



(11) **EP 1 611 840 B1**

(12) **EUROPEAN PATENT SPECIFICATION**

(45) Date of publication and mention of the grant of the patent:
13.08.2008 Bulletin 2008/33

(51) Int Cl.:
A61B 5/00 ^(2006.01) **A61B 3/12** ^(2006.01)
A61B 5/021 ^(2006.01)

(21) Application number: **05253758.6**

(22) Date of filing: **16.06.2005**

(54) **Analysis of retinal metabolism over at least a portion of a cardiac cycle**

Analyse des rétinaux Stoffwechsels für mindestens einen Teil eines Herzzyklus

Analyse du métabolisme rétinien pour au moins une partie d'un cycle cardiaque

(84) Designated Contracting States:
DE ES FR GB IT

(30) Priority: **29.06.2004 GB 0414570**

(43) Date of publication of application:
04.01.2006 Bulletin 2006/01

(73) Proprietor: **Kerr, Patrick**
Reigate,
Surrey RH2 7DJ (GB)

(72) Inventor: **Kerr, Patrick**
Reigate,
Surrey RH2 7DJ (GB)

(74) Representative: **Want, Clifford James**
Harrison Goddard Foote
40-43 Chancery Lane
London WC2A 1JA (GB)

(56) References cited:
WO-A-02/080759 **US-B1- 6 244 712**

- **SCOTT V A ET AL: "Retinal pulse oximetry: towards a method for measuring cerebral oxygen saturation" ENGINEERING IN MEDICINE AND BIOLOGY SOCIETY, 1995., IEEE 17TH ANNUAL CONFERENCE MONTREAL, QUE., CANADA 20-23 SEPT. 1995, NEW YORK, NY, USA, IEEE, US, vol. 2, 20 September 1995 (1995-09-20), pages 1555-1556, XP010214854 ISBN: 0-7803-2475-7**
- **VANZETTA I ET AL: "Novel Intrinsic Optical Signals in Feline and Human Retina Evoked by Photic Stimulation." ANNUAL MEETING OF THE ASSOCIATION FOR RESEARCH IN VISION AND OPHTHALMOLOGY; FORT LAUDERDALE, FLORIDA, USA; MAY 05-10, 2002, vol. 2002, 2002, page Abstract No. 4363, XP008052444 ANNUAL MEETING OF THE ASSOCIATION FOR RESEARCH IN VISION AND OPHTHALMOLOGY; FORT LAUDERDALE, FLORIDA, USA; MAY 05-10, 2002**
- **ALONI E H ET AL: "Non-invasive Imaging of Retinal Blood Flow and Oximetry by a new Retinal Function Imager." ARVO ANNUAL MEETING ABSTRACT SEARCH AND PROGRAM PLANNER, vol. 2002, 2002, page Abstract No. 2552, XP008052213 & ANNUAL MEETING OF THE ASSOCIATION FOR RESEARCH IN VISION AND OPHTHALMOLOGY; FORT LAUDERDALE, FLORIDA, USA; MAY 05-10, 2002**

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).

EP 1 611 840 B1

Description

[0001] This invention relates to analysis of retinal metabolism over at least a portion of a cardiac cycle to provide an objective assessment of retinal metabolism. In an embodiment of the invention retinal metabolic response to optical stimuli is determined.

[0002] The retina is a complex structure that contains photoreceptor cells, a network of nerve cells, blood vessels and a metabolically active retinal pigment epithelium. Common retinal diseases that affect retinal metabolism include age-related macular degeneration, diabetic retinopathy and glaucoma, which may cause visual loss and blindness.

[0003] Known retinal imaging technology provides some structural and functional information about retinal function and hence retinal diseases.

[0004] Thus a scanning laser ophthalmoscope disclosed in R. H. Webb, G. W. Hughes, and O. Pomerantz-eff, Flying spot TV ophthalmoscope, Applied Optics 19, 2991-2997, 1980, uses a laser scanning light source to image a retina, subsequently combined with a confocal optical filter to select some light reflected from the retina.

[0005] Scanning laser ophthalmoscope indirect mode imaging, for analysing indirectly-reflected light, which uses an annular aperture and stop to block on-axis reflections to form an image from laterally scattered light reflections, is disclosed in Ann E Elser, Stephen A. Burns, John J Weitter, Francois C Delori, Infrared imaging of sub-retinal structures in the human ocular fundus, Vision Res. Vol. 36, No 1, pp. 191-205, 1996.

[0006] Retinal oximetry produces a numerical value measurement of the percentage oxygen saturation of blood in retinal arteries and veins. A small region of multi-spectral retinal images of a retinal blood vessel and small adjacent region of the retina is selected for analysis and a numerical percentage oxygen saturation of blood in the blood vessel calculated.

[0007] The retinal metabolic image changes over the duration of a heartbeat. With the arrival of a retinal arterial pulse there is an increase of retinal haemoglobin oxygenation. This is followed by a fall in retinal haemoglobin oxygenation, due to oxygen consumption within the metabolically active retinal tissue, before the next retinal arterial pulse.

[0008] US 6,244,712 discloses optical scanning spectroscopic retinal blood vessel oximetry using a plurality of wavelengths to illuminate successive portions of the retina and form an interlaced retinal data frame, to avoid over-illuminating an eye by scanning with the plurality of wavelengths simultaneously. The interlaced retinal data frame may be de-interlaced to form plural monochromatic retinal images corresponding to the respective wavelengths. Signals from the eye may be filtered or selected with confocal or anti-confocal filters before being delivered to a detector. The laser scans may be triggered, for example in response to an r-wave of an electrocardiogram, at a predetermined point in a cardiac cycle thereby

permitting a detailed analysis of one or more phases of the cardiac cycle.

[0009] US 2002/0188203 discloses measurement of blood oxygen saturation in a retinal blood vessel by detecting light that has made a single pass through the retinal blood vessel, i.e. retinal vein or artery, and then been diffused laterally through retinal and/or choroidal layers and left the eye without again passing through the retinal blood vessel. An anti-confocal optical filter, with an aperture and central stop, is used to isolate such single-pass optical signals and thereby simplify calculation of retinal blood oxygen saturation, resulting in increased accuracy of measurement of oxygen saturation. This provides an objective assessment of retinal haemoglobin oxygenation within a small portion of the retinal blood vessel. It is suggested that such measurements can be used to monitor cardiac output of a subject or detect and determine a rate of blood loss.

[0010] A retinal metabolic image changes in response to a light stimulus of the retina caused by an increase in retinal neuronal metabolic activity and therefore of oxygen consumption.

[0011] US 2004/0075812 discloses detection of changes in reflectance of near-infrared light from the retina of human subjects, caused by changes in oxygen saturation in response to visual activation of the retina by a light pattern or other light stimulus. This provides an objective assessment of inner retinal function, allowing detection, at an early stage, of a regional defect caused by glaucoma.

[0012] Haemoglobin oxygenation saturation light absorption is disclosed by, Van Assendelft OW. Spectrophotometry of haemoglobin derivatives. Royal Vangorcum, Assen, The Netherlands: Thomas, 1970.

[0013] WO 02/080759 discloses a retinal function camera in which use of isoreflexive points enables isolation of retinal haemoglobin oxygenation image data from multi-spectral retinal images, and formation of a retinal metabolic image based on haemoglobin oxygenation. The retinal metabolic image provides a subjective assessment of retinal haemoglobin oxygenation image data captured within the imaging time period. The retinal function image may be synchronised with an R wave of a subject's electrocardiogram, to study retinal metabolism at a predetermined time in a cardiac cycle of the subject. That is, a predetermined time delay may be allowed between detection of the R wave and formation of a scanned image.

[0014] The prior art, therefore, provides: oximetry, an objective assessment of retinal haemoglobin oxygenation within a small portion of a retinal blood vessel; an objective assessment of infrared light reflectance change of the retina to light stimulus; and a subjective assessment of retinal haemoglobin oxygenation image data. The prior art does not provide an objective assessment of retinal metabolism based on haemoglobin oxygenation.

[0015] According to the present invention, there is pro-

vided a method of analysing retinal metabolism over at least a portion of a cardiac cycle, the method comprising the steps of: a) illuminating a portion of a retina of an eye with light of a first wavelength by providing a first light source and emitting light of the first wavelength and providing scanning means to produce a scanning beam and scanning the portion of the retina with light from the first light source; b) producing a first image of the portion of the retina illuminated with the light of the first wavelength; c) illuminating the portion of the retina with light of a second wavelength, the first and second wavelengths being selected such that absorptivity of light of the first wavelength by oxygenated blood is greater than absorptivity of light of the second wavelength and the absorptivity of light of the first wavelength by deoxygenated blood is less than absorptivity of light of the second wavelength; d) producing a second image of the portion of the retina illuminated with the light of the second wavelength; e) processing the first and second images with image processing means to map relative oxygenation of the portion of the retina as an indication of retinal metabolic function of the portion of the retina by determining isoreflective points of the respective images at which absorption of light of the first wavelength is substantially equal to absorption of light of the second wavelength and determining areas of the respective images having differential absorptivity for the first and second wavelengths; isolating haemoglobin oxygenation image data from the first and second wavelength images by subtracting the isoreflective point from respective first and second wavelength images, the contrast of each portion of the processed images thereby being proportional to a difference in oxygenation of that portion of the processed images from the oxygenation of the isoreflective point, to obtain retinal metabolic image data based on haemoglobin oxygenation; and f) repeating steps a) to e) within the at least a portion of a cardiac cycle to analyse metabolic function changes of the portion of the retina to detect a haemoglobin oxygenation image data retinal arterial waveform within the at least a portion of a cardiac cycle to provide synchronisation with a cardiac cycle of a subject for analysing retinal metabolic image data over the at least a portion of the cardiac cycle.

[0016] Advantageously, the step of illuminating the portion of the retina with light of a second wavelength comprises providing: a second light source and emitting light of the second wavelength and first optical beam combiner means and selectively directing light from the first light source and from the second light source to the scanning means to produce a scanning beam and scanning the portion of the retina with light from the second light source subsequently to scanning the portion of the retina with light from the first light source.

[0017] Conveniently, at least one of the first light source and the second light source is synchronised with the scanning means.

[0018] Conveniently, the method includes a further step of providing first focusing means and focusing the

scanning beam from the scanning means onto the at least a portion of the retina with the first focusing means.

[0019] Advantageously, the method includes a step of providing refractive error correcting means and correcting for refractive errors of the eye with the refractive error correcting means, to permit focusing of the scanning beam on the at least a portion of the retina.

[0020] Conveniently, the step of providing refractive error correcting means comprises providing adaptive optics and correcting for optical aberrations of the eye.

[0021] Advantageously, the step of providing refractive error correcting means comprises providing wave-front sensor means and wave-front compensation means.

[0022] Conveniently, the step of providing wave-front compensation means comprises providing deformable mirror means.

[0023] Conveniently, the step of producing first and second images includes providing beam splitter means for splitting a beam reflected from the at least a portion of the retina when illuminated by light of the first wavelength and light of the second wavelength to form a split beam; image sensor means and second focusing means and focusing the split beam on the image sensor means for sensing an image of the at least a portion of the retina and image capture means for capturing the sensed image.

[0024] Advantageously, the step of providing image capture means comprises providing frame grabber means.

[0025] Advantageously, the scanning means is synchronised with at least one of the image sensor means, the image capture means and the image processing means.

[0026] Conveniently, the step of illuminating the portion of the retina with light of the first wavelength comprises illuminating the portion of the retina with light of wavelengths centred on one of 830 nm, 850 nm and 910 nm.

[0027] Conveniently, the step of illuminating the portion of the retina with light of the second wavelength comprises illuminating the portion of the retina with light of wavelengths centred on one of 635 nm, 670 nm and 760 nm.

[0028] Advantageously, step f) further comprises determining from the metabolic function changes a waveform reference point in a haemoglobin oxygenation image data waveform corresponding to a cardiac cycle.

[0029] Conveniently, the step of determining a waveform reference point comprises determining an amplitude trough between a trailing edge and a leading edge of an arterial pulse of the haemoglobin oxygenation image data waveform.

[0030] Optionally, the step of determining a waveform reference point comprises determining an amplitude peak of an arterial pulse of the haemoglobin oxygenation image data waveform.

[0031] Optionally, the step of determining a waveform reference point comprises determining an inflection point

between an amplitude trough and an amplitude peak of an arterial pulse of the haemoglobin oxygenation image data waveform.

[0032] Conveniently, the method is adapted for determining oxygenation changes associated with the cardiac cycle.

[0033] Conveniently, the method is adapted for determining characteristics of at least one of amplitude, pattern, shape and duration of the cardiac cycle.

[0034] Advantageously, the step of determining oxygenic changes comprises using Fourier analysis for determining a cardiac cycle waveform.

[0035] Advantageously, step f) further comprises subtracting effects of changes in oxygenation caused by the cardiac cycle to determine metabolic function of the at least a portion of the retina independently of oxygenation changes caused by the cardiac cycle.

[0036] Advantageously, the method includes a further step of subjecting the at least a portion of the retina to optical stimulation and analysing effects of the optical stimulation on retinal metabolic function.

[0037] Conveniently, the optical stimulation of the at least a portion of the retina is synchronised with reference to the waveform reference point.

[0038] Advantageously, step f) further comprises subtracting effects of changes in oxygenation caused by the cardiac cycle from the effects of changes in oxygenation caused by the optical stimulation to isolate changes in retinal metabolism caused by the optical stimulation.

[0039] Advantageously, the method further comprises analysing isolated responses of retinal metabolism caused by the optical stimulation to determine waveform characteristics of at least one of latency between the optical stimulation and the retinal metabolism response, amplitude, pattern, shape and duration of the response.

[0040] Conveniently, the step of subjecting the at least a portion of the retina to optical stimulation comprises providing an optical signal stimulus light source; and second beam combining means and combining a light beam from the optical signal stimulus light source selectively with one of light of the first wavelength and light of the second wavelength.

[0041] Advantageously, the step of providing an optical signal stimulus light source comprises providing an array of light-emitting diodes and one of simultaneously or sequentially stimulating a plurality of points on the at least a portion of the retina.

[0042] Conveniently, the optical signal stimulus light source is modulated to generate an optical signal stimulus pattern on the at least a portion of a retina.

[0043] Conveniently, the optical signal stimulus light source comprises an optical signal stimulus laser.

[0044] Advantageously, the method is adapted for determining cardiac function.

[0045] Advantageously, the method is adapted for determining cerebral arterial circulatory function.

[0046] Conveniently the method is adapted for evaluating effects of therapeutic agents on retinal metabolism.

[0047] Conveniently the method is adapted for evaluating effects of therapeutic agents on retinal metabolic response to optical stimulation.

[0048] According to a second aspect of the invention, there is provided a retinal function camera comprising: a first source of light of a first wavelength band; a second source of light of a second wavelength band, the absorptivity of light of the first wavelength band by oxygenated blood being greater than the absorptivity of light of the second wavelength band and the absorptivity of light of the first wavelength band by deoxygenated blood being less than the absorptivity of light of the second wavelength band; an optical signal stimulus light source distinct from the first source of light and the second source of light; means for focusing light from the optical signal stimulus light source and selectively from the first and second sources onto a portion of a retina of an eye; imaging means for producing respective images of the portion of the retina illuminated with the respective wavelength bands and stimulated by the optical signal stimulus light source; and image processing means adapted to process the respective images obtained by the imaging means to determine isorefective points of the respective images at which absorption of light of the first wavelength is substantially equal to absorption of light of the second wavelength and areas of the respective images having differential absorptivity for the first and second wavelengths, repeatedly to obtain a retinal function image based on haemoglobin oxygenation within at least a portion of a cardiac cycle to analyse metabolic functional changes of the portion of the retina to detect a haemoglobin oxygenation image data retinal arterial waveform within the at least a portion of a cardiac cycle to provide synchronisation with a cardiac cycle of a subject for analysing retinal metabolic image data over the at least a portion of the cardiac cycle when the retina is subjected to optical stimulation.

[0049] Conveniently, the optical signal stimulus light source comprises an optical signal stimulus laser.

[0050] Conveniently, the retinal function camera further comprises scanning means arranged for producing a scanning beam for scanning the portion of the retina with light from the optical signal stimulus light source and selectively from the first and second sources.

[0051] Advantageously, the retinal function camera further comprises first optical beam combiner means arranged for selectively directing light from the first light source and from the second light source to the scanning means.

[0052] Advantageously, the retinal function camera further comprises first synchronisation means arranged for synchronising at least one of the first light source and the second light source with the scanning means.

[0053] Advantageously, the retinal function camera further comprises refractive error correcting means arranged for correcting for refractive errors of the eye to permit focusing of the scanning beam on the at least a portion of the retina.

[0054] Conveniently, the refractive error correcting means comprises adaptive optics arranged for correcting for optical aberrations of the eye.

[0055] Conveniently, the refractive error correcting means comprises wave-front sensor means and wave-front compensation means.

[0056] Advantageously, the wave-front compensation means comprises deformable mirror means.

[0057] Conveniently, the retinal function camera further comprises beam splitter means arranged for splitting a beam reflected from the at least a portion of the retina when illuminated by light of the first wavelength band and light of the second wavelength band to form a split beam; image sensor means and second focusing means arranged for focusing the split beam on the image sensor means for sensing an image of the at least a portion of the retina and image capture means for capturing the sensed image.

[0058] Conveniently, the image capture means comprises frame grabber means.

[0059] Advantageously, the retinal function camera further comprises second synchronisation means arranged for synchronising at least one of the image sensor means, the image capture means and the image processing means with the scanning means.

[0060] Conveniently, the first wavelength band comprises wavelengths centred on one of 830 nm, 850 nm and 910 nm.

[0061] Conveniently, the second wavelength band comprises wavelengths centred on one of 635 nm, 670 nm and 760 nm.

[0062] Further, there is provided computer executable software code stored on a computer readable medium, the code being for analysing retinal metabolism over at least a portion of a cardiac cycle, comprising the steps of: a) illuminating a portion of a retina of an eye with light of a first wavelength; b) producing a first image of the portion of the retina illuminated with the light of the first wavelength; c) illuminating the portion of the retina with light of a second wavelength, the first and second wavelengths being selected such that absorptivity of light of the first wavelength by oxygenated blood is greater than absorptivity of light of the second wavelength and the absorptivity of light of the first wavelength by deoxygenated blood is less than absorptivity of light of the second wavelength; d) producing a second image of the portion of the retina illuminated with the light of the second wavelength; e) processing the first and second images with image processing means to map relative oxygenation of the portion of the retina as an indication of retinal metabolic function of the portion of the retina; and f) repeating steps a) to e) within the at least a portion of a cardiac cycle to analyse metabolic function changes of the portion of the retina to detect a haemoglobin oxygenation image data retinal arterial waveform within the at least a portion of a cardiac cycle to provide synchronisation with a cardiac cycle of a subject for analysing retinal metabolic image data over the at least a portion of the cardiac cycle.

[0063] Further, there is provided one or more programmed computers for analysing retinal metabolism over at least a portion of a cardiac cycle comprising the steps of: a) illuminating a portion of a retina of an eye with light of a first wavelength; b) producing a first image of the portion of the retina illuminated with the light of the first wavelength; c) illuminating the portion of the retina with light of a second wavelength, the first and second wavelengths being selected such that absorptivity of light of the first wavelength by oxygenated blood is greater than absorptivity of light of the second wavelength and the absorptivity of light of the first wavelength by deoxygenated blood is less than absorptivity of light of the second wavelength; d) producing a second image of the portion of the retina illuminated with the light of the second wavelength; e) processing the first and second images with image processing means to map relative oxygenation of the portion of the retina as an indication of retinal metabolic function of the portion of the retina; and f) repeating steps a) to e) within the at least a portion of a cardiac cycle to analyse metabolic function changes of the portion of the retina to detect a haemoglobin oxygenation image data retinal arterial waveform within the at least a portion of a cardiac cycle to provide synchronisation with a cardiac cycle of a subject for analysing retinal metabolic image data over the at least a portion of the cardiac cycle.

[0064] Other aspects and features of the present invention will become apparent to those ordinarily skilled in the art upon review of the following description of specific embodiments of the invention in conjunction with the accompanying figures.

[0065] Embodiments of the present invention will now be described, by way of example only, with reference to the accompanying drawing in which Figure 1 is a schematic diagram of an apparatus according to the invention.

[0066] Throughout the description, identical reference numerals are used to identify like parts.

[0067] A retinal function camera 200 according to the invention, as shown in Figure 1, has arranged sequentially on a first optical axis: a first light source 21 for emitting light of a first wavelength, a first optical beam combiner 30, a second optical beam combiner 50, an optical beam splitter 60, a first focusing lens 70, a vertical and horizontal optical beam scanner 80 and an imaging lens 90. A second light source 22, for alternatively emitting light of a second wavelength different from the first wavelength, is arranged so that light from the second light source incident on the first optical beam combiner 30 is alternatively transmittable along the first optical axis. An optical signal stimulus laser 40 is arranged so that light selectively emitted from the optical signal stimulus laser and incident on the second optical beam combiner 50 is transmittable along the first optical axis.

[0068] The optical beam splitter 60 is also located on a second optical axis, substantially perpendicular to the first optical axis. Sequentially arranged on the second optical axis, downstream from the optical beam splitter

60, are an orthogonal polarising filter 100, a second focusing lens 110, a confocal optical filter 120 and an image sensor 130. The imaging sensor may be a photomultiplier tube, an avalanche photodiode, a CCD imaging sensor or a CMOS imaging sensor. Operationally coupled to the image sensor is a frame grabber 140, and operationally coupled to the frame grabber 140 is an image processor 150. As an alternative to the frame grabber, any other known device for capturing the image may be provided.

[0069] One or more of the optical components may optionally be coupled by one or more optical fibres respectively (not shown) in either or both of illuminating and imaging paths, so that the coupled components are not necessarily physically aligned on one of the optical axes.

[0070] An endoscope arrangement may alternatively be used.

[0071] Moreover, although the first optical axis is shown schematically as substantially perpendicular to the second optical axis, it will be understood that other relative orientations of the axes may be employed.

[0072] The first and second light sources 21, 22 are provided such that absorptivity of light emitted by the first light source by oxygenated blood is greater than absorptivity of light emitted by the second light source and the absorptivity of light emitted by the first light source by deoxygenated blood is less than absorptivity of light emitted by the second light source. The first light source 21 and the second light source 22 emit light with respective wavelengths preferably in a range between 488 nm and 1000 nm. The first light source 21 emits wavelengths preferably centred substantially on one of 830 nm, 850 nm and 910 nm. The second light source 22 emits wavelengths preferably centred substantially on one of 635 nm, 670 nm and 760 nm. The first light source 21 is preferably a laser or a superluminescent diode and the second light source 22 is also preferably a laser or a superluminescent diode.

[0073] In use, light from the first light source 21 and the second light source 22 is sequentially scanned over a same portion of the retina of an eye 10 of a subject. The portion of the retina may be, for example, a pixel, a line scan or a complete image scan.

[0074] In detail, a first incident light beam 211 emitted by the first light source 21 is incident on a first face of the first optical beam combiner 30 at a first angle of incidence of substantially 45 degrees, such that the light beam is transmitted through the first optical beam combiner to be incident on, and pass through, the second optical beam combiner 50. The light beam then passes sequentially through the optical beam splitter 60 and a focus lens 70 for correcting for refractive errors of the eye 10, to permit eventual focusing of the light beam on a retina of the eye 10. The corrected light beam is incident on a vertical and horizontal scanner 80 and subsequently focused by the imaging lens 90 on the retina of the eye 10.

[0075] The scanner 80 and imaging lens 90 scan focused light in a raster manner across at least a portion

of the retina. The scanner 80 is a two-axis scanner including a horizontal scanner for scanning the focussed light horizontally across at least a horizontally extending portion of the retina and a vertical scanner for scanning the focussed light vertically across at least a vertically extending portion of the retina. The horizontal scanner may include one of a rotatable polygonal mirror and a vibratable plane mirror and the vertical scanner may include a galvanometer scanner.

[0076] The scanner 80 is synchronised to the first light source 21 of light and to the second light source 22 by a controller (not shown).

[0077] The scanner 80 is synchronised by the controller (not shown) to the image sensor 130, the frame grabber 140 and the image processor 150.

[0078] A fixation target light emitting diode (not shown) may be positioned between the scanner 80 and the eye 10 to enable a subject to fix a direction of the eye for analysis.

[0079] A second incident light beam 221, subsequently emitted from the second light source 22 in a direction substantially perpendicular to the first light beam 211, is incident on a second face of the first optical beam combiner 30, opposed to the first face, at a second angle of incidence of substantially 45 degrees such that the second light beam has access to the first optical axis of the retinal function camera 200.

[0080] That is, a same portion of the retina is sequentially scanned with light from the first light source 21 and then from the second light source 22.

[0081] Light reflected from the retina at a back of the eye 10 re-passes through the imaging lens 90, is de-scanned by the scanner 80 and re-passes through the first focusing lens 70 to the optical beam splitter 60.

[0082] The optical beam splitter 60 directs light reflected from the retina to the orthogonal polarising filter 100. The polarising filter blocks light reflected from eye surfaces with substantially the same polarisation as the first light source 21 and the second light source 22.

[0083] A second focusing lens 110 focuses light reflected from the retina towards an optical filter 120.

[0084] The optical filter 120 is substantially confocal to the imaged retina. The optical filter may be a confocal aperture or an anti-confocal stop. Alternatively, the optical filter may be a combined confocal aperture and an anti-confocal stop forming an annular aperture. The filter blocks a portion of the reflected light from the retina reaching and being detected by the imaging sensor 130. For example, an anti-confocal filter will allow laterally scattered light signals to be detected while blocking on-axis light signals.

[0085] The imaging sensor 130 is sensitive to light emitted by the first light source 21 and the second light source 22. The imaging sensor converts the reflected light signals to electrical signals. The electrical signals are amplified, synchronised with the scanning means 80 and frame grabber 140 to form a time and space resolved image data frame.

[0086] The image processor 150 processes the image data frames. The image processor includes at least a computer and image processing and analysis software. Suitable software includes LabVIEW® image processing and analysis software available from National Instruments Corporation, Austin, Texas, United States of America.

[0087] Adaptive optics (not shown) may be used to correct for optical aberrations of the eye. A wave-front sensor such as a Shack-Hartmann wave-front sensor may be used to measure wave aberration of the light that is used to form the image. Wave-front compensation with a deformable mirror, before the raster scanner, corrects optical aberrations on the illuminating and return imaging light paths. The deformable mirror shape may be controlled by LabVIEW® software with data derived from the wave-front sensor.

[0088] A retinal metabolic image data based on haemoglobin oxygenation is obtained by using the first and second light sources with such wavelengths that the absorptivity of light of the first wavelength by oxygenated blood is greater than the absorptivity of light of the second wavelength and the absorptivity of light of the first wavelength by deoxygenated blood is less than the absorptivity of light of the second wavelength. The respective images of the portion of the retina illuminated with the respective wavelength bands are processed to determine isorefective points of the respective images at which absorption of light of the first wavelength is substantially equal to absorption of light of the second wavelength and areas of the respective images having differential absorptivity for the first and second wavelengths. The haemoglobin oxygenation image data is isolated from the respective first and second wavelength images by subtracting the reflectivity at the isorefective point from respective first and second wavelength images. The contrast of each portion of the processed images is then proportional to a difference in oxygenation of that respective point from the oxygenation of the isorefective point, thereby providing retinal metabolic image data based on haemoglobin oxygenation.

[0089] The retinal metabolic haemoglobin oxygenation image data may be time synchronised to a heartbeat of the subject. Conventional synchronisation to an external signal, which may be used, includes synchronisation to an R-wave of an electrocardiogram or to pulse blood volume waveform detection with an infrared photoplethysmograph. However, in an embodiment of the invention, the haemoglobin oxygenation image data from a same portion of successive data frames is analysed to detect the haemoglobin oxygenation image data retinal arterial waveform to provide synchronisation with a cardiac cycle of the subject. For example, line 1 of a data frame may be compared with line 1 of successive data frames, line 2 may be compared with line 2 of successive data frames, line x may be compared with line x of successive data frames. The amplitude of successive line haemoglobin oxygenation image data is determined to

generate a pattern or shape of successive line haemoglobin oxygenation image data. A steep rise waveform related to an arterial pulse is determined from the falling haemoglobin oxygenation image data slope. An amplitude foot at a junction between falling slope and steep arterial pulse on the haemoglobin oxygenation image data waveform may be identified and used as a retinal arterial waveform reference point. Optionally, an amplitude peak at a peak arterial pulse on the haemoglobin oxygenation image data waveform may be identified and used as the retinal arterial waveform reference point. Optionally again, an inflection point between an amplitude foot and an amplitude peak, i.e. when a second derivative of the waveform changes sign, on the haemoglobin oxygenation image data waveform may be identified and used as the retinal arterial waveform reference point. A period between a retinal arterial waveform reference point and a next retinal arterial waveform reference point defines a functional time period of a cardiac cycle, or heartbeat. The functional time period of a heartbeat may be defined from at least a portion of the image data.

[0090] A portion of the retinal metabolic image may be selected for analysis. The portion may be, for example, at least a pixel, a line scan, a data frame or an ophthalmologist-defined portion of the macula or optic disc. The portion may be each pixel of a data frame. The portion may be a subset of the haemoglobin oxygenation image data and may correspond to more haemoglobin oxygenation than the isorefective point. Alternatively, the portion may be haemoglobin oxygenation image data corresponding to less haemoglobin oxygenation than the isorefective point. The portion may be haemoglobin oxygenation image data that changes from more haemoglobin oxygenation than the isorefective point to less haemoglobin oxygenation than the isorefective point during the functional time period of a heartbeat. The haemoglobin oxygenation image data may be analysed to determine waveform characteristics of amplitude, pattern, shape, duration and response to optical signal stimuli. At least a portion of the image data may be analysed. A portion of the image data may be analysed over time. A portion of the image data may be analysed by Fourier analysis. A portion of the image data may be analysed over a functional time period of at least a heartbeat. A portion of the image data may be analysed over a functional time period of at least two heartbeats to determine average data waveform characteristics. Image alignment processing, optionally including pattern recognition, may be used.

[0091] The imaged eye may be position-stabilised by contra-lateral eye fixing on an external optical fixation target. That is, an optical fixation point may be used to help to maintain retinal image positional stability. The fixation point may be internal or external of the imaging optics. A typical fixation point is a light emitting diode. An optical fixation and optical signal stimulus laser 40 may be used instead of a light emitting diode source fixation point. The wavelength of the optical fixation point source

may be selected between 488 nm and 760 nm. The optical fixation point wavelength may be centred substantially on 532 nm. The optical signal stimulus laser may be synchronised to the scanner 80. The optical signal stimulus laser may be synchronised to a retinal arterial waveform reference point.

[0092] A band-pass filter (not shown) may be used to block the green 532 nm optical signal stimulus light reflected from the eye from reaching the imaging sensor 130.

[0093] The optical signal stimulus laser may be modulated to generate an optical signal stimulus pattern on the retina of the eye 10.

[0094] That is, the optical fixation and optical signal stimulus laser 40 emits an optical stimulus light beam 401, centred substantially on a wavelength of 532 nm. The optical stimulus light beam 401 is incident at an angle of substantially 45 degrees on a first face of the second optical beam combiner 50 located on the first optical axis, downstream of the first optical beam combiner 30, on which the first incident beam 211 is incident on a second face, opposed to the first face, to combine the first incident beam 211 with the optical stimulus light beam 401 to form a combined beam 501 with access to the first optical axis. The combined beam passes sequentially through the optical beam splitter 60 and the first focusing lens 70 for correcting for refractive errors of the eye 10 to permit eventual focusing of the corrected combined beam 701 on a retina of the eye 10. The corrected combined beam 701 is incident on a vertical and horizontal scanner 80 to form a scanning beam 801 substantially on the first optical axis. The imaging lens 90 focuses the scanning beam 801 as a focussed scanning beam 901 onto the retina of the eye 10.

[0095] Alternatively, light from the second light source may be combined with the optical stimulus light beam to form a combined beam.

[0096] The optical signal stimulus laser 40 may be synchronised to the scanner 80 by a controller (not shown). The optical signal stimulus laser 40 is modulated to generate an optical signal stimulus pattern on the retina of the eye 10.

[0097] Illuminating the internal fixation target light emitting diode or optical fixation and optical signal stimulus laser 40, allows retinal metabolic image changes in response to the optical signal stimulus to be imaged. The retinal metabolic image changes in response to the optical signal stimulus may be imaged and isolated by image subtraction from the background functional retinal metabolic image data obtained over a heartbeat, preferably after image alignment. Beat-to-beat variation of the background functional retinal metabolic image data obtained over a heartbeat may be averaged. The retinal metabolic image changes in response to the optical signal stimulus may be imaged and isolated by image subtraction from the averaged background functional retinal metabolic image data obtained over a heartbeat and fitted for amplitude and duration. The isolated retinal met-

abolic image changes in response to the optical signal stimulus may be analysed to determine waveform characteristics of latency, amplitude, pattern, shape, and duration. The isolated retinal metabolic image changes in response to the optical signal stimulus may be displayed in image, numerical or graphic format. The background functional retinal metabolic image data obtained over a heartbeat is dynamically varying and a function of the arterial haemoglobin oxygen supply and retinal metabolic oxygen consumption. The isolation of a subtle alteration in retinal tissue oxygenation due to an optical signal stimulus from a dynamically varying background is enhanced by the functional time synchronised reference background functional retinal metabolic image data.

[0098] An array of light emitting diodes arranged in a grid may be used to allow multiple points on the retina of the eye 10 to be stimulated simultaneously. The optical signal stimulus laser 40 may be modulated to generate an optical signal stimulus pattern on the retina of the eye 10. The optical signal stimulus laser 40 may be modulated to generate an optical signal stimulus pattern on the retina of the eye 10 with a single optical fixation point. The optical signal stimulus laser 40 may be modulated to generate an optical signal stimulus pattern on the retina of the eye 10 with a single optical fixation point and then switch on and off a pattern while retaining a single optical fixation point. The duration of illumination may be controlled. The light intensity may be controlled.

[0099] An array of external light emitting diodes may be arranged in a circular grid as an external optical fixation target. Sequentially fixing the eye on sequentially illuminated light emitting diodes allows the retinal periphery to be imaged.

[0100] The retinal metabolic image changes over the duration of a heartbeat. With the arrival of a retinal arterial pulse there is an increase of retinal haemoglobin oxygenation. This is followed by a fall in retinal haemoglobin oxygenation, due to oxygen consumption within the metabolically active retinal tissue, before the next retinal arterial pulse. Retinal metabolic image data may be analysed over a functional time period of a heartbeat to provide an objective assessment of retinal metabolism. The retinal metabolic image data is derived from the dynamic function of arterial oxygenated haemoglobin supply and the retinal metabolic oxygen consumption. Changes in the arterial oxygenated haemoglobin supply may be detected. If the cardiac output is impaired then tissue perfusion with oxygenated blood will be impaired. Retinal metabolic image data may detect a reduced cardiac output. Retinal metabolic image data may detect a reduced cardiac output secondary to blood loss. Retinal metabolic image data may be analysed over a functional time period of a heartbeat to provide an objective assessment of retinal metabolism with a reduced cardiac output.

[0101] Changes in the arterial oxygenated haemoglobin supply between the heart and the retina may be detected. A cerebral artery haemorrhage may increase the intracranial pressure. Increased intracranial pressure

may impair both retinal arterial supply and retinal vein drainage. A cerebral artery occlusion does not necessarily increase intracranial pressure. Retinal metabolic image data may be analysed over a functional time period of a heartbeat to provide an objective assessment of retinal metabolism with raised intracranial pressure. Characteristics of retinal metabolic image data analysed over a functional time period of a heartbeat may help to differentiate between a cerebral artery occlusion and a cerebral artery haemorrhage.

[0102] Retinal metabolic image data may be analysed over time. The time period may be a functional time period, such as a heartbeat or a discrete time interval. This generates an objective assessment of retinal metabolism by generating numerical or graphical output of retinal metabolism compared with the images requiring subjective assessment in the prior art.

[0103] The retinal metabolic image changes in response to stimulus optical signals may be isolated by image subtraction from the background functional retinal metabolic image data obtained over a heartbeat.

[0104] A resulting isolated retinal metabolism response to the optical stimulation may be analysed for one or more of latency between the optical stimulation and the metabolism response and the amplitude, pattern, shape and duration of a waveform of the response.

[0105] This invention overcomes the lack of an objective assessment of retinal metabolism by analysing the retinal haemoglobin oxygenation image data over a functional time period of at least a portion of a heartbeat to produce an objective assessment of retinal metabolism.

[0106] This invention provides the advantage of isolating retinal haemoglobin oxygenation image data from multi-spectral retinal images, analysing the data over time, determining retinal metabolic response to optical stimuli and providing an objective assessment of retinal metabolism. The invention enables earlier diagnosis of eye disease than in the prior art. The invention enables objective monitoring of eye disease and enables objective monitoring of eye disease response to therapy. Clinical trials may objectively evaluate the effects of therapeutic agents on retinal metabolism. Clinical trials may objectively evaluate the effects of therapeutic agents on retinal metabolic response to optical stimulation.

[0107] Alternative embodiments of the invention can be implemented as a computer program product for use with a computer system, the computer program product being, for example, a series of computer instructions stored on a tangible data recording medium, such as a diskette, CD-ROM, ROM, or fixed disk, or embodied in a computer data signal, the signal being transmitted over a tangible medium or a wireless medium, for example microwave or infrared. The series of computer instructions can constitute all or part of the functionality described above, and can also be stored in any memory device, volatile or non-volatile, such as semiconductor, magnetic, optical or other memory device.

[0108] Although the present invention has been de-

scribed with reference to preferred embodiments, workers skilled in the art will recognize that changes may be made in form and detail without departing from the scope of the invention with defined in the claims..

Claims

1. A method of analysing retinal metabolism over at least a portion of a cardiac cycle, the method comprising the steps of:

- a. illuminating a portion of a retina of an eye (10) with light of a first wavelength by providing a first light source (21) and emitting light (211) of the first wavelength and providing scanning means (80) to produce a scanning beam (801) and scanning the portion of the retina with light from the first light source;
- b. producing a first image of the portion of the retina illuminated with the light of the first wavelength;
- c. illuminating the portion of the retina with light (221) of a second wavelength, the first and second wavelengths being selected such that absorptivity of light of the first wavelength by oxygenated blood is greater than absorptivity of light of the second wavelength and the absorptivity of light of the first wavelength by deoxygenated blood is less than absorptivity of light of the second wavelength;
- d. producing a second image of the portion of the retina illuminated with the light of the second wavelength; the method being **characterized by** further comprising the steps of:
- e. processing the first and second images with image processing means (150) to map relative oxygenation of the portion of the retina as an indication of retinal metabolic function of the portion of the retina by determining isoreflective points of the respective images at which absorption of light (211) of the first wavelength is substantially equal to absorption of light (221) of the second wavelength and determining areas of the respective images having differential absorptivity for the first and second wavelengths; isolating haemoglobin oxygenation image data from the first and second wavelength images by subtracting the reflectivity at the isoreflective point from respective first and second wavelength images, the contrast of each portion of the processed images thereby being proportional to a difference in oxygenation of that portion of the processed images from the oxygenation of the isoreflective point, to obtain retinal metabolic image data based on haemoglobin oxygenation; and
- f. repeating steps a) to e) within the at least a

- portion of a cardiac cycle thereby analysing metabolic function changes of the portion of the retina, detecting a haemoglobin oxygenation image data waveform within the at least a portion of a cardiac cycle and providing synchronisation with a cardiac cycle of a subject for analysing retinal metabolic image data over the at least a portion of the cardiac cycle.
2. A method as claimed in claim 1, wherein the step of illuminating the portion of the retina with light (221) of a second wavelength comprises providing: a second light source (22) and emitting light of the second wavelength and first optical beam combiner means (30) and selectively directing light (211) from the first light source (21) and from the second light source (22) to the scanning means (80) to produce a scanning beam (801) and scanning the portion of the retina with light from the second light source subsequently to scanning the portion of the retina with light from the first light source.
 3. A method as claimed in claim 2, wherein at least one of the first light source (21) and the second light source (22) is synchronised with the scanning means (80).
 4. A method as claimed in any of the preceding claims, including a step of providing first focusing means (90) and focusing the scanning beam (801) from the scanning means (80) onto the at least a portion of the retina with the first focusing means.
 5. A method as claimed in claim 4, including a step of providing refractive error correcting means and correcting for refractive errors of the eye with the refractive error correcting means, to permit focusing of the scanning beam on the at least a portion of the retina.
 6. A method as claimed in any of the preceding claims, wherein the step of producing first and second images includes providing beam splitter means (60) for splitting a beam reflected from the at least a portion of the retina when illuminated by light (211) of the first wavelength and light (221) of the second wavelength to form a split beam; image sensor means (130) and second focusing means (110) and focusing the split beam on the image sensor means for sensing an image of the at least a portion of the retina and image capture means (140) for capturing the sensed image.
 7. A method as claimed in any of the preceding claims, wherein the step of illuminating the portion of the retina with light (211) of the first wavelength comprises illuminating the portion of the retina with light of wavelengths centred on one of 830 nm, 850 nm and 910 nm.
 8. A method as claimed in any of the preceding claims, wherein the step of illuminating the portion of the retina with light (221) of the second wavelength comprises illuminating the portion of the retina with light of wavelengths centred on one of 635 nm, 670 nm and 760 nm.
 9. A method as claimed in any of the preceding claims, wherein step f) further comprises determining from the metabolic function changes a waveform reference point in a haemoglobin oxygenation image data waveform corresponding to a cardiac cycle.
 10. A method as claimed in claim 9, wherein the step of determining a waveform reference point comprises determining an amplitude trough between a trailing edge and a leading edge of an arterial pulse of the haemoglobin oxygenation image data waveform.
 11. A method as claimed in claims 9 or 10, for determining oxygenation changes associated with the cardiac cycle.
 12. A method as claimed in claim 11, for determining characteristics of at least one of amplitude, pattern, shape and duration of the haemoglobin oxygenation image data waveform.
 13. A method as claimed in claim 11 or 12, wherein step f) further comprises subtracting effects of changes in oxygenation caused by the cardiac cycle to determine metabolic function of the at least a portion of the retina independently of oxygenation changes caused by the cardiac cycle.
 14. A method as claimed in any of the preceding claims, including a further step of subjecting the at least a portion of the retina to optical stimulation and analysing effects of the optical stimulation on retinal metabolic function.
 15. A method as claimed in claim 14 when dependent on claim 9, wherein the optical stimulation of the at least a portion of the retina is synchronised with reference to the waveform reference point.
 16. A method as claimed in claim 15, comprising isolating retinal metabolic image changes in response to the optical stimulation by image subtraction from background functional retinal metabolic image data obtained over a heartbeat.
 17. A method as claimed in claim 15, comprising analysing a resulting isolated retinal metabolism response to the optical stimulation for one or more of latency between the optical stimulation and the metabolism response and the amplitude, pattern, shape and duration of a waveform of the response.

18. A method as claimed in claim 14 or 15, wherein step f) further comprises subtracting effects of changes in oxygenation caused by the cardiac cycle from the effects of changes in oxygenation caused by the optical stimulation to isolate changes in retinal metabolism caused by the optical stimulation. 5
19. A method as claimed in claim 18, comprising analysing isolated responses of retinal metabolism caused by the optical stimulation to determine waveform characteristics of at least one of latency between the optical stimulation and the retinal metabolism response, amplitude, pattern, shape and duration of the response. 10
20. A method as claimed in any of claims 14 to 19, wherein the optical signal stimulus light source is modulated to generate an optical signal stimulus pattern on the at least a portion of a retina. 15
21. A method as claimed in any of claims 14 to 20, wherein the step of providing an optical signal stimulus light source comprises providing an optical signal stimulus laser. 20
22. A retinal function camera (200) comprising: a first source (21) of light (211) of a first wavelength band; a second source (22) of light (221) of a second wavelength band, the absorptivity of light of the first wavelength band by oxygenated blood being greater than the absorptivity of light of the second wavelength band and the absorptivity of light of the first wavelength band by deoxygenated blood being less than the absorptivity of light of the second wavelength band; **Characterized by** further comprising an optical signal stimulus light source (40) distinct from the first source of light and the second source of light; means (90) for focusing light from the optical signal stimulus light source and selectively from the first and second sources onto a portion of a retina of an eye (10); imaging means (110) for producing respective images of the portion of the retina illuminated with the respective wavelength bands and stimulated by the optical signal stimulus light source (40); and processing means (150) adapted to process the respective images obtained by the imaging means to determine isorefective points of the respective images at which absorption of light of the first wavelength is substantially equal to absorption of light of the second wavelength and areas of the respective images having differential absorptivity for the first and second wavelengths, repeatedly to obtain a retinal function image based on haemoglobin oxygenation within at least a portion of a cardiac cycle to analyse metabolic functional changes of the portion of the retina to detect a haemoglobin oxygenation image data waveform within the at least a portion of a cardiac cycle to provide synchronisation with a cardiac cycle of a subject for analysing retinal metabolic image data over the at least a portion of the cardiac cycle when the retina is subjected to optical stimulation. 25
23. A retinal function camera as claimed in claim 22, wherein the optical signal stimulus light source (40) is an optical signal stimulus laser. 30
24. A retinal function camera as claimed in claim 22 or 23, further comprising scanning means (80) arranged for producing a scanning beam for scanning the portion of the retina with light from the optical signal stimulus light source (40) and selectively from the first and second sources (21, 22). 35
25. A retinal function camera as claimed in claim 24, further comprising first optical beam combiner means (30) arranged for selectively directing light from the first light source and from the second light source to the scanning means (80). 40
26. A retinal function camera as claimed in claims 24 or 25, further comprising first synchronisation means arranged for synchronising at least one of the first light source (21) and the second light source (22) with the scanning means (80). 45
27. A retinal function camera as claimed in any of claims 24 to 26, further comprising refractive error correcting means arranged for correcting for refractive errors of the eye to permit focusing of the scanning beam (801) on the at least a portion of the retina. 50
28. A retinal function camera as claimed in any of claims 22 to 27, further comprising beam splitter means (60) arranged for splitting a beam reflected from the at least a portion of the retina when illuminated by light (211) of the first wavelength band and light (221) of the second wavelength band to form a split beam; image sensor means and second focusing means (110) arranged for focusing the split beam on the image sensor means (130) for sensing an image of the at least a portion of the retina and image capture means (140) for capturing the sensed image. 55
29. A retinal function camera as claimed in claim 28, wherein the beam splitter means is arranged to direct light reflected from the retina to an orthogonal polarising filter (100) such that the polarising filter blocks light reflected from eye surfaces with substantially a same polarisation as the first light source (21) and the second light source (22).
30. A retinal function camera as claimed in claim 28, further comprising second synchronisation means arranged for synchronising at least one of the image sensor means (130), the image capture means (140)

and image processing means (150) with the scanning means (80).

31. A retinal function camera as claimed in any of claims 22 to 30, wherein the first wavelength band comprises wavelengths centred on one of 830 nm, 850 nm and 910 nm. 5
32. A retinal function camera as claimed in any of claims 22 to 31, wherein the second wavelength band comprises wavelengths centred on one of 635 nm, 670 nm and 760 nm. 10

Patentansprüche 15

1. Verfahren zum Analysieren des Netzhautstoffwechsels über zumindest einen Teil eines Herzzyklus, wobei das Verfahren die folgenden Schritte umfasst: 20

a) das Beleuchten eines Abschnitts der Netzhaut eines Auges (10) mit Licht einer ersten Wellenlänge durch Bereitstellen einer ersten Lichtquelle (21) und Emittieren von Licht (211) der ersten Wellenlänge und das Bereitstellen einer Abtasteinrichtung (80) zur Erzeugung eines Abtaststrahls (801) und das Abtasten des Abschnitts der Netzhaut mit Licht von der ersten Lichtquelle; 25

b) das Erzeugen eines ersten Bilds des Abschnitts der Netzhaut, der mit dem Licht der ersten Wellenlänge beleuchtet wird; 30

c) das Beleuchten des Abschnitts der Netzhaut mit Licht (221) einer zweiten Wellenlänge, wobei die ersten und zweiten Wellenlängen so ausgewählt werden, dass das Absorptionsvermögen des Lichts der ersten Wellenlänge durch mit Sauerstoff angereichertes Blut größer ist als das Absorptionsvermögen des Lichts der zweiten Wellenlänge, und wobei das Absorptionsvermögen des Lichts der ersten Wellenlänge durch deoxygeniertes Blut geringer ist als das Absorptionsvermögen des Lichts der zweiten Wellenlänge; 35

d) das Erzeugen eines zweiten Bilds des Abschnitts der Netzhaut, der mit dem Licht der zweiten Wellenlänge beleuchtet wird; wobei das Verfahren **dadurch gekennzeichnet ist, dass** es ferner die folgenden Schritte umfasst: 40

e) das Verarbeiten der ersten und zweiten Bilder mit einer Bildverarbeitungseinrichtung (150), um die relative Sauerstoffanreicherung des Abschnitts der Netzhaut als eine Anzeige der Stoffwechselfunktion der Netzhaut des Abschnitts der Netzhaut abzubilden, indem isoreflektierende Punkte der entsprechenden Bilder bestimmt werden, an denen die Absorption von Licht (211) der ersten Wellenlänge im Wesentlichen der Ab- 45

sorption von Licht (221) der zweiten Wellenlänge entspricht, und wobei Bereiche der entsprechenden Bilder bestimmt werden, die für die ersten und zweiten Wellenlängen unterschiedliche Absorptionsvermögenswerte aufweisen; wobei Hämoglobin-Sauerstoffanreicherungsbilddaten aus den Bildern der ersten und der zweiten Wellenlänge isoliert werden, indem das Reflexionsvermögen an dem isoreflektierenden Punkt von den entsprechenden Bildern der ersten und zweiten Wellenlänge subtrahiert wird, wobei der Kontrast jedes Abschnitts der verarbeiteten Bilder **dadurch** proportional ist zu einer Differenz zwischen der Sauerstoffanreicherung des Abschnitts der verarbeiteten Bilder und der Sauerstoffanreicherung des isoreflektierenden Punkts, so dass Netzhautstoffwechsel-Bilddaten auf der Basis der Hämoglobin-Sauerstoffanreicherung erhalten werden; und 50

f) das Wiederholen der Schritte a) bis e) innerhalb zumindest eines Teils eines Herzzyklus, wodurch Stoffwechselfunktionsveränderungen des Abschnitts der Netzhaut analysiert werden, wobei eine Hämoglobin-Sauerstoffanreicherungsbilddatenkurvenform mindestens in einem Abschnitt eines Herzzyklus detektiert wird, und wobei eine Synchronisation mit einem Herzzyklus eines Subjekts bereitgestellt wird, um Netzhautstoffwechsel-Bilddaten über zumindest den einen Abschnitt des Herzzyklus zu analysieren. 55

2. Verfahren nach Anspruch 1, wobei der Schritt des Beleuchtens des Abschnitts der Netzhaut mit Licht (221) einer zweiten Wellenlänge das folgende Bereitstellen umfasst: einer zweiten Lichtquelle (22) und das Emittieren von Licht der zweiten Wellenlänge; und einer ersten optischen Strahlverknüpfungseinrichtung (30) sowie das selektive Leiten von Licht (211) von der ersten Lichtquelle (21) und von der zweiten Lichtquelle (22) zu der Abtasteinrichtung (80), so dass ein Abtaststrahl (801) erzeugt wird, und das Abtasten des Abschnitts der Netzhaut mit Licht von der zweiten Lichtquelle nach dem Abtasten des Abschnitts der Netzhaut mit Licht von der ersten Lichtquelle. 60

3. Verfahren nach Anspruch 2, wobei die mindestens eine der ersten Lichtquelle (21) und der zweiten Lichtquelle (22) mit der Abtasteinrichtung (80) synchronisiert wird. 65

4. Verfahren nach einem der vorstehenden Ansprüche, wobei dieses einen Schritt des Bereitstellens einer ersten Fokussierungseinrichtung (90) und des Fokussierens des Abtaststrahls (801) von der Abtasteinrichtung (80) auf den mindestens einen Abschnitt der Netzhaut mit der ersten Fokussierungseinrichtung umfasst. 70

5. Verfahren nach Anspruch 4, wobei das Verfahren den Schritt des Bereitstellens einer Einrichtung zur Korrektur von Brechungsfehlern und zur Korrektur von Brechungsfehlern des Auges mit einer Einrichtung zur Korrektur von Brechungsfehlern aufweist, um das Fokussieren des Abtaststrahls auf den mindestens einen Abschnitt der Netzhaut zu ermöglichen. 5
6. Verfahren nach einem der vorstehenden Ansprüche, wobei der Schritt des Erzeugens erster und zweiter Bilder das Bereitstellen einer Strahlenteilereinrichtung (60) zum Teilen eines Strahls umfasst, der von dem mindestens einen Abschnitt der Netzhaut reflektiert wird, wenn dieser mit Licht (211) der ersten Wellenlänge und mit Licht (221) der zweiten Wellenlänge beleuchtet wird, so dass ein geteilter Strahl erzeugt wird; mit einer Bildsensoreinrichtung (130) und einer zweiten Fokussierungseinrichtung (110), und wobei der geteilte Strahl auf die Bildsensoreinrichtung fokussiert wird, um ein Bild des mindestens einen Abschnitts der Netzhaut zu messen, und mit einer Bilderfassungseinrichtung (140) zum Erfassen des gemessenen Bilds. 10 15 20 25
7. Verfahren nach einem der vorstehenden Ansprüche, wobei der Schritt des Beleuchtens des Abschnitts der Netzhaut mit Licht (211) der ersten Wellenlänge das Beleuchten des Abschnitts der Netzhaut mit Licht von Wellenlängen umfasst, die auf eine der Wellenlängen 830 nm, 850 nm und 910 nm zentriert ist. 30
8. Verfahren nach einem der vorstehenden Ansprüche, wobei der Schritt des Beleuchtens des Abschnitts der Netzhaut mit Licht (221) der zweiten Wellenlänge das Beleuchten des Abschnitts der Netzhaut mit Licht von Wellenlängen umfasst, die auf eine der Wellenlängen 635 nm, 670 nm und 760 nm zentriert ist. 35
9. Verfahren nach einem der vorstehenden Ansprüche, wobei der Schritt f) ferner aus Stoffwechselfunktionsveränderungen das Bestimmen eines Kurvenformreferenzpunkts in einer Hämoglobin-Sauerstoffanreicherungsbilddatenkurvenform entsprechend einem Herzzyklus umfasst. 40 45
10. Verfahren nach Anspruch 9, wobei der Schritt des Bestimmens eines Kurvenformreferenzpunkts das Bestimmen eines Amplitudentiefpunkts zwischen einer abfallenden Flanke und einer Anstiegsflanke eines arteriellen Pulses der Hämoglobin-Sauerstoffanreicherungsbilddatenkurvenform umfasst. 50
11. Verfahren nach Anspruch 9 oder 10 zum Bestimmen von Stoffwechselveränderungen, die dem Herzzyklus zugeordnet sind. 55
12. Verfahren nach Anspruch 11 zur Bestimmung von mindestens einer der folgenden Eigenschaften: der Amplitude, des Musters, der Form oder der Dauer der Hämoglobin-Sauerstoffanreicherungsbilddatenkurvenform.
13. Verfahren nach einem der Ansprüche 11 oder 12, wobei der Schritt f) ferner das Subtrahieren von Effekten von Veränderungen der Sauerstoffanreicherung bewirkt durch den Herzzyklus umfasst, um die Stoffwechselfunktion des mindestens einen Abschnitts der Netzhaut unabhängig von durch den Herzzyklus verursachten Stoffwechselveränderungen zu bestimmen.
14. Verfahren nach einem der vorstehenden Ansprüche, wobei das Verfahren ferner den Schritt des Aussetzens des mindestens einen Abschnitts der Netzhaut optischen Stimulation umfasst sowie das Analysieren der Effekte der optischen Stimulation auf die Stoffwechselfunktion der Netzhaut.
15. Verfahren nach Anspruch 14, wenn dieses von Anspruch 9 abhängig ist, wobei die optische Stimulation des mindestens einen Abschnitts der Netzhaut synchronisiert wird in Bezug auf den Kurvenformreferenzpunkt.
16. Verfahren nach Anspruch 15, wobei das Verfahren das Isolieren von Netzhautstoffwechselbildveränderungen als Reaktion auf die optische Stimulation durch Bildsubtraktion von den funktionalen Hintergrund-Netzhautstoffwechselbilddaten, die über einen Herzschlag erhalten werden, umfasst.
17. Verfahren nach Anspruch 15, wobei das Verfahren ferner das Analysieren einer resultierenden isolierten Netzhautstoffwechselreaktion auf die optische Stimulation für eine oder mehrere Latenzen zwischen der optischen Stimulation und der Stoffwechselreaktion sowie der Amplitude, dem Muster, der Form und der Dauer einer Kurvenform der Reaktion umfasst.
18. Verfahren nach Anspruch 14 oder 15, wobei der Schritt f) ferner das Subtrahieren von Effekten von Veränderungen der Sauerstoffanreicherung bewirkt durch den Herzzyklus durch die Auswirkungen der Veränderungen der Sauerstoffanreicherung bewirkt durch die optische Stimulation umfasst, um die Veränderungen des Netzhautstoffwechsels zu isolieren, die durch die optische Stimulation verursacht werden.
19. Verfahren nach Anspruch 18, wobei das Verfahren das Analysieren der isolierten Reaktionen des Netzhautstoffwechsels umfasst, verursacht durch die optische Stimulation, um Kurvenformeingenschaften

mindestens einer der Latenzen zwischen der optischen Stimulation und der Netzhautstoffwechselreaktion, der Amplitude, dem Muster, der Form und der Dauer der Reaktion umfasst.

20. Verfahren nach einem der Ansprüche 14 bis 19, wobei die optische Signalreizlichtquelle so moduliert wird, dass ein optisches Signalreizmuster an dem mindestens einen Abschnitt einer Netzhaut erzeugt wird.
21. Verfahren nach einem der Ansprüche 14 bis 20, der Schritt des Bereitstellens einer optischen Signalreizlichtquelle das Bereitstellen eines optischen Signalreizlasers umfasst.
22. Netzhautfunktionskamera (200), die folgendes umfasst: eine erste Quelle (21) für Licht (211) eines ersten Wellenlängenbands; eine zweite Quelle (22) für Licht (221) eines zweiten Wellenlängenbands, wobei das Absorptionsvermögen für Licht des ersten Wellenlängenbands von mit Sauerstoff angereicherter Blut größer ist als das Absorptionsvermögen von Licht des zweiten Wellenlängenbands, und wobei das Absorptionsvermögen von Licht des ersten Wellenlängenbands von deoxygeniertem Blut geringer ist als das Absorptionsvermögen von Licht des zweiten Wellenlängenbands; **dadurch gekennzeichnet, dass** die Kamera ferner folgendes umfasst: eine optische Signalreizlichtquelle (40), die sich von der ersten Lichtquelle und von der zweiten Lichtquelle unterscheidet; eine Einrichtung (90) zum Fokussieren von Licht von der optischen Signalreizlichtquelle und selektiv von den ersten und zweiten Quellen auf einen Abschnitt der Netzhaut eines Auges (10); eine Bilddarstellungseinrichtung (110) zur Erzeugung entsprechender Bilder des Abschnitts der Netzhaut, der mit den entsprechenden Wellenlängenbändern beleuchtet und durch die optische Signalreizlichtquelle (40) stimuliert wird; und eine Verarbeitungseinrichtung (150), die die entsprechenden Bilder verarbeiten kann, die von der Bilddarstellungseinrichtung erhalten werden, um isoreflektierende Punkte der entsprechenden Bilder zu bestimmen, an denen die Absorption von Licht der ersten Wellenlänge im Wesentlichen der Absorption von Licht der zweiten Wellenlänge und von Bereichen der entsprechenden Bilder mit differenziellem Absorptionsvermögen für die ersten und zweiten Wellenlängen entspricht, und zwar wiederholt, um ein Netzhautfunktionsbild auf der Basis der Hämoglobin-Sauerstoffanreicherung in mindestens einem Teil des Herzzyklus zu erhalten, um Stoffwechselfunktionsveränderungen des Abschnitts der Netzhaut zu analysieren, um eine Hämoglobin-Sauerstoffanreicherungsbilddatenkurvenform in dem mindestens einen Abschnitt eines Herzzyklus zu detektieren, um eine Synchronisation mit einem Herzzyklus eines Subjekts bereitzustellen.

len, um Netzhautstoffwechsel-Bilddaten über den mindestens einen Abschnitt des Herzzyklus zu analysieren, wenn die Netzhaut optischer Stimulation ausgesetzt wird.

- 5
23. Netzhautfunktionskamera nach Anspruch 22, wobei die optische Signalreizlichtquelle (40) einen optischen Signalreizlaser darstellt.
- 10
24. Netzhautfunktionskamera nach Anspruch 22 oder 23, wobei die Kamera ferner eine Abtasteinrichtung (80) umfasst, die so angeordnet ist, dass sie einen Abtaststrahl zum Abtasten des Abschnitts der Netzhaut mit Licht von der optischen Signalreizlichtquelle (40) und selektiv von den ersten und zweiten Quellen (21, 22) erzeugt.
- 15
25. Netzhautfunktionskamera nach Anspruch 24, wobei die Kamera ferner eine erste optische Strahlverknüpfungseinrichtung (30) umfasst, die so angeordnet ist, dass sie selektiv Licht von der ersten Lichtquelle und von der zweiten Lichtquelle zu der Abtasteinrichtung (80) leitet.
- 20
26. Netzhautfunktionskamera nach Anspruch 24 oder 25, wobei die Kamera ferner eine erste Synchronisationseinrichtung umfasst, die so angeordnet ist, dass sie mindestens eine der ersten Lichtquelle (21) oder der zweiten Lichtquelle (22) mit der Abtasteinrichtung (80) umfasst.
- 25
27. Netzhautfunktionskamera nach einem der Ansprüche 24 bis 26, wobei die Kamera ferner eine Einrichtung zur Korrektur von Brechungsfehlern umfasst, die so angeordnet ist, dass sie Brechungsfehler des Auges korrigiert, um die Fokussierung des Abtaststrahls (801) an dem mindestens einen Abschnitt der Netzhaut zu ermöglichen.
- 30
- 35
- 40
- 45
- 50
- 55
28. Netzhautfunktionskamera nach einem der Ansprüche 22 bis 27, wobei die Kamera ferner eine Strahlenteilerinrichtung (60) umfasst, die so angeordnet ist, dass ein von dem mindestens einen Abschnitt der Netzhaut reflektierter Strahl geteilt wird, wenn der Bereich durch Licht (211) des ersten Wellenlängenbands und durch Licht (221) des zweiten Wellenlängenbands beleuchtet wird, so dass ein geteilter Strahl erzeugt wird; mit einer Bildsensoreinrichtung und einer zweiten Fokussierungseinrichtung (110), die so angeordnet ist, dass der geteilte Strahl an der Bildsensoreinrichtung (130) fokussiert wird, um ein Bild des mindestens einen Abschnitts der Netzhaut zu messen, und mit einer Bilderfassungseinrichtung (140) zum Erfassen des gemessenen Bilds.
29. Netzhautfunktionskamera nach Anspruch 28, wobei die Strahlenteilerinrichtung so angeordnet ist, dass

sie von der Netzhaut reflektiertes Licht zu einem orthogonalen Polarisationsfilter (100) leitet, so dass der Polarisationsfilter von Augenoberflächen reflektiertes Licht im Wesentlichen mit der gleichen Polarisation wie die erste Lichtquelle (21) und die zweite Lichtquelle (22) polarisiert.

30. Netzhautfunktionskamera nach Anspruch 28, wobei die Kamera ferner eine zweite Synchronisationseinrichtung umfasst, die so angeordnet ist, dass sie mindestens eine der Bildsensoreinrichtung (130), der Bilderfassungseinrichtung (140) oder der Bildverarbeitungseinrichtung (150) mit der Abtasteinrichtung (80) synchronisiert.
31. Netzhautfunktionskamera nach einem der Ansprüche 22 bis 30, wobei das erste Wellenlängenband Wellenlängen umfasst, die auf eine der Wellenlängen 830 nm, 850 nm oder 910 nm zentriert sind.
32. Netzhautfunktionskamera nach einem der Ansprüche 22 bis 31, wobei das zweite Wellenlängenband Wellenlängen umfasst, die auf eine der Wellenlängen 635 nm, 670 nm oder 760 nm zentriert sind.

Revendications

1. Procédé d'analyse du métabolisme rétinien sur au moins une partie d'un cycle cardiaque, le procédé comprenant les étapes consistant à :
- a. éclairer une partie d'une rétine d'un oeil (10) avec la lumière d'une première longueur d'onde en fournissant une première source lumineuse (21) et en émettant la lumière (211) de la première longueur d'onde et fournir des moyens de balayage (80) pour produire un faisceau de balayage (801) et balayer la partie de la rétine avec la lumière provenant de la première source lumineuse ;
- b. produire une première image de la partie de la rétine éclairée avec la lumière de la première longueur d'onde ;
- c. éclairer la partie de la rétine avec la lumière (221) d'une seconde longueur d'onde, les première et seconde longueurs d'onde étant sélectionnées de sorte que l'absorptivité de la lumière de la première longueur d'onde par le sang oxygéné est supérieure à l'absorptivité de la lumière de la seconde longueur d'onde et que l'absorptivité de la lumière de la première longueur d'onde par le sang désoxygéné est inférieure à l'absorptivité de la lumière de la seconde longueur d'onde ;
- d. produire une seconde image de la partie de la rétine éclairée avec la lumière de la seconde longueur d'onde ; le procédé étant **caractérisé**

en ce qu'il comprend en outre les étapes consistant à :

e. traiter les première et seconde images avec des moyens de traitement d'image (150) pour mapper l'oxygénation relative de la partie de la rétine comme une indication de la fonction métabolique rétinienne de la partie de la rétine en déterminant des points isorélecteurs des images respectives auxquels l'absorption de lumière (211) de la première longueur d'onde est sensiblement égale à l'absorption de lumière (211) de la seconde longueur d'onde et déterminer des zones des images respectives ayant une absorptivité différentielle pour les première et seconde longueurs d'onde ; isoler les données d'image d'oxygénation d'hémoglobine des première et seconde images de longueur d'onde en soustrayant la réflectivité au point isorélecteur des première et seconde images de longueur d'onde respectives, le contraste de chaque partie des images traitées étant ainsi proportionnel à une différence en oxygénation de cette partie des images traitées par rapport à l'oxygénation du point isorélecteur, pour obtenir les données d'image métabolique rétinienne basées sur l'oxygénation de l'hémoglobine ; et

f. répéter les étapes a) à e) dans l'au moins une partie d'un cycle cardiaque pour analyser ainsi les changements de la fonction métabolique de la partie de la rétine, détecter un signal de données d'image d'oxygénation d'hémoglobine dans l'au moins une partie d'un cycle cardiaque et fournir une synchronisation avec un cycle cardiaque d'un sujet pour analyser les données d'image métabolique rétinienne sur l'au moins une partie du cycle cardiaque.

2. Procédé selon la revendication 1, dans lequel l'étape d'éclairage de la partie de la rétine avec la lumière (221) d'une seconde longueur d'onde comprend les étapes consistant à fournir une seconde source de lumière (22) et à émettre la lumière de la seconde longueur d'onde et des premiers moyens de combinaison de faisceau optique (30) et diriger sélectivement la lumière (211) de la première source de lumière (21) et de la seconde source de lumière (22) vers les moyens de balayage (80) pour produire un faisceau de balayage (801) et balayer la partie de la rétine avec la lumière provenant de la seconde source de lumière subséquentement au balayage de la partie de la rétine avec la lumière provenant de la première source de lumière.
3. Procédé selon la revendication 2, dans lequel au moins l'une de la première source de lumière (21) et de la seconde source de lumière (22) est synchronisée avec les moyens de balayage (80).

4. Procédé selon l'une quelconque des revendications précédentes, incluant une étape consistant à fournir des premiers moyens de focalisation (90) et à focaliser le faisceau de balayage (801) provenant des moyens de balayage (80) sur l'au moins une partie de la rétine avec les premiers moyens de focalisation.
5. Procédé selon la revendication 4, incluant une étape consistant à fournir des premiers moyens de correction d'erreur réfractive et à corriger les erreurs réfractives de l'oeil avec les moyens de correction d'erreur réfractive, pour permettre la focalisation du faisceau de balayage sur l'au moins une partie de la rétine.
6. Procédé selon l'une quelconque des revendications précédentes, dans lequel l'étape de production de première et seconde images inclut les étapes consistant à fournir des moyens de séparation de faisceau (60) pour séparer un faisceau reflété depuis l'au moins une partie de la rétine en cas d'éclairage par la lumière (211) de la première longueur d'onde (221) et par la lumière (221) de la seconde longueur d'onde pour former un faisceau séparé ; des moyens capteurs d'image (130) et des seconds moyens de focalisation (110) et à focaliser le faisceau séparé sur les moyens capteurs d'image pour capter une image de l'au moins une partie de la rétine et des moyens de capture d'image (140) pour capturer l'image captée.
7. Procédé selon l'une quelconque des revendications précédentes, dans lequel l'étape d'éclairage de la partie de la rétine avec la lumière (211) de la première longueur d'onde comprend l'étape consistant à éclairer la partie de la rétine avec la lumière de longueurs d'onde centrée sur l'un de 830 nm, 850 nm et 910 nm.
8. Procédé selon l'une quelconque des revendications précédentes, dans lequel l'étape d'éclairage de la partie de la rétine avec la lumière (211) de la seconde longueur d'onde comprend l'étape consistant à éclairer la partie de la rétine avec la lumière de longueurs d'onde centrée sur l'un de 635 nm, 670 nm et 760 nm.
9. Procédé selon l'une quelconque des revendications précédentes, dans lequel l'étape f) comprend en outre l'étape consistant à déterminer à partir des changements de fonction métabolique un point de référence de signal dans un signal de données d'image d'oxygénation d'hémoglobine correspondant à un cycle cardiaque.
10. Procédé selon la revendication 9, dans lequel l'étape de détermination d'un point de référence de signal comprend l'étape consistant à déterminer une amplitude entre un front descendant et un front montant d'un pouls artériel du signal de données d'image d'oxygénation d'hémoglobine.
11. Procédé selon la revendication 9 ou 10, pour déterminer les changements d'oxygénation associés au cycle cardiaque.
12. Procédé selon la revendication 11, pour déterminer les caractéristiques d'au moins l'un d'une amplitude, d'un motif, d'une forme et d'une durée du signal de données d'image d'oxygénation d'hémoglobine.
13. Procédé selon la revendication 11 ou 12, dans lequel l'étape f) comprend en outre l'étape consistant à soustraire les effets des changements dans l'oxygénation causés par le cycle cardiaque pour déterminer la fonction métabolique de l'au moins une partie de la rétine indépendamment des changements d'oxygénation causés par le cycle cardiaque.
14. Procédé selon l'une quelconque des revendications précédentes, comprenant en outre l'étape consistant à soumettre l'au moins une partie de la rétine à une stimulation optique et à analyser les effets de la stimulation optique sur la fonction métabolique rétinienne.
15. Procédé selon la revendication 14 telle que dépendante de la revendication 9, dans lequel la stimulation optique de l'au moins une partie de la rétine est synchronisée en référence au point de référence de signal.
16. Procédé selon la revendication 15, comprenant l'étape consistant à isoler les changements d'image métabolique rétinienne en réponse à la stimulation optique par soustraction d'image depuis des données d'image métabolique rétinienne fonctionnelle d'arrière-plan obtenues sur un battement du coeur.
17. Procédé selon la revendication 15, comprenant l'étape consistant à analyser une réponse de métabolisme rétinien isolé résultante à la stimulation optique pour une ou plusieurs latences entre la stimulation optique et la réponse de métabolisme et l'amplitude, le motif, la forme et la durée d'un signal de la réponse.
18. Procédé selon la revendication 14 ou 15, dans lequel l'étape f) comprend en outre l'étape consistant à soustraire les effets des changements en oxygénation causés par le cycle cardiaque à partir des effets des changements en oxygénation causés par la stimulation optique pour isoler les changements en métabolisme rétinien causés par la stimulation optique.
19. Procédé selon la revendication 18, comprenant l'étape

- pe consistant à analyser les réponses isolées de métabolisme rétinien causées par la stimulation optique pour déterminer les caractéristiques de signal d'au moins l'une des latences entre la stimulation optique et la réponse de métabolisme rétinien, l'amplitude, le motif, la forme et la durée de la réponse.
- 5
20. Procédé selon l'une quelconque des revendications 14 à 19, dans lequel la source de lumière de stimulus de signal optique est modulée pour générer un motif de stimulus de signal optique sur l'au moins une partie d'une rétine.
- 10
21. Procédé selon l'une quelconque des revendications 14 à 20, dans lequel l'étape de fourniture d'une source de lumière de stimulus de signal optique comprend l'étape consistant à fournir un laser de stimulus de signal optique.
- 15
22. Caméra à fonction rétinienne (200) comprenant :
 une première source (21) de lumière (211) d'une première bande de longueur d'onde ; une seconde source (22) de lumière (221) d'une seconde bande de longueur d'onde, l'absorptivité de lumière de la première bande de longueur d'onde par le sang oxygéné étant supérieure à l'absorptivité de la lumière de la seconde bande de longueur d'onde et l'absorptivité de la lumière de la première bande de longueur d'onde par le sang désoxygéné étant inférieure à l'absorptivité de la lumière de la seconde bande de longueur d'onde ; **caractérisée en ce qu'elle** comprend en outre une source de lumière de stimulus de signal optique (40) différente de la première source de lumière et de la seconde source de lumière ; des moyens (90) pour focaliser la lumière provenant de la source de lumière de stimulus de signal optique et sélectivement des première et seconde sources sur une partie d'une rétine d'un oeil (10) ; des moyens d'imagerie (110) pour produire des images respectives de la partie de la rétine éclairée avec les bandes de longueur d'onde respectives et stimulés par la source de lumière de stimulus de signal optique (40) ; et des moyens de traitement (150) adaptés pour traiter les images respectives obtenues par les moyens d'imagerie pour déterminer les points isorélecteurs des images respectives auxquels l'absorption de lumière de la première longueur d'onde est sensiblement égale à l'absorption de lumière de la seconde longueur d'onde et les zones des images respectives ayant une absorptivité différentielle pour les première et seconde longueurs d'onde, de façon répétitive pour obtenir une image de fonction rétinienne basée sur l'oxygénation d'hémoglobine dans au moins une partie d'un cycle cardiaque pour analyser les changements fonctionnels métaboliques de la partie de la rétine pour détecter un signal de données d'image d'oxygénation d'hémoglobine dans l'au moins une partie d'un cycle cardiaque pour four-
- 20
- 25
- 30
- 35
- 40
- 45
- 50
- 55
- nir la synchronisation avec un cycle cardiaque d'un sujet pour analyser les données d'image métabolique rétinienne sur l'au moins une partie du cycle cardiaque lorsque la rétine est soumise à une stimulation optique.
23. Caméra à fonction rétinienne selon la revendication 22, dans lequel la source de lumière de stimulus de signal optique (40) est un laser de stimulus de signal optique.
24. Caméra à fonction rétinienne selon la revendication 22 ou 23, comprenant en outre des moyens de balayage (80) agencés pour produire un faisceau de balayage pour balayer la partie de la rétine avec la lumière provenant de la source de lumière de stimulus de signal optique (40) et provenant sélectivement des première et seconde sources (21, 22).
25. Caméra à fonction rétinienne selon la revendication 24, comprenant en outre des premiers moyens combineurs de faisceau optique (30) agencés pour diriger sélectivement la lumière provenant de la première source de lumière et de la seconde source de lumière vers les moyens de balayage (80).
26. Caméra à fonction rétinienne selon la revendication 24 ou 25, comprenant en outre des premiers moyens de synchronisation pour synchroniser au moins l'une de la première source de lumière (21) et de la seconde source de lumière (22) avec les moyens de balayage (80).
27. Caméra à fonction rétinienne selon l'une quelconque des revendications 24 à 26, comprenant en outre des premiers moyens de correction d'erreur réfractive agencés pour corriger les erreurs réfractives de l'oeil pour permettre la focalisation du faisceau de balayage (801) sur l'au moins une partie de la rétine.
28. Caméra à fonction rétinienne selon l'une quelconque des revendications 22 à 27, comprenant en outre des moyens de séparation de faisceau (60) agencés pour séparer un faisceau reflété depuis l'au moins une partie de la rétine en cas d'éclairage par la lumière (211) de la première bande de longueur d'onde et la lumière (221) de la seconde bande de longueur d'onde pour former un faisceau séparé ; des moyens capteurs d'image et des seconds moyens de focalisation (110) agencés pour focaliser le faisceau séparé sur les moyens capteurs d'image (130) pour capter une image de l'au moins une partie de la rétine et des moyens de capture d'image (140) pour capturer l'image captée.
29. Caméra à fonction rétinienne selon la revendication 28, dans laquelle les moyens séparateurs de faisceau sont agencés pour diriger la lumière reflétée

depuis la rétine sur un filtre polarisant orthogonal (100) de sorte que le filtre polarisant bloque la lumière réfléctée des surfaces de l'oeil avec sensiblement une même polarisation comme la première source de lumière (21) et la seconde source de lumière (22). 5

30. Caméra à fonction rétinienne selon la revendication 28, comprenant en outre des seconds moyens de synchronisation agencés pour synchroniser au moins l'un des moyens capteurs d'image (130), des moyens de capture d'image (140) et des moyens de traitement d'image (150) avec les moyens de balayage (80). 10
31. Caméra à fonction rétinienne selon l'une quelconque des revendications 22 à 30, dans lequel la première bande de longueur d'onde comprend des longueurs d'onde centrées sur l'un de 830 nm, 850 nm et 910 nm. 15
20
32. Caméra à fonction rétinienne selon l'une quelconque des revendications 22 à 31, dans lequel la seconde bande de longueur d'onde comprend des longueurs d'onde centrées sur l'un de 635 nm, 670 nm et 760 nm. 25

30

35

40

45

50

55

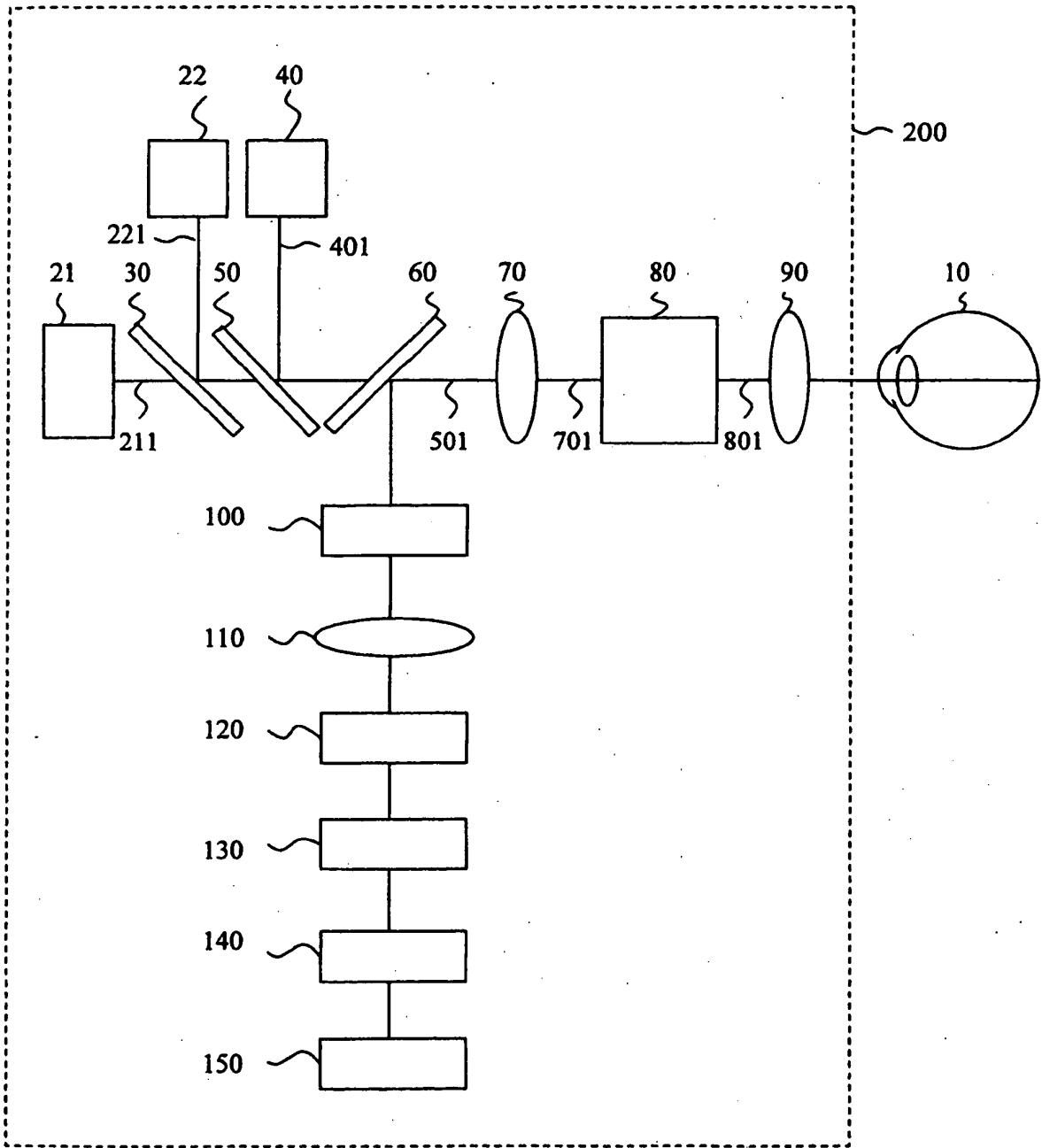


Figure 1

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 6244712 B [0008]
- US 20020188203 A [0009]
- US 20040075812 A [0011]
- WO 02080759 A [0013]

Non-patent literature cited in the description

- **R. H. WEBB ; G. W. HUGHES ; O. POMERANTZ-EFF.** Flying spot TV ophthalmoscope. *Applied Optics*, 1980, vol. 19, 2991-2997 [0004]
- **ANN E ELSER ; STEPHEN A. BURNS ; JOHN J WEITTER ; FRANCOIS C DELORI.** Infrared imaging of sub-retinal structures in the human ocular fundus. *Vision Res.*, 1996, vol. 36 (1), 191-205 [0005]

专利名称(译)	分析至少一部分心动周期的视网膜代谢		
公开(公告)号	EP1611840B1	公开(公告)日	2008-08-13
申请号	EP2005253758	申请日	2005-06-16
[标]申请(专利权)人(译)	KERR PATRICK		
申请(专利权)人(译)	KERR , PATRICK		
当前申请(专利权)人(译)	KERR , PATRICK		
[标]发明人	KERR PATRICK		
发明人	KERR, PATRICK		
IPC分类号	A61B5/00 A61B3/12 A61B5/021		
CPC分类号	A61B3/1015 A61B3/1225 A61B5/021 A61B5/14555		
优先权	2004014570 2004-06-29 GB		
其他公开文献	EP1611840A1		
外部链接	Espacenet		

摘要(译)

通过首先用第一波长的光照射眼睛10的视网膜的一部分并产生第一图像，在心动周期的至少一部分上用视网膜功能相机分析视网膜代谢。随后用第二波长的光照射视网膜的部分，选择第一和第二波长，使得氧合血液对第一波长的光的吸收率大于第二波长的光的吸收率和第二波长的光的吸收率。通过脱氧血液的第一波长小于第二波长的光的吸收率，以产生第二图像。处理第一和第二图像以绘制视网膜部分的相对氧合作为视网膜部分的视网膜代谢功能的指示。在心动周期的至少一部分上重复该过程，以分析心动周期的至少一部分内的视网膜部分的代谢功能变化。在一些实施方案中，对视网膜的至少一部分进行光学刺激，并且分析光学刺激对视网膜代谢功能的影响。

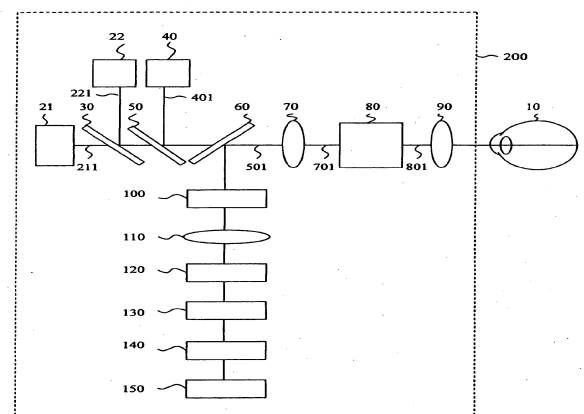


Figure 1