherein the probe is formed of a flexible material.

7. A system for aligning multiple fluorescent images, the system comprising:

the pen-type medical fluorescent imaging device of claim 1

a fluorescent multi-image acquirer configured to acquire an actual image of a target part, a fluorescent vascular image in which blood vessels are marked by the first fluorescent material, and a fluorescent glioma image in which a glioma is marked by the second fluorescent material;

a blood vessel extractor configured to extract a vascular shape from the acquired fluorescent vascular image;

a glioma image processor configured to process the acquired fluorescent glioma image so that a boundary of the glioma becomes distinct;

a fluorescent multi-image aligner configured to align the extracted fluorescent vascular image and the fluorescent glioma image in real time; and

a fluorescent image display configured to display a fluorescent multi-image in which the fluorescent vascular image and the fluorescent glioma image are aligned.

8. The system of claim 7, wherein the fluorescent multi-image acquirer includes:

a fluorescent vascular shape image acquirer configured to acquire the fluorescent vascular image in which positions of the blood vessels are marked by the first fluorescent material; and

a fluorescent glioma image acquirer configured to acquire the fluorescent glioma image in which a position of the glioma is marked by the second fluorescent material.

9. The system of claim 7, wherein the fluorescent multi-image acquirer acquires transformation parameters corresponding to the highest similarity by measuring pixel value-based similarities of a moving image with respect to the actual image, and matches the moving image to the actual image by performing two-dimensional (2D) centered similarity registration based on the acquired transformation parameters so that coordinate errors between the actual image and the moving image are minimized, and

the moving image is one of the fluorescent vascular image and the fluorescent glioma image.

10. The system of claim 7, wherein the fluorescent image display displays at least one of the actual image, the fluorescent vascular image, the fluorescent glioma image, and the fluorescent multi-image according to a selection of an operator or a user.

11. The system of claim 10, wherein the fluorescent image display sets individual opacity values of the actual image, the fluorescent vascular image, the fluorescent glioma

image, and the fluorescent multi-image to simultaneously display a plurality of images overlapping each other, and adjusts the opacity values of the individual images to display only a specific image as necessary.

12. The system of claim 7, wherein the pen-type medical fluorescent imaging device includes a transmitter configured to transmit the actual image, the fluorescent vascular image, and the fluorescent glioma image captured by the image capturing section to the fluorescent multi-image acquirer through wireless communication.

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专利名称(译)	笔型医用荧光成像装置和使用该装置对准多个荧光图像的系统		
公开(公告)号	US20180218508A1	公开(公告)日	2018-08-02
申请号	US15/749871	申请日	2016-09-06
[标]申请(专利权)人(译)	NAT癌症CENT		
申请(专利权)人(译)	国家癌症中心		
当前申请(专利权)人(译)	国家癌症中心		
[标]发明人	LEE SEUNG HOON KIM KWANG GI YOO HEON		
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IPC分类号	G06T7/37 G06T7/12 G06T7/174 A61B5/00 A61K49/00		
CPC分类号	G06T7/37 G06T7/12 G06T7/174 A61B5/0071 A61B5/7425 A61K49/0034 G06T2207/10064 G06T2207/10016 G06T2207/30096 G06T2207/30101 A61B2560/0418 A61B2562/0233 A61B1/043		
优先权	1020150109516 2015-08-03 KR		
外部链接	Espacenet USPTO		

摘要(译)

笔型医用荧光成像装置技术领域本发明涉及一种笔型医用荧光成像装置，包括：探针，其在纵向上是细长的并且在一端具有图像捕获单元；围绕图像捕获单元的多个光源单元；控制单元，用于控制光源单元，其中光源单元包括：第一光源，用于发射第一波长范围的光，使得血管由第一荧光材料标记；用于发射第二波长范围的光的第二光源，使得神经胶质瘤由第二荧光材料标记，其中第一和第二光源由控制单元选择性地控制。

