



(11) **EP 1 272 102 B1**

(12) **EUROPEAN PATENT SPECIFICATION**

(45) Date of publication and mention of the grant of the patent:
11.02.2009 Bulletin 2009/07

(21) Application number: **01929448.7**

(22) Date of filing: **22.03.2001**

(51) Int Cl.:
A61B 5/00 (2006.01) G01N 21/47 (2006.01)

(86) International application number:
PCT/EP2001/003362

(87) International publication number:
WO 2001/074238 (11.10.2001 Gazette 2001/41)

(54) **METHOD AND DEVICE FOR LOCALIZING A DEVIANT REGION IN A TURBID MEDIUM**

VERFAHREN UND VORRICHTUNG ZUR LOKALISIERUNG EINES ABWEICHENDEN BEREICHES IN EINEM TRÜBEN MEDIUM

PROCEDE ET DISPOSITIF PERMETTANT DE LOCALISER UNE REGION DEVIANTE DANS UN MILIEU TROUBLE

(84) Designated Contracting States:
AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE TR

(30) Priority: **31.03.2000 EP 00201164**

(43) Date of publication of application:
08.01.2003 Bulletin 2003/02

(73) Proprietor: **Koninklijke Philips Electronics N.V. 5621 BA Eindhoven (NL)**

(72) Inventors:
• **VAN DER MARK, Martinus, B. NL-5656 AA Eindhoven (NL)**

• **'T HOOFT, Gert, W. NL-5656 AA Eindhoven (NL)**
• **WACHTERS, Arthur, J., H. NL-5656 AA Eindhoven (NL)**

(74) Representative: **Schouten, Marcus Maria Philips Intellectual Property & Standards P.O. Box 220 5600 AE Eindhoven (NL)**

(56) References cited:
WO-A-99/03394 WO-A-99/66832
US-A- 5 719 398

EP 1 272 102 B1

Note: Within nine months of the publication of the mention of the grant of the European patent in the European Patent Bulletin, any person may give notice to the European Patent Office of opposition to that patent, in accordance with the Implementing Regulations. Notice of opposition shall not be deemed to have been filed until the opposition fee has been paid. (Art. 99(1) European Patent Convention).

Description

[0001] The invention relates to a method of localizing a deviant region in a turbid medium that is represented by a set of volume elements, which method includes:

- a first measuring step in which:
 - (a) the turbid medium is irradiated by means of light comprising radiation of mainly of first wavelength;
 - (b) intensities of a part of the light comprising radiation of mainly the first wavelength that is transported along a plurality of light paths through the turbid medium are measured;
- at least a second measuring step in which:
 - (c) the turbid medium is irradiated by means of light comprising radiation of mainly a second wavelength which is not equal to the first wavelength;
 - (d) intensities of a part of the light comprising radiation of mainly the second wavelength that is transported along a plurality of light paths through the turbid medium are measured;
- an imaging step for reconstructing an image of the turbid medium.

[0002] The invention also relates to a device for imaging a deviant region in a turbid medium which is represented by a set of volume elements, which device includes:

- means for irradiating the turbid medium by means of light comprising radiation of mainly a first wavelength;
- means for measuring intensities of a part of the light comprising radiation of mainly the first wavelength that is transported along a plurality of light paths through the turbid medium;
- means for irradiating the turbid medium by means of light comprising radiation of mainly a second wavelength which is not equal to the first wavelength;
- means for measuring intensities of a part of light comprising radiation of mainly the second wavelength that is transported along a plurality of light paths through the turbid medium;
- means for reconstructing an image of the turbid medium from the measured intensities.

[0003] In the context of the present application the term light is to be understood to mean electromagnetic radiation of a wavelength in the visible or infrared range between approximately 400 and 1400 nm. A turbid medium is to be understood to mean a substance consisting of a highly light dispersive material. More specifically, in the

context of the present application the term turbid medium is to be understood to mean biological tissue. A deviant region is to be understood to mean a region in which the turbid medium deviates in any way or form from the turbid medium in the surrounding region. More specifically, in the context of the present application such an area is to be understood to mean a region comprising tumor tissue. The turbid medium is represented by a set of volume elements. Volume elements of this kind are also known as voxels. The size and shape of the volume elements may be the same for all volume elements. However, it is alternatively possible for the volume elements to have mutually different dimensions and shapes.

[0004] A method and a device of this kind are known from "Clinical Optical Tomography and NIR Spectroscopy for Breast Cancer Detection", S.B. Colak et al, IEEE Journal of Selected Topics in Quantum Electronics, Vol. 5, No. 4, July/August 1999. The known method and device are used for imaging the interior of biological tissues. The method and the device can be used inter alia in medical diagnostics for in vivo breast examinations for visual localization of any tumors present in the breast tissue of a human or animal female body. According to the known method a turbid medium is successively irradiated by light from various irradiation positions. Subsequently, the intensity of the light having been transported along different light paths through the turbid medium that extend from their irradiation position is measured in a number of measuring positions. The intensities measured are used for the reconstruction of an image of the turbid medium. A spatial distribution of the attenuation of the light through the tissue is reproduced in this image. Light is attenuated by tissue in that the tissue disperses and absorbs the light.

[0005] A possibly deviant region can be visually localized in the image if the attenuation of the light by the tissue in the deviant region deviates sufficiently from the attenuation of the light by the tissue in the surrounding region. A plurality of individual images can be formed by carrying out the known method repeatedly while using light of different wavelengths.

[0006] The US-patent US 5 719 398 mentions to determine differences of optical parameters obtained at different wavelengths.

[0007] The known method and device have a drawback in that the images thus obtained are insufficiently clear so as to enable accurate visual localization of deviant regions.

[0008] It is an object of the invention to provide a method that yields an image of the turbid medium in which deviant regions can be accurately localized visually.

[0009] This object is achieved by a method of the kind set forth in the preamble, which is characterized in that the imaging step includes the following steps:

- a first calculation step in which a value is assigned to at least a first parameter in dependence on the intensities measured for a volume element in the

measuring steps, said first parameter representing a property of the turbid medium:

- a reconstruction step for representing the value assigned to the first parameter for the volume element in the image of the turbid medium.

[0010] The invention is based on the one hand on the insight that the combining of measured intensities derived from a plurality of measurements with light of different wavelengths can yield values for parameters representing properties of the tissue, which properties enable (separately or in combination), normal tissue to be distinguished from deviant tissue. The parameters thus obtained can represent inter alia components constituting the turbid medium. For example, it is possible for a parameter to represent a spatial distribution of a quantity of water through the turbid medium.

[0011] The invention is also based on the recognition of the fact that regions that are deemed to be deviant on the basis of the parameter values obtained can be marked in the image. The attention of a user, for example a radiologist, is thus drawn to the region in the image that is considered to be deviant.

[0012] The imaging of such parameters yields an image in which a deviant region can be localized more accurately than in an image of the attenuation of the light by the tissue. This is the case inter alia when the parameters represent constituent components of the turbid medium. For example, in an image of the spatial distribution of deoxygenation ratio of blood, $(Hb)/(Hb+HbO_2)$, in female breast tissue a region with tumor tissue will be visually perceptible because, generally speaking, the deoxygenation ratio in tumor tissue is higher than that in normal tissue. Similarly, in an image of the quantity of blood, $(Hb+HbO_2)$, per unit of volume a region with tumor tissue will be visually perceptible, because the blood circulation in tumor tissue generally is greater than that in normal tissue.

[0013] An image can be reconstructed from the parameter values by application of reconstruction techniques that are also used for the reconstruction of an image of the attenuation of the light by tissue from the measured intensities. Examples of such reconstruction techniques are the 3-D Algebraic Reconstruction Technique according to Hoogenraad and the 3-D Conjugent Gradient Technique according to Wachters.

[0014] It is to be noted that the described method is suitable not only for the imaging of biological tissue with a view to detecting deviant regions, but also for the imaging of functional properties of biological tissue. For example, images can be formed of the blood circulation and of the oxygen content of the blood in the brain of infants.

[0015] A version of the method in accordance with the invention is characterized in that the imaging step further comprises the following additional steps:

- a second calculation step in which a significance value is assigned to the volume element in dependence

- on the value of at least the first parameter;
- a display step for displaying the significance value of the corresponding volume element in the image of the turbid medium for a point in the image that corresponds to one of the volume elements.

[0016] The significance value can be represented in the image of the turbid medium, for example by utilizing colors. Distinct colors can thus be assigned to different significance values. Should the significance value vary continuously, alternatively smoothly varying colors can be added to the image. Known color scales, for example the rainbow scale, can be used for this purpose.

[0017] A version of the method in accordance with the invention is characterized in that it includes a third measuring step in which the turbid medium is irradiated by means of light comprising radiation of mainly a third wavelength that is not equal to the first wavelength or the second wavelength and in which intensities of a part of the light comprising radiation of mainly the third wavelength that is transported along a plurality of light paths through the turbid medium are measured, and that in the first calculation step a value is assigned to a second parameter in dependence on the intensities measured for a volume element in the measuring steps, said second parameter representing a property of the turbid medium that is not the same as that represented by the first parameter, and that in the second calculation step a significance value is assigned to the volume element in dependence on the value of the first parameter and on the value of the second parameter.

[0018] Each measuring step in this version produces a set of measured data. The values for two parameters are determined from the three sets of measured data obtained. When the wavelengths of the light are sufficiently far apart in the three measuring steps, the sets of measured data will not be correlated and, using known mathematical techniques, it will in all cases be possible to determine values for the mutually independent parameters from the sets of measured data.

[0019] A version of the method in accordance with the invention is characterized in that the first wavelength has a value of between 830 nm and 900 nm, that the second wavelength has a value of between 750 nm and 830 nm, and that the third wavelength has a value of between 655 nm and 750 nm. Light with the above-mentioned wavelengths can easily be generated by state of the art semiconductor lasers.

[0020] When the wavelengths of the light used in the three measuring steps lie in the above ranges, the spatial distribution of inter alia hemoglobin (Hb) and oxyhemoglobin (HbO_2) through the tissue can be suitably approximated. This is because the attenuation of the light due to dispersion and absorption by biological tissue is low in these regions and because for hemoglobin (Hb) the attenuation of the light as a function of the wavelength of the light deviates from that for oxyhemoglobin (HbO_2).

[0021] A particularly attractive result is obtained when

the first wavelength has a value of approximately 867 nm while the second wavelength has a value of approximately 780 nm and the third wavelength has a value of approximately 715 nm or, as an alternative, a value of approximately 682 nm.

[0022] If an accurate determination of the ratio of fat to water in the tissue is desirable, a fourth measuring step may be added in which the light used contains radiation of a wavelength of between 1040 and 1070 nm or, as an alternative, of between 750 nm and 770 nm. A more accurate determination of the fat to water ratio also enables a more accurate determination of the spatial distribution of hemoglobin (Hb) and oxyhemoglobin (HbO₂) through the tissue.

[0023] An advantageous version of the method in accordance with the invention is characterized in that the first parameter represents a quantity of blood in a volume in the turbid medium that corresponds to one of the volume elements, and that the second parameter represents a deoxygenation ratio of blood in a volume in the turbid medium that corresponds to one of the volume elements.

[0024] In the context of the present invention a quantity of blood is to be understood to mean the sum of the quantities of hemoglobin (Hb) and oxyhemoglobin (HbO₂), while the deoxygenation ratio of blood is to be understood to mean the ratio $(Hb)/(Hb+HbO_2)$.

[0025] Clinical tests performed by applicant have demonstrated that tumor tissue contains an increased quantity of blood per unit of volume and that, moreover, the deoxygenation ratio of the blood in tumor tissue is high in comparison with that in normal tissue. Furthermore, the quantity of blood and the deoxygenation ratio increase as a tumor is more malignant. Therefore, the quantity of blood in a volume and the deoxygenation ratio are suitable parameters, both individually and in combination, for use in the method in accordance with the invention for the to localization of a tumor, if any, in the image.

[0026] A further advantageous version of the method in accordance with the invention is characterized in that the parameters define a parameter space, that in the second calculation step the volume elements are assigned a position in the parameter space in dependence on the values assigned to the parameters for the relevant volume element, and that the significance value assigned to a volume element is determined by the position of the volume element in the parameter space.

[0027] Depending on the number of parameters, such a parameter space may be one-dimensional (a line), two-dimensional (a plane), three-dimensional (a space or volume) or even more dimensional. The parameters then represent the system of co-ordinates of the parameter space. A volume element is assigned a position in said parameter space, the co-ordinates of said position being formed by the values associated with the parameters. For example, in a two-dimensional parameter space a volume element with a value X assigned to a first param-

eter and a value Y assigned to a second parameter will be assigned a position in the parameter space that has the co-ordinate (X, Y).

[0028] Such a parameter space may be virtually present in the device that is used to carry out the method. Moreover, the parameter space can also be shown to the user. For example, a two-dimensional parameter space can be displayed on a display screen.

[0029] A version of the method in accordance with the invention is characterized in that the parameter space is subdivided into at least a first sub-space and a second sub-space, that the volume elements are assigned a first significance value when they occupy a position in the first sub-space and that the volume elements are assigned a second significance value when they occupy a position in the second sub-space.

[0030] The parameter space can be subdivided into sub-spaces by hand as well as automatically. For example, a user can manually indicate a sub-space in a two-dimensional parameter space displayed on a display screen; the volume elements that are situated in said sub-space are then assigned a significance value that is indicative of a deviant region. The user can choose this sub-space, for example, by comparing the distribution of the positions of the volume elements across the parameter space with a reference distribution. During an examination of the breast tissue of a human female body the distribution of the positions of the volume elements across the parameter space that is found for one breast can thus be compared with that found for the other breast.

[0031] The automatic subdivision of the parameter space into sub-spaces can be performed, for example, by subdividing the parameter space into predetermined sub-spaces. Performing a number of reference measurements, thus enables determination in advance as to how the sub-spaces should be distributed across the parameter space and as to what significance value should be assigned to the volume elements in each of the sub-spaces. Alternatively, it is possible to realize the automatic subdivision by means of known segmentation techniques. In that case, for example, the percentage of the number of volume elements that should be situated in a first sub-space and the percentage of the number of volume elements that should be situated in a second sub-space can be indicated in advance.

[0032] Even though the parameter space is subdivided into two sub-spaces in the above description, it will be evident to those skilled in the art that a subdivision into a larger number of sub-spaces is also possible. For each sub-space a distinct significance value can then be assigned to the volume elements situated within each of the sub-spaces. It is also possible to assign for a plurality of sub-spaces the same significance value to the volume elements that are situated in such sub-spaces.

[0033] A version of the method in accordance with the invention is characterized in that the significance value is determined at least by the distance between the position assigned to the volume element and a selected po-

sition in the parameter space.

[0034] For example, the distance between two points in conformity with Euclid's geometric principles can be used as a measure of the distance. For example, the distance between two points $P1(x1, y1)$ and $P2(x2, y2)$ that are situated in a two-dimensional parameter space, where $x1$ represents a first value and $x2$ represents a second value for the parameter X and where $y1$ represents a first value and $y2$ represents a second value for the parameter Y, is equal to .

[0035] For example, the center of gravity of the distribution of the positions of the volume elements in the parameter space can be selected as the selected position. The significance value assigned to a volume element may be a continuously varying value in dependence on the distance. However, the distance may also be subdivided into a number of discrete steps, a significance value being associated with each step.

[0036] A further version of the method in accordance with the invention is characterized in that the significance value is determined inter alia by the situation of the position assigned to the volume element relative to the selected position.

[0037] In addition to the distance from the selected position, the situation of the position assigned to the volume element relative to the selected position is also decisive in respect of the significance value to be assigned to the volume element. For example, a volume element with high parameter values may be assigned a significance value other than the significance value assigned to a volume element that is situated at the same distance from the selected point but has low parameter values.

[0038] It is a further object of the invention to provide a device for producing an image of the turbid medium in which deviant regions can be accurately localized visually.

[0039] This object is achieved by means of a device of the kind set forth in the opening paragraph, which is characterized in that it also includes:

- means for assigning, in dependence on the measured intensities for a volume element, a value to at least a first parameter which represents a property of the turbid medium;
- means for representing the value assigned to the first parameter for the volume element in the image of the turbid medium from at least the value assigned to the first parameter for the volume element.

[0040] A version of the device in accordance with the invention is characterized in that the device further includes:

- means for assigning a significance value to the volume element in dependence on the value of at least the first parameter;
- means for displaying the significance value of the corresponding volume element in the image of the

turbid medium for each point in the image that corresponds to one of the volume elements.

[0041] The devices mentioned in the previous paragraphs are, for example, implemented in the form of a medical diagnostic workstation.

[0042] The invention further relates to a computer program for carrying out the methods according to the invention. This computer program may comprise instructions for carrying out all of the versions of the methods according to the invention or, alternatively, may comprise instructions for carrying out a sub-set of the versions of the methods. The instructions of this computer program are, for example, loaded into the memory of a medical diagnostic workstation and are subsequently executed by a processor in this workstation. According to a further object of the invention, the computer program is recorded on a record carrier, such as for example a CD-ROM disc or a magnetic tape. The instructions of the computer program are transferred from the record carrier into the memory of, for example, a medical diagnostic workstation. Alternatively, the instructions of the computer program are transferred from the record carrier into the memory of a PC or any other standard computer system. It is also possible to load the computer program into a memory through a network such as, for example, the "World-Wide Web".

[0043] The above and other, more detailed aspects of the invention will be described in detail hereinafter, by way of example, with reference to the drawings. Therein:

Fig. 1 illustrates diagrammatically a first method in accordance with the invention.

Fig. 2 shows an example of a two-dimensional parameter space, and

Fig. 3 illustrates diagrammatically a second method in accordance with the invention.

[0044] Fig. 1 shows diagrammatically a first method in accordance with the invention. During a first measuring step 1 the turbid medium is irradiated by means of light-comprising radiation of mainly a wavelength of 867 nm and the intensity of a part of the light that is transported along a plurality of light paths through the turbid medium is measured. Such a measuring step is described in greater detail in European patent application EP-A-0 925 017. Subsequently, a second, similar measuring step 2 is performed by means of light comprising radiation of mainly a wavelength of 780 nm, followed by a third measuring step 3 while using light comprising radiation of mainly a wavelength of 715 nm.

[0045] Using a reconstruction algorithm 5 in an imaging step 4, an image 9 of the turbid medium is reconstructed from the intensities measured in the third measuring step 3. Even though an image is reconstructed from the intensities measured in the third measuring step 3 in the present example, the intensities measured in the first measuring step 1 or the second measuring step 2 can

also be used for this purpose.

[0046] In a first calculation step 6 for each volume element two parameters are determined from the three sets of intensities measured in the three measuring steps. These parameters, representing local characteristics of the tissue examined, are the quantity of blood in a volume and the deoxygenation ratio of the blood in a volume in the present example. In this context a quantity of blood in a volume is to be understood to mean the sum of the hemoglobin and the oxyhemoglobin ($\text{Hb} + \text{HbO}_2$) present in the volume. The term deoxygenation ratio of the blood in a volume is to be understood to mean herein the ratio of the hemoglobin present in the volume to the quantity of blood present in a volume ($\text{Hb}/(\text{Hb} + \text{HbO}_2)$). These parameters are particularly suitable for the localization of tumor tissue in breast tissue, because tumors are generally characterized by a higher blood circulation and a higher oxygen consumption, leading to an increase of both parameters.

[0047] In a second calculation step 7 the volume elements are mapped in a two-dimensional parameter space 20, an example of which is shown in Fig. 2. This parameter space 20 is defined by the two parameters determined in the first calculation step 6. In the present example the quantity of blood in a volume is represented by the horizontal X axis 22 and the deoxygenation ratio of the blood in a volume is represented by the vertical Y axis 21. A volume element in this parameter space is assigned a position 23 which is dependent on the values of the two parameters for the relevant volume element. Thus, a volume element having a value X_1 for the quantity of blood in a volume and a value Y_1 for the deoxygenation ratio of the blood in a volume will be assigned a position having the co-ordinate (X_1, Y_1) in the parameter space.

[0048] Subsequently, each volume element is assigned a significance value which is dependent on the position 23 of the volume element in the parameter space 20. In the present example a volume element is assigned a significance value "1" if it is deemed to be situated in a deviant region and a significance value "0" if it is deemed to be situated in a region outside the deviant region. Fig. 2A shows a sub-region 24 that contains high values for both parameters and is indicative of tumor tissue as discussed above. Volume elements having a position 25 in the sub-region 24 are assigned a significance value "1". The other volume elements are assigned a significance value "0". Even though only two distinct significance values are used in the present example, a plurality of discrete significance values can be used when the parameter space is subdivided into a plurality of sub-regions. It is also possible to use a continuously varying significance value. The significance value then represents, for example a probability value.

[0049] Subsequently, the significance values assigned to the volume elements in the image 9 of the turbid medium are reproduced in the display step 8. A dot in the image 9 that corresponds to a volume element is assigned, for example, a red color if the significance value

assigned to the relevant volume element is "1" and a grey color if the significance value assigned to the relevant volume element is "0". Thus, an image 9 is formed of the turbid medium which contains grey values and a region 10 with red values. The region 10 corresponds to a deviant region in the turbid medium. It will be evident to those skilled in the art that other colors and other methods can also be used to mark a region in an image.

[0050] Fig. 2 shows an example of a two-dimensional parameter space 20. The parameter space 20 is defined by two parameters. A first parameter is represented by the horizontal X axis 22 and a second parameter by the vertical Y axis 21. A volume element in this parameter space is assigned a position 23 which is dependent on the values of the two parameters for the relevant volume element. Thus, a volume element having a value X_1 for the first parameter and a value Y_1 for the second parameter will be assigned a position in the parameter space that has the co-ordinate (X_1, Y_1) .

[0051] In Fig. 2A the parameter space is subdivided into two sub-regions by manual formation of a sub-region 24. The volume elements 25 that are situated in the sub-region 24 were assigned a significance value that deviates from that assigned to the volume elements situated outside the sub-region 24. During the imaging of breast tissue the position and the shape of the sub-region 24 can be selected by comparing a parameter space obtained from measurements performed on one breast with a parameter space obtained from measurements performed on the other breast.

[0052] Fig. 2B shows an alternative method of assigning significance values to the volume elements. A volume element is now assigned a significance value that is dependent on the distance 27 between the position 30 of the volume element in the parameter space 20 and a selected position 26. In the present example the selected position 26 is the geometrical center of gravity of the distribution of the positions of the volume elements in the parameter space 20. The significance value to be assigned may be a continuous function of the distance 27. The distance 27 may alternatively be subdivided into a number of discrete steps, each step having an associated significance value.

[0053] Even though the position 30 of a first volume element and the position 31 of a second volume element are situated at the same distance from the selected position 26, a different significance value can still be assigned to the two volume elements as an alternative. The orientation of the positions of the volume elements relative to the selected point 26 is then also important. For example, the volume element with the position 30 that is situated in the upper right-hand quadrant 28 of the parameter space 20 will be assigned a significance value which differs from that assigned to the volume element with the position 31 that is situated in the lower left-hand quadrant 29 of the parameter space 20. This makes sense, for example, if the quantity of blood in a volume and the deoxygenation ratio of the blood in a volume are

chosen as the parameters. As has already been described, an increased value of both parameters is indicative of tumor tissue. This is the case for the volume elements situated in the upper right-hand quadrant 28; therefore, these volume elements should be assigned a high significance value. However, a reduced value of one or both parameters has no significance. Therefore, the volume elements that are situated in one of the other three quadrants should not be assigned a high significance value on the basis of these two parameters.

[0054] Fig. 3 illustrates diagrammatically a second method in accordance with the invention. During a first measuring step 1 the turbid medium is irradiated by means of light-comprising radiation of mainly a wavelength of 867 nm and the intensity of a part of the light that is transported along a plurality of light paths through the turbid medium is measured. Subsequently, a second, similar measuring step 2 is performed while using light comprising radiation of mainly a wavelength of 780 nm and also a third measuring step 3 that utilizes light comprising radiation of mainly a wavelength of 715 nm.

[0055] In a first calculation step 6 for each volume element two parameters are determined from the three sets of intensities measured in the three measuring steps. Subsequently, in a reconstruction step 13 the value of the first parameter for a volume element is imaged in an image 11 and the value of the second parameter is imaged in an image 12. A point in the image 11 and a point in the image 12 then correspond to the volume element. This yields an image of the spatial distribution of a parameter across the turbid medium.

[0056] When the parameters are suitably chosen, a deviant region will be clearly visible in the images. For example, when the quantity of blood in a volume is imaged in the image 11 and the deoxygenation ratio of the blood in a volume is imaged in the image 12, a region with tumor tissue that is present in the breast tissue will be visible because both parameters have an increased value in the tumor tissue.

[0057] The two parameters are separately imaged in the present example. However, it is also possible to image only one of the two parameters. It is furthermore possible to image more than two parameters separately if more than two parameters are determined in the first calculation step 6. Furthermore, the images of a plurality of parameters can be combined so as to form one common image.

Claims

1. A method of localizing a deviant region in a turbid medium that is represented by a set of volume elements, which method includes:

- a first measuring step (1) in which:

(a) the turbid medium is irradiated by means

of light comprising radiation of mainly of first wavelength;

(b) intensities of a part of the light comprising radiation of mainly the first wavelength that is transported along a plurality of light paths through the turbid medium are measured;

- at least a second measuring step (2) in which:

(c) the turbid medium is irradiated by means of light comprising radiation of mainly a second wavelength which is not equal to the first wavelength;

(d) intensities of a part of the light comprising radiation of mainly the second wavelength that is transported along a plurality of light paths through the turbid medium are measured;

- an imaging step (4) for reconstructing an image of the turbid medium;

characterized in that

the imaging step (4) includes the following steps:

- a first calculation step (6) in which a value is assigned to at least a first parameter in dependence on the intensities measured for a volume element in the measuring steps, said first parameter representing a property of the turbid medium;

- a reconstruction step (13) for representing the value assigned to the first parameter for the volume element in the image (11) of the turbid medium.

2. A method as claimed in claim 1, wherein the imaging step (4) further comprises the following additional step:

- a second calculation step (7) in which a significance value is assigned to the volume element in dependence on the value of at least the first parameter;

- a display step (8) for displaying the significance value (10) of the corresponding volume element in the image of the turbid medium (9) for a point in the image that corresponds to one of the volume elements.

3. A method as claimed in claim 1-2, **characterized in that** the method includes a third measuring step (3) in which the turbid medium is irradiated by means of light comprising radiation of mainly a third wavelength that is not equal to the first wavelength or the second wavelength, and in which intensities of a part of the light comprising radiation of mainly the third wavelength that is transported along a plurality of light paths through

the turbid medium are measured,
and that in the first calculation step (6) a value is assigned to a second parameter in dependence on the intensities measured for a volume element in the measuring steps, said second parameter representing a property of the turbid medium that is not the same as that represented by the first parameter,
and that in the second calculation step (7) a significance value is assigned to the volume element in dependence on the value of the first parameter and on the value of the second parameter.

4. A method as claimed in claim 3, **characterized in that**

the first wavelength has a value of between 830 nm and 900 nm, that the second wavelength has a value of between 750 nm and 830 nm, and that the third wavelength has a value of between 655 nm and 750 nm.

5. A method as claimed in claim 4, **characterized in that**

the first parameter represents a quantity of blood in a volume in the turbid medium that corresponds to one of the volume elements,
and that the second parameter represents a deoxygenation ratio of blood in a volume in the turbid medium that corresponds to one of the volume elements.

6. A method as claimed in claim 1-2, **characterized in that**

the parameters define a parameter space (20), that in the second calculation step (7) the volume elements are assigned a position (23) in the parameter space (20) in dependence on the values assigned to the parameters for the relevant volume element, and that the significance value assigned to a volume element is determined by the position of the volume element in the parameter space.

7. A method as claimed in claim 6, **characterized in that**

the parameter space (20) is subdivided into at least a first sub-space (24) and a second sub-space, that the volume elements are assigned a first significance value when they occupy a position in the first sub-space,
and that the volume elements are assigned a second significance value when they occupy a position in the second sub-space.

8. A method as claimed in claim 6, **characterized in that**

the significance value is determined at least by the distance between the position (23) assigned to the volume element and a selected position (26) in the parameter space.

9. A method as claimed in claim 8, **characterized in that**

the significance value is determined inter alia by the situation of the position assigned to the volume element relative to the selected position.

10. A device for imaging a deviant region in a turbid medium which is represented by a set of volume elements, which device includes:

- means for irradiating the turbid medium by means of light comprising radiation of mainly a first wavelength;
- means for measuring intensities of a part of the light comprising radiation of mainly the first wavelength that is transported along a plurality of light paths through the turbid medium;
- means for irradiating the turbid medium by means of light comprising radiation of mainly a second wavelength which is not equal to the first wavelength;
- means for measuring intensities of a part of light comprising radiation of mainly the second wavelength that is transported along a plurality of light paths through the turbid medium;
- means for reconstructing an image of the turbid medium from the measured intensities

characterized in that

the device also includes:

- means for assigning, in dependence on the measured intensities for a volume element, a value to at least a first parameter which represents a property of the turbid medium;
- means for representing the value assigned to the first parameter for the volume element in the image of the turbid medium from at least the value assigned to the first parameter for the volume element.

11. A device as claimed in claim 10, wherein the device further includes:

- means for assigning a significance value to the volume element in dependence on the value of at least the first parameter;
- means for displaying the significance value of the corresponding volume element in the image of the turbid medium for each point in the image that corresponds to one of the volume elements.

12. Computer program comprising a set of instructions for carrying out the method according to claim 1 or 9.

13. Record carrier comprising the computer program according to claim 12.

Patentansprüche

1. Verfahren zur Lokalisierung eines abweichenden Bereiches in einem trüben Medium, das durch einen Satz Volumenelemente dargestellt wird, wobei das Verfahren Folgendes umfasst:
- einen ersten Messschritt (1), in dem:
 - (a) das trübe Medium mit Licht bestrahlt wird, das Strahlung von hauptsächlich einer ersten Wellenlänge enthält;
 - (b) Intensitäten eines Teils des Strahlung von hauptsächlich der ersten Wellenlänge enthaltenden Lichts gemessen werden, das auf einer Vielzahl von Lichtpfaden durch das trübe Medium transportiert wird;
 - mindestens einen zweiten Messschritt (2), in dem:
 - (c) das trübe Medium mit Licht bestrahlt wird, das Strahlung von hauptsächlich einer zweiten Wellenlänge enthält, die ungleich der ersten Wellenlänge ist;
 - (d) Intensitäten eines Teils des Strahlung von hauptsächlich der zweiten Wellenlänge enthaltenden Lichts gemessen werden, das auf einer Vielzahl von Lichtpfaden durch das trübe Medium transportiert wird;
 - einen Bildgebungsschritt (4) zum Rekonstruieren eines Bildes des trüben Mediums; **dadurch gekennzeichnet, dass** der Bildgebungsschritt (4) die folgenden Schritte umfasst:
 - einen ersten Berechnungsschritt (6), in dem mindestens einem ersten Parameter ein Wert in Abhängigkeit von den für ein Volumenelement in den Messschritten gemessenen Intensitäten zugewiesen wird, wobei der genannte erste Parameter eine Eigenschaft des trüben Mediums darstellt;
 - einen Rekonstruktionsschritt (13) zum Darstellen des dem ersten Parameter für das Volumenelement zugewiesenen Wertes in dem Bild (11) des trüben Mediums.
2. Verfahren nach Anspruch 1, wobei der Bildgebungsschritt (4) weiterhin die folgenden zusätzlichen Schritte umfasst:
- einen zweiten Berechnungsschritt (7), in dem dem Volumenelement in Abhängigkeit von dem Wert von mindestens dem ersten Parameter ein Signifikanzwert zugewiesen wird;
 - einen Anzeigeschritt (8) zum Anzeigen des Signifikanzwertes (10) des entsprechenden Volumenelements in dem Bild des trüben Mediums (9) für einen Punkt im Bild, der einem der Volumenelemente entspricht.
3. Verfahren nach Anspruch 1 bis 2, **dadurch gekennzeichnet, dass** das Verfahren einen dritten Messschritt (3) umfasst, in dem das trübe Medium mit Licht bestrahlt wird, das Strahlung von hauptsächlich einer dritten Wellenlänge enthält, die ungleich der ersten Wellenlänge oder der zweiten Wellenlänge ist, und in dem Intensitäten eines Teils des Strahlung von hauptsächlich der dritten Wellenlänge enthaltenden Lichts gemessen werden, das auf einer Vielzahl von Lichtpfaden durch das trübe Medium transportiert wird, und dass im ersten Berechnungsschritt (6) einem zweiten Parameter ein Wert in Abhängigkeit von den für ein Volumenelement in den Messschritten gemessenen Intensitäten zugewiesen wird, wobei der genannte zweite Parameter eine Eigenschaft des trüben Mediums darstellt, die nicht die gleiche ist wie durch den ersten Parameter dargestellt, und dass in dem zweiten Berechnungsschritt (7) dem Volumenelement in Abhängigkeit von dem Wert des ersten Parameters und von dem Wert des zweiten Parameters ein Signifikanzwert zugewiesen wird.
4. Verfahren nach Anspruch 3, **dadurch gekennzeichnet, dass** die erste Wellenlänge einen Wert zwischen 830 nm und 900 nm hat, dass die zweite Wellenlänge einen Wert zwischen 750 nm und 830 nm hat und dass die dritte Wellenlänge einen Wert zwischen 655 nm und 750 nm hat.
5. Verfahren nach Anspruch 4, **dadurch gekennzeichnet, dass** der erste Parameter eine Blutmenge in einem Volumen im trüben Medium darstellt, das einem der Volumenelemente entspricht, und dass der zweite Parameter das Desoxygenationsverhältnis des Bluts in einem Volumen im trüben Medium darstellt, das einem der Volumenelemente entspricht.
6. Verfahren nach Anspruch 1 bis 2, **dadurch gekennzeichnet, dass** die Parameter einen Parameterraum (20) definieren, dass in dem zweiten Berechnungsschritt (7) den Volumenelementen in Abhängigkeit von den Werten, die den Parametern für das relevante Volumenelement zugewiesen wurden, eine Position (23) im Parameterraum (20) zugewiesen wird, und dass der einem Volumenelement zugewiesene Signifikanzwert durch die Position des Volumenelements im Parameterraum bestimmt wird.
7. Verfahren nach Anspruch 6, **dadurch gekennzeichnet,**

- zeichnet, dass** der Parameterraum (20) in mindestens einen ersten Teilraum (24) und einen zweiten Teilraum unterteilt ist, dass den Volumenelementen ein erster Signifikanzwert zugewiesen wird, wenn sie eine Position im ersten Teilraum einnehmen, und dass den Volumenelementen ein zweiter Signifikanzwert zugewiesen wird, wenn sie eine Position im zweiten Teilraum einnehmen.
8. Verfahren nach Anspruch 6, **dadurch gekennzeichnet, dass** der Signifikanzwert mindestens durch den Abstand zwischen der dem Volumenelement zugewiesenen Position (23) und einer ausgewählten Position (26) im Parameterraum bestimmt wird.
9. Verfahren nach Anspruch 8, **dadurch gekennzeichnet, dass** der Signifikanzwert unter anderem durch die Lage der dem Volumenelement zugewiesenen Position relativ zu der ausgewählten Position bestimmt wird.
10. Vorrichtung zur Bildgebung eines abweichenden Bereiches in einem trüben Medium, das durch einen Satz Volumenelemente dargestellt wird, wobei die Vorrichtung Folgendes umfasst:
- Mittel zum Bestrahlen des trüben Mediums mit Licht, das Strahlung von hauptsächlich einer ersten Wellenlänge enthält;
 - Mittel zum Messen der Intensitäten eines Teils des Strahlung von hauptsächlich der ersten Wellenlänge enthaltenden Lichts, das auf einer Vielzahl von Lichtpfaden durch das trübe Medium transportiert wird;
 - Mittel zum Bestrahlen des trüben Mediums mit Licht, das Strahlung von hauptsächlich einer zweiten Wellenlänge enthält, die ungleich der ersten Wellenlänge ist;
 - Mittel zum Messen der Intensitäten eines Teils des Strahlung von hauptsächlich der zweiten Wellenlänge enthaltenden Lichts, das auf einer Vielzahl von Lichtpfaden durch das trübe Medium transportiert wird;
 - Mittel zum Rekonstruieren eines Bildes des trüben Mediums anhand der gemessenen Intensitäten
- dadurch gekennzeichnet, dass** die Vorrichtung außerdem Folgendes umfasst:
- Mittel, um in Abhängigkeit von den für ein Volumenelement gemessenen Intensitäten mindestens einem ersten Parameter einen Wert zuzuweisen, wobei der genannte erste Parameter eine Eigenschaft des trüben Mediums darstellt;
 - Mittel, um den dem ersten Parameter für das Volumenelement zugewiesenen Wert in dem Bild des trüben Mediums darzustellen, ausgehend von mindestens dem Wert, der dem ersten
- Parameter für das Volumenelement zugewiesen wurde.
11. Vorrichtung nach Anspruch 10, wobei die Vorrichtung weiterhin Folgendes umfasst:
- Mittel, um dem Volumenelement in Abhängigkeit von dem Wert von mindestens dem ersten Parameter einen Signifikanzwert zuzuweisen;
 - Mittel, um den Signifikanzwert des entsprechenden Volumenelements in dem Bild des trüben Mediums für jeden Punkt im Bild darzustellen, der einem der Volumenelemente entspricht.
12. Computerprogramm mit einem Anweisungssatz zum Ausführen des Verfahrens nach Anspruch 1 oder 9.
13. Aufzeichnungsträger mit dem Computerprogramm nach Anspruch 12.

Revendications

1. Procédé de localisation d'une région déviante dans un milieu trouble qui est représenté par un ensemble d'éléments de volume, lequel procédé comprend :
- une première étape de mesure (1) dans laquelle :
 - (a) le milieu trouble est irradié au moyen de lumière comprenant du rayonnement de principalement une première longueur d'onde ;
 - (b) il est mesuré des intensités d'une partie de la lumière comprenant du rayonnement de principalement la première longueur d'onde qui est transporté le long d'une pluralité de trajets de lumière à travers le milieu trouble ;
 - au moins une deuxième étape de mesure (2) dans laquelle :
 - (c) le milieu trouble est irradié au moyen de lumière comprenant du rayonnement de principalement une deuxième longueur d'onde qui n'est pas égale à la première longueur d'onde ;
 - (d) il est mesuré des intensités d'une partie de la lumière comprenant du rayonnement de principalement la deuxième longueur d'onde qui est transporté le long d'une pluralité de trajets de lumière à travers le milieu trouble ;
 - une étape d'imagerie (4) pour reconstruire une

- image du milieu trouble ;
caractérisé en ce que :
 l'étape d'imagerie (4) comprend les étapes suivantes :
- une première étape de calcul (6) dans laquelle une valeur est assignée à au moins un premier paramètre en fonction des intensités qui sont mesurées pour un élément de volume dans les étapes de mesure, ledit premier paramètre représentant une propriété du milieu trouble ;
 - une étape de reconstruction (13) pour représenter la valeur qui est assignée au premier paramètre pour l'élément de volume dans l'image (11) du milieu trouble.
2. Procédé selon la revendication 1, dans lequel l'étape d'imagerie (4) comprend encore l'étape additionnelle suivante :
- une deuxième étape de calcul (7) dans laquelle une valeur de signification est assignée à l'élément de volume en fonction de la valeur d'au moins le premier paramètre ;
 - une étape d'affichage (8) pour afficher la valeur de signification (10) de l'élément de volume correspondant dans l'image du milieu trouble (9) pour un point dans l'image qui correspond à un des éléments de volume.
3. Procédé selon les revendications précédentes 1 à 2, **caractérisé en ce que** :
- le procédé comprend une troisième étape de mesure (3) dans laquelle le milieu trouble est irradié au moyen de lumière comprenant du rayonnement de principalement une troisième longueur d'onde qui n'est pas égale à la première longueur d'onde ou à la deuxième longueur d'onde, et dans lequel il est mesuré des intensités d'une partie de la lumière comprenant du rayonnement de principalement la troisième longueur d'onde qui est transporté le long d'une pluralité de trajets de lumière à travers le milieu trouble ; et **en ce que** dans la première étape de calcul (6) une valeur est assignée à un deuxième paramètre en fonction des intensités qui sont mesurées pour un élément de volume dans les étapes de mesure, ledit deuxième paramètre représentant une propriété du milieu trouble qui n'est pas la même que celle qui est représentée par le premier paramètre ; et **en ce que** dans la deuxième étape de calcul (7) une valeur de signification est assignée à l'élément de volume en fonction de la valeur du premier paramètre et de la valeur du deuxième paramètre.
4. Procédé selon la revendication 3, **caractérisé en ce que** :
- la première longueur d'onde présente une valeur dans la gamme comprise entre 830 nm et 900 nm, **en ce que** la deuxième longueur d'onde présente une valeur dans la gamme comprise entre 750 nm et 830 nm et **en ce que** la troisième longueur d'onde présente une valeur dans la gamme comprise entre 655 nm et 750 nm.
5. Procédé selon la revendication 4, **caractérisé en ce que** :
- le premier paramètre représente une quantité de sang dans un volume dans le milieu trouble qui correspond à un des éléments de volume, et **en ce que** le deuxième paramètre représente un rapport de désoxygénation de sang dans un volume dans le milieu trouble qui correspond à un des éléments de volume.
6. Procédé selon les revendications précédentes 1 à 2, **caractérisé en ce que** : les paramètres définissent un espace de paramètre (20), **en ce que** dans la deuxième étape de calcul (7) une position (23) dans l'espace de paramètre (20) est assignée aux éléments de volume en fonction des valeurs qui sont assignées aux paramètres pour l'élément de volume concerné, et **en ce que** la valeur de signification qui est assignée à un élément de volume est déterminée par la position de l'élément de volume dans l'espace de paramètre.
7. Procédé selon la revendication 6, **caractérisé en ce que** :
- l'espace de paramètre (20) est subdivisé en au moins un premier sous-espace (24) et en un deuxième sous-espace, **en ce qu'**une première valeur de signification est assignée aux éléments de volume lorsqu'ils occupent une position dans le premier sous-espace, et **en ce qu'**une deuxième valeur de signification est assignée aux éléments de volume lorsqu'ils occupent une position dans le deuxième sous-espace.
8. Procédé selon la revendication 6, **caractérisé en ce que** :
- la valeur de signification est déterminée au moins par la distance comprise entre la position (23) qui est assignée à l'élément de volume et une position sélectionnée (26) dans l'espace de paramètre.
9. Procédé selon la revendication 8, **caractérisé en**

ce que :

la valeur de signification est déterminée, entre autres, par la situation de la position qui est assignée, entre autres, par la situation de la position qui est assignée à l'élément de volume par rapport à la position sélectionnée.

10. Dispositif d'imagerie d'une région déviante dans un milieu trouble qui est représenté par un ensemble d'éléments de volume, lequel dispositif comprend :

- des moyens pour irradier le milieu trouble au moyen de lumière comprenant du rayonnement de principalement une première longueur d'onde ;
- des moyens pour mesurer des intensités d'une partie de la lumière comprenant du rayonnement de principalement la première longueur d'onde qui est transporté le long d'une pluralité de trajets de lumière à travers le milieu trouble ;
- des moyens pour irradier le milieu trouble au moyen de lumière comprenant du rayonnement de principalement une deuxième longueur d'onde qui n'est pas égale à la première longueur d'onde ;
- des moyens pour mesurer des intensités d'une partie de la lumière comprenant du rayonnement de principalement la deuxième longueur d'onde qui est transporté le long d'une pluralité de trajets de lumière à travers le milieu trouble ;
- des moyens pour reconstruire une image du milieu trouble à partir des intensités mesurées ;

caractérisé en ce que le dispositif comprend également :

- des moyens pour assigner, en fonction des intensités mesurées pour un élément de volume, une valeur à au moins un premier paramètre qui représente une propriété du milieu trouble ;
- des moyens pour représenter la valeur qui est assignée au premier paramètre pour l'élément de volume dans l'image du milieu trouble à partir d'au moins la valeur qui est assignée au premier paramètre pour l'élément de volume.

11. Dispositif selon la revendication 10, dans lequel le dispositif comprend encore :

- des moyens pour assigner une valeur de signification à l'élément de volume en fonction de la valeur d'au moins le premier paramètre ;
- des moyens pour afficher la valeur de signification de l'élément de volume correspondant dans l'image du milieu trouble pour chaque point dans l'image qui correspond à un des éléments de volume.

12. Programme informatique comprenant un ensemble d'instructions pour la mise en oeuvre du procédé selon la revendication 1 ou selon la revendication 9.

13. Support d'enregistrement comprenant le programme informatique selon la revendication 12.

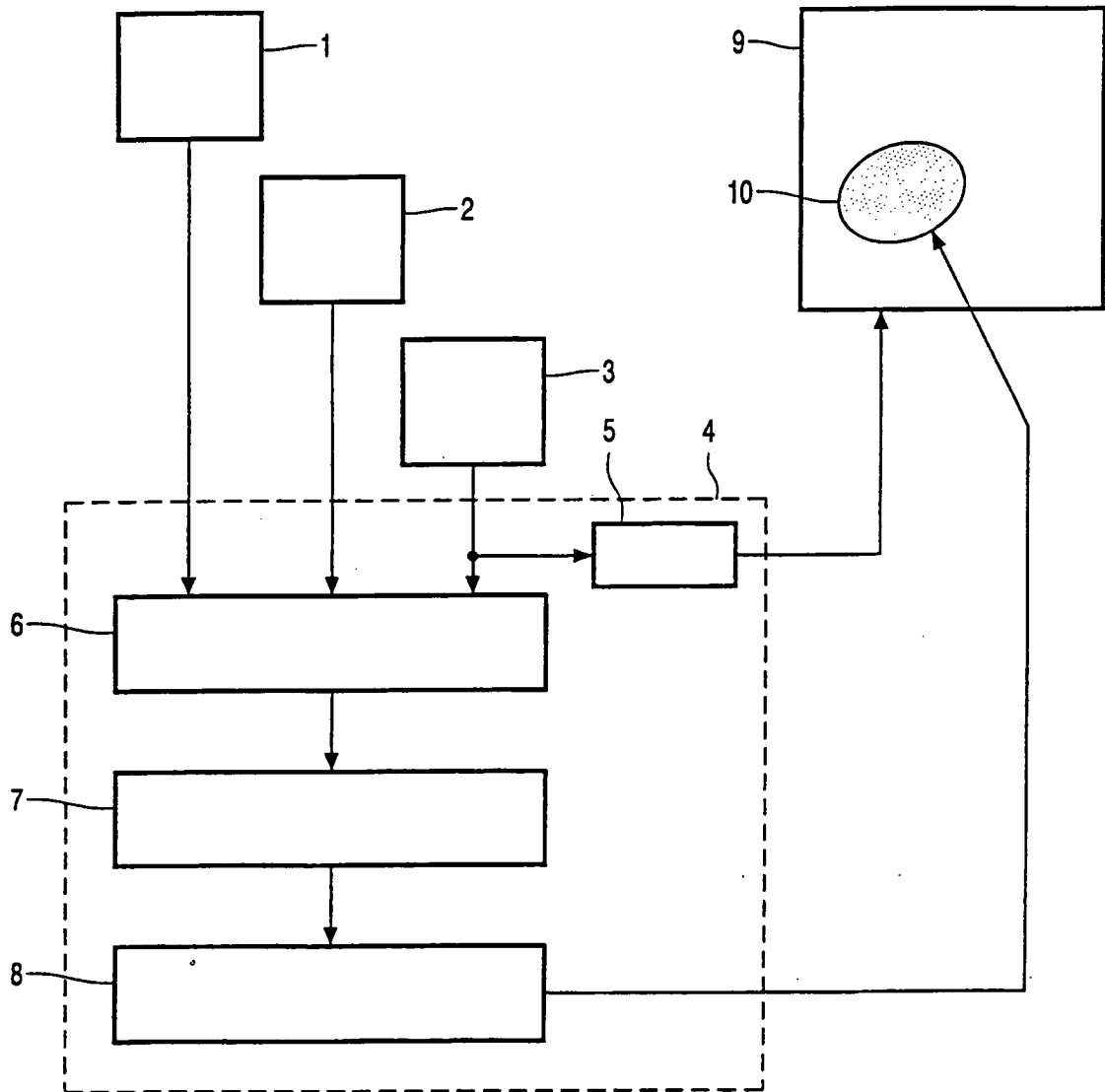


FIG. 1

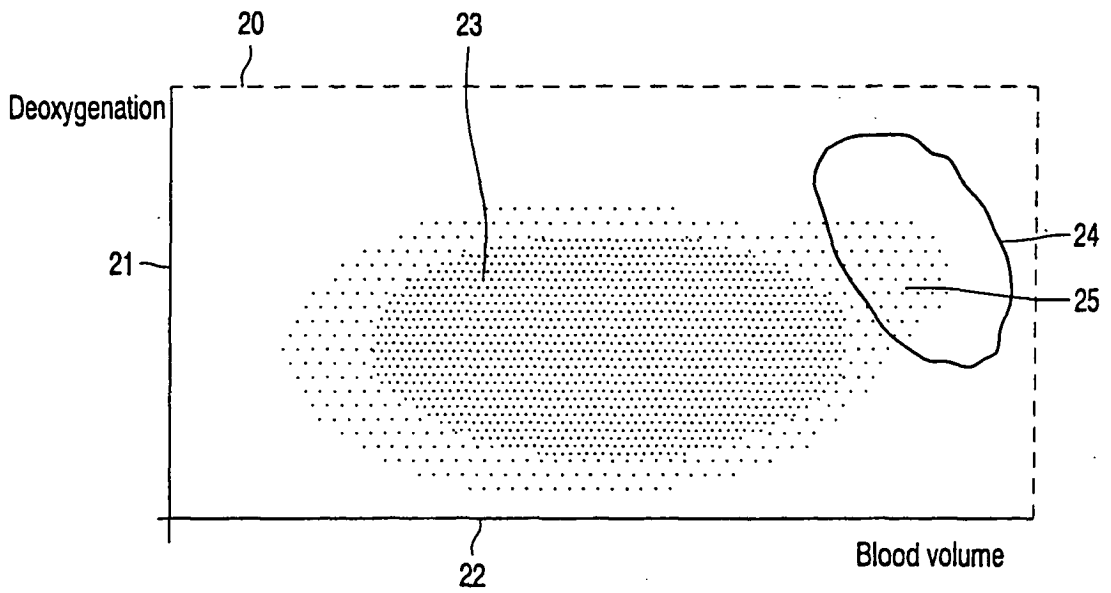


FIG. 2A

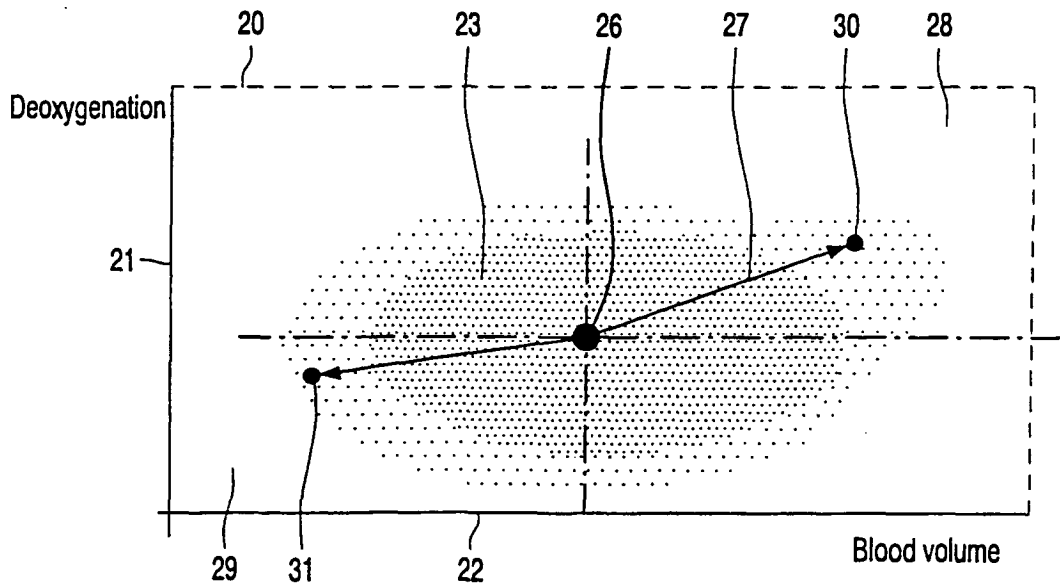


FIG. 2B

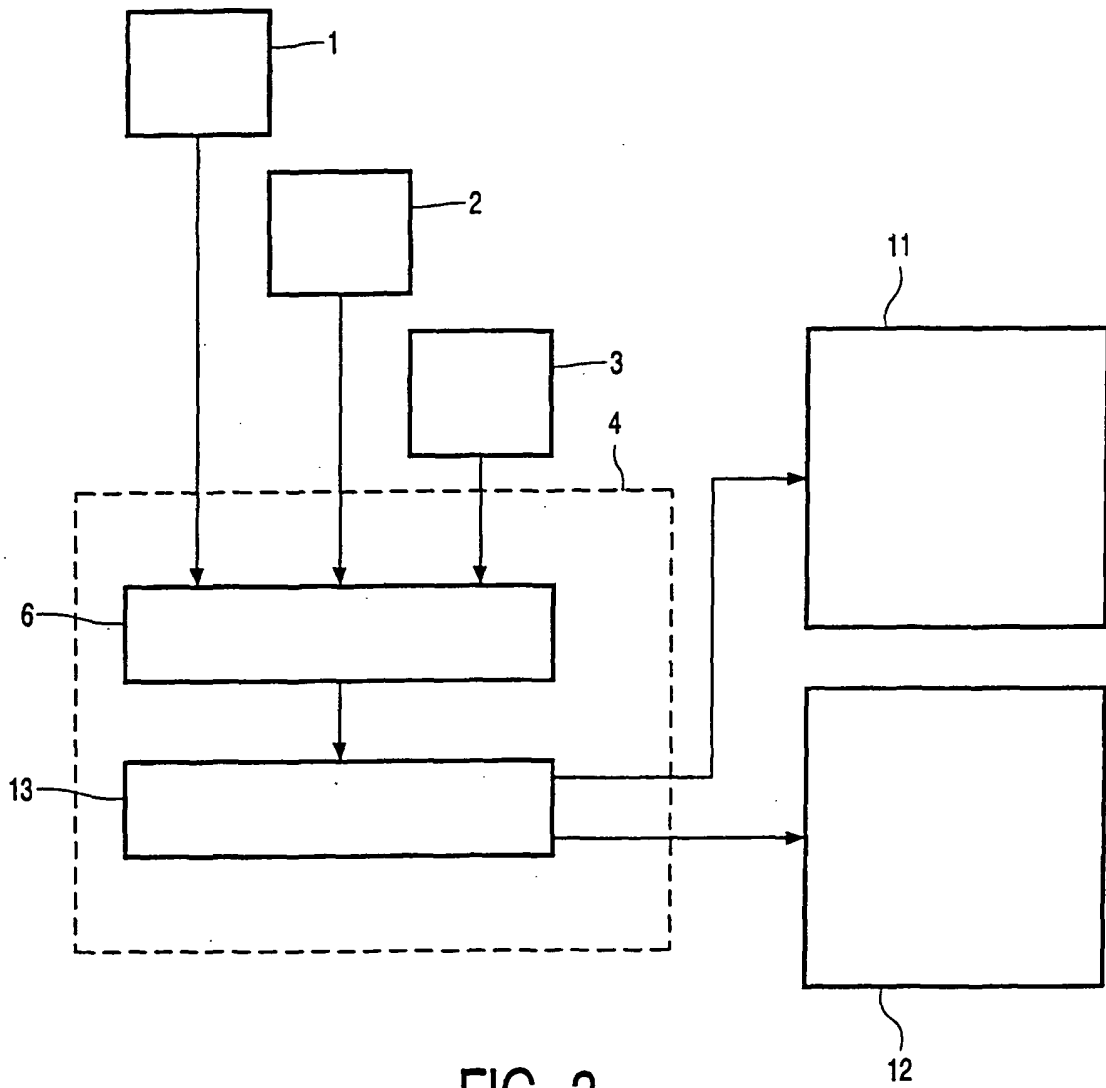


FIG. 3

REFERENCES CITED IN THE DESCRIPTION

This list of references cited by the applicant is for the reader's convenience only. It does not form part of the European patent document. Even though great care has been taken in compiling the references, errors or omissions cannot be excluded and the EPO disclaims all liability in this regard.

Patent documents cited in the description

- US 5719398 A [0006]
- EP 0925017 A [0044]

Non-patent literature cited in the description

- **S.B. COLAK et al.** Clinical Optical Tomography and NIR Spectroscopy for Breast Cancer Detection. *IEEE Journal of Selected Topics in Quantum Electronics*, July 1999, vol. 5 (4 [0004]

专利名称(译)	用于定位混浊介质中的偏差区域的方法和设备		
公开(公告)号	EP1272102B1	公开(公告)日	2009-02-11
申请号	EP2001929448	申请日	2001-03-22
[标]申请(专利权)人(译)	皇家飞利浦电子股份有限公司		
申请(专利权)人(译)	皇家飞利浦电子N.V.		
当前申请(专利权)人(译)	皇家飞利浦电子N.V.		
[标]发明人	VAN DER MARK MARTINUS B T HOOFT GERT W WACHTERS ARTHUR J H		
发明人	VAN DER MARK, MARTINUS, B. 'T HOOFT, GERT, W. WACHTERS, ARTHUR, J., H.		
IPC分类号	A61B5/00 G01N21/47 G01N21/27 A61B5/145 A61B5/1455 A61B10/00 G01N21/35		
CPC分类号	A61B5/0091 A61B5/1455 A61B5/4312 G01N21/4795		
优先权	2000201164 2000-03-31 EP		
其他公开文献	EP1272102A1		
外部链接	Espacenet		

摘要(译)

本发明涉及定位混浊介质中的偏差区域的方法。本发明还涉及用于实施这些方法的装置。所述方法可以用于光学乳房X射线照相术，其中通过光检查女性身体的乳房。所述方法产生图像，其中可以清楚地识别任何偏差，例如肿瘤。这尤其通过在混浊介质的图像(9)中提供标记(10)来实现。

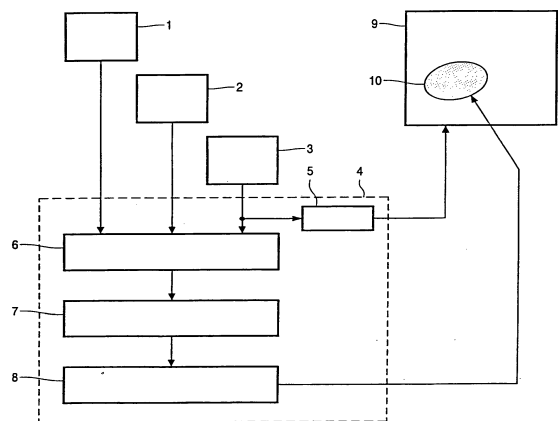


FIG. 1