



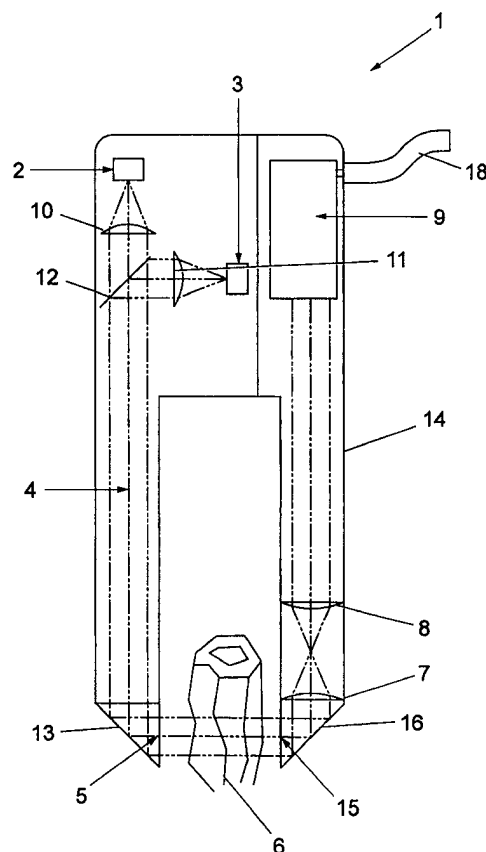
INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

<p>(51) International Patent Classification ⁶ : A61B 5/00</p>	<p>A1</p>	<p>(11) International Publication Number: WO 98/52460 (43) International Publication Date: 26 November 1998 (26.11.98)</p>
<p>(21) International Application Number: PCT/GB98/01524 (22) International Filing Date: 26 May 1998 (26.05.98) (30) Priority Data: 9710561.3 23 May 1997 (23.05.97) GB (71) Applicant (for all designated States except US): MEDICAL LASER TECHNOLOGIES LIMITED [GB/GB]; Unit 4, Belleknowes Industrial Estate, Inverkeithing, Fife KY11 1HY (GB). (72) Inventor; and (75) Inventor/Applicant (for US only): COLLES, Michael, John [GB/GB]; Boglesknowe, Hartree by Biggar, Lanarkshire ML12 6JJ (GB). (74) Agent: MURGITROYD & COMPANY; 373 Scotland Street, Glasgow G5 8QA (GB).</p>		<p>(81) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG).</p> <p>Published <i>With international search report. Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.</i></p>

(54) Title: NON-INVASIVE DIAGNOSTIC METHOD AND APPARATUS

(57) Abstract

Two light sources (2, 3; 20, 22) of wavelengths within the region 1.1 to 1.6 μm, modulated at the same frequency but in antiphase to each other, are transmitted through a tooth (6; 24). Transmitted light is measured by a detector (9; 26) provided with an electronic filter set (28) to pass only the modulated frequency signal corresponding to differences in absorption between the two wavelengths. The differential absorption detection serves as a basis for caries detection. Scatter of the light in tissue is offset by the use of optical spatial filtering.



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1

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3 Non-invasive diagnostic method and apparatus

4

5 This invention relates to a non-invasive diagnostic
6 method and apparatus and in particular to a method of
7 and apparatus for two-dimensional imaging by
8 differential absorption detection.

9

10 Most particularly this invention relates to an infra-
11 red imaging system which provides visualisation of
12 hidden tooth structures for, for example, the detection
13 and identification of dental hard tissue disease in
14 dentistry and oral medicine.

15

16 Dental hard tissue decay or caries, although reduced by
17 the modern use of fluorides in toothpastes or tablets
18 and an intensive programme of dental health education,
19 remains a major cause of tooth loss. The
20 identification of caries at its extreme is through pain
21 to the patient and a subsequent visit to the dentist.
22 At this stage the disease is usually extensive and its
23 cure requires substantial drilling and removal of both
24 diseased and sound tissue (the latter to gain
25 appropriate access to disease underlying enamel),

1 before filling.

2

3 At a regular dental examination caries may be detected
4 at a somewhat earlier stage by visual or more often
5 mechanical means. The latter, basically probing with a
6 pointed instrument, can find regions where overlying
7 enamel has softened but not yet fully decomposed. It
8 is less likely that detection will be possible when the
9 site is at a point of contact or near contact between
10 adjacent teeth. Unfortunately, since this region is
11 the hardest to maintain free from lodged food
12 particles, this is the most likely site for decay.

13

14 Earlier detection can be achieved with X-ray
15 examination. This technique is, however, not
16 straightforward when applied to caries detection since
17 the discrimination levels are low and shadowing effects
18 require both careful initial set-up and reasonably
19 expert interpretation. Nevertheless the technique does
20 provide for the earliest detection of caries and for
21 detection at sites that are visually and mechanically
22 hidden.

23

24 Various other techniques have been investigated at a
25 research level. These include visible light
26 transillumination and fluorescence detection. No
27 device with imaging potential has been made available
28 based on these techniques, a reflection of the
29 difficulties of obtaining adequate discrimination or
30 fluorescence site localisation.

31

32 Thus the earliest detection of dental disease is
33 available through X-ray imaging. This has the
34 disadvantage of requiring both very careful initial
35 setting up and very careful interpretation. In
36 addition, although dental X-ray images are taken at

1 very low fluxes, the radiation is still, of course,
2 ionising, and therefore carries a potential safety
3 hazard.

4

5 It is an object of the present invention to provide a
6 method and apparatus for imaging of dental tissue which
7 overcomes the above disadvantages.

8

9 According to a first aspect of the present invention
10 there is provided a method of two-dimensional imaging
11 of tissue by differential absorption detection
12 comprising the steps of transmitting radiation of two
13 wavelengths through tissue, detecting transmitted
14 radiation, measuring the absorption of said transmitted
15 radiation, and comparing absorption of said two
16 wavelengths.

17

18 Preferably the radiation has wavelengths of size
19 sufficient to preclude scattering in tissue. The
20 radiation is preferably infra-red light. More
21 preferably the light comprises two wavelengths within
22 the range 1.1 to 1.6 micrometres.

23

24 The first wavelength is preferably greater than 1.4
25 micrometres, most preferably within the range 1.4 to
26 1.5 micrometres, and may show a dominant change in
27 transmission characteristics.

28

29 The second wavelength is preferably less than 1.4
30 micrometres, most preferably within the range 1.2 to
31 1.3 micrometres, and may act as a reference.

32

33 Preferably the two wavelengths are modulated at the
34 same frequency. More preferably the two wavelengths
35 are in antiphase.

36

1 The method may include detection of the transmitted
2 radiation by optical means. Preferably the method
3 includes the step of filtering out all transmitted
4 radiation not at the modulated frequency.

5
6 The method may include the step of restricting
7 detection to radiation transmitted in a substantially
8 direct path through the tissue. Said restriction may
9 be by optical spatial filtering.

10
11 Preferably the method is applied to the detection of
12 dental caries. Alternatively the method is applied to
13 distinguish between different dental structures such as
14 enamel, dentine, root canals and pulp chambers.

15
16 Further according to the present invention there is
17 provided an apparatus for differential absorption
18 detection comprising two modulated light sources, each
19 source producing light of a different wavelength, optic
20 transmission means to train the light from said light
21 sources to illuminate a first side of a tissue sample
22 *in vivo*, and optic detection means to detect light
23 transmitted through said tissue sample to a second side
24 of said tissue sample.

25
26 Preferably the apparatus further includes means to
27 combine the output of said sources to provide a single
28 beam of incident light.

29
30 More preferably the optic detection means includes an
31 electronic filter.

32
33 The apparatus may comprise means to spatially filter
34 transmitted light.

35
36 Preferably the apparatus comprises means to display

1 detected transmitted light as an image.

2

3 Further according to the present invention there is
4 provided an apparatus for detection of dental caries
5 and/or visualisation of dental structures by
6 differential absorption detection comprising two
7 modulated light sources, optic means to combine light
8 from said sources to train a single beam on a first
9 side of a tooth, receiving means on a second side of
10 the tooth for receiving light transmitted through the
11 tooth, an optic filter to spatially filter the
12 transmitted light, and optic detection means to detect
13 the spatially filtered light.

14

15 The apparatus may comprise means to display detected
16 transmitted light as an image of the tooth. More
17 preferably the apparatus comprises means to display the
18 image in real time. The apparatus may comprise means
19 to electronically enhance the image. The apparatus may
20 comprise means to produce hard copies of said image.

21

22 Embodiments of the present invention will now be
23 described by way of example only with reference to the
24 accompanying drawings in which:

25

26 Figure 1 is a comparative graphic representation
27 of transmission characteristics of light through
28 healthy and decayed dental tissue;

29

30 Figure 2a is a schematic drawing of a laboratory
31 arrangement of a differential absorption detection
32 device in accordance with an aspect of the present
33 invention;

34

35 Figure 2b is a schematic drawing of a laboratory
36 arrangement of a differential absorption detection

1 device in accordance with a further aspect of the
2 present invention; and

3

4 Figure 3 is a schematic drawing of a differential
5 absorption detection device in accordance with an
6 aspect of the present invention.

7

8 Referring to the drawings, dental decay can give rise
9 to significant changes in the transmission
10 characteristics of light through enamel and dentine.
11 These changes occur in a spectral region not previously
12 investigated in the prior art and are substantial
13 enough to provide the basis for the selective imaging
14 of tooth decay.

15

16 These changes result from the breakdown of the
17 inorganic structures associated with dental hard
18 tissue, and their replacement with organic materials
19 having a higher water content. This gives rise to
20 significant changes in absorption as illustrated in
21 Figure 1, in which line 40 shows the relative
22 transmission of light through a sample of caries over a
23 range of wavelengths, while line 42 shows the relative
24 transmission of light through a sample of dentine over
25 the same range.

26

27 For light having a wavelength of between 1000 and
28 1400nm, caries exhibits higher relative transmission,
29 and hence lower absorption than dentine, while between
30 1400 and 1500nm, caries exhibits lower relative
31 transmission. The crossover point 44 occurs at about
32 1400nm.

33

34 This change in absorption is, however, only one aspect
35 of what is required to realise an effective image.

36 Figure 1 refers to spectra recorded through relatively

1 thin sections of teeth.

2

3 In practice there are substantial variations in:

4 (i) the thickness of teeth,

5 (ii) the thickness of various elements of a tooth such
6 as the enamel and the dentine, and

7 (iii) the direction of light incident on the tooth from
8 a given source point outside that tooth.

9

10 All of these variations act to disguise the smaller
11 variations due to the presence of caries. Thus
12 examining the tooth at a single wavelength only
13 provides image detection of caries under the most
14 favourable of circumstances.

15

16 This approach examines the difference in transmission
17 between two wavelengths in the same region one of which
18 shows the dominant changes due to caries, and the other
19 of which acts as a less strongly affected reference.

20

21 From Figure 1 it is clear that these wavelengths are in
22 the region 1.1 to 1.6 μm , and are most advantageously
23 chosen to be between 1.4 and 1.5 μm , and between 1.2 and
24 1.3 μm .

25

26 However when detecting other tissue it may be found
27 that the absorption characteristics of the tissue mean
28 that other wavelengths are chosen. The important
29 criterion in selecting the wavelengths is that the
30 relative absorptions of the tissue to be detected and
31 its surrounding tissue must be related such that the
32 relationship at the first frequency is different from
33 the relationship at the second frequency. In the
34 present example the absorption of the caries tissue at
35 the first frequency (1.3 μm) is 50% greater than that of
36 the dentine tissue, while at the second frequency

1 (1.44 μ m) it is 30% less.

2

3 The principle of differential detection of absorption
4 is well known and documented as it applies to the
5 detection of minor species in the presence of
6 interfering absorbers. It is most often encountered in
7 the laboratory and in commercial infra-red gas
8 analyzers. Its application to two-dimensional imaging
9 has not previously been investigated.

10

11 Figure 2 illustrates a laboratory arrangement used to
12 demonstrate that a practical step in realising this
13 differential absorption detection in a simple fashion
14 is to modulate the two light sources at the same
15 frequency but in antiphase to each other. Light
16 Emitting Diodes (LEDs) 20 and 22, operating with
17 relatively narrow bandwidths of several nm at
18 wavelengths within the regions previously noted, are
19 the preferred sources. Light transmitted through a
20 sample 24 is viewed with a detector 26 provided with an
21 electronic filter 28. This electronic filter 28 is set
22 to pass only the modulated frequency and thus the
23 detector 26 "sees" only the differential signal, that
24 is the signal corresponding to differences in
25 absorption between the two wavelengths. This approach
26 provides the basis for caries detection.

27

28 The use of any optical wavelength, as opposed to an X-
29 ray wavelength, has the added complication of the high
30 level of light scattering in tissue. This scattering
31 complicates the process of trying to retain detailed
32 image information on objects buried within such
33 scattering media. The longer wavelengths used in this
34 method and device help to reduce the level of scatter
35 but not to the point where even teeth become free of
36 the problem.

1 Thus an additional feature of this invention is the
2 inclusion of a means by which the effects of scatter
3 can be significantly reduced. Specifically, the rays
4 allowed to fall on the detector 28 are restricted to
5 only those having direct or nearly direct paths through
6 the sample 24 from the sources 20,22. This is
7 accomplished through the use of appropriate optical
8 spatial filtering 32 and a re-imaging lens 34. These
9 rays, sometimes referred to as 'ballistic' rays, are
10 those which pass through the sample 24 substantially
11 unscattered and are therefore better able to preserve
12 the information or the hidden detail associated, in
13 this case, with differential absorption between caries
14 and the surrounding sound tissue.

15

16 The laboratory arrangement of Figure 2b is used to
17 observe details of carious lesions. The detector used
18 is a CCD camera 30. This arrangement provides two
19 dimensional imaging of the ballistic rays. The phrase
20 "ballistic shadowgram" has been used to refer to this
21 2D image.

22

23 In summary, the use of a pair of wavelengths within a
24 newly identified region, offering differing absorption
25 characteristics between caries and sound tissue,
26 provides a novel means for the detection of dental
27 disease.

28

29 Figure 3 illustrates a practical embodiment of a device
30 incorporating this principle wherein two modulated
31 light sources 2,3 at $1.25\mu\text{m}$ and $1.43\mu\text{m}$ provide
32 simultaneous illumination of a tooth 6. The output
33 from these sources 2,3 is substantially collimated by
34 means of lenses 10, 11 and a transmitting mirror 12 and
35 combined along a single axis.

36

1 The combined beams are transmitted along a first arm 4
2 of the device and reflected by a reflector 13 to an
3 aperture 5 placed on one side of a tooth or teeth 6. A
4 second arm 14 of the device has an aperture 15 which is
5 disposed on the opposite side of the same tooth or
6 teeth 6, and this second arm 14 contains lenses 7,8 and
7 a reflector 16 such that a spatially filtered image of
8 the tooth 6 is presented to a two-dimensional array
9 detector 9. The image produced by the detector 9 is
10 transmitted by cable 18 to a computer and/or visual
11 display unit for display of the image.

12

13 The main features of this arrangement could be achieved
14 equally well using fibre optic delivery for the source
15 light, and image preserving fibre optic coupling for
16 the detection path.

17

18 Equally the resulting image could be scanned in one
19 dimension across a line array of detectors, or scanned
20 in both directions across a single point detector.
21 Such arrangements may be required since two dimensional
22 array detectors at these wavelengths are slow and
23 expensive.

24

25 Whatever approach is used to achieve a two dimensional
26 image of the tooth or teeth, the resulting image is
27 displayed, in real time, on the screen of a video
28 monitor (not shown). The image may be subject to
29 processing by software running on an associated
30 computer or microprocessor (not shown) to
31 electronically enhance the raw image, and means such as
32 a printer may be provided for producing hard copies of
33 images for maintaining patient records.

34

35 Since other variations in absorption occur in this
36 spectral region, including differences between enamel

1 and dentine, the technique may be extended to include
2 visualisation of normal dental structures such as root
3 canals and pulp chambers.

4

5 The embodiment described above offers significant
6 advantages over currently available devices for the
7 early detection of dental disease. Specifically it
8 employs non-ionising radiation and its use is therefore
9 intrinsically safer than the use of X-rays.

10

11 The device as described offers real time images through
12 teeth combined with the availability of hard copies of
13 those images. Finally the device is constructed from
14 relatively inexpensive materials and processes
15 resulting in a diagnostic offered at a much lower price
16 than current X-ray machines.

17

18 Whilst the foregoing describes application of the
19 invention to the examination of dental tissue, it is
20 apparent that the invention is equally applicable to
21 examination of other animal tissue structures.

22

23 Modifications and improvements may be made to the above
24 without departing from the scope of the invention.

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1 CLAIMS

2

3 1. A method of two-dimensional imaging of tissue by
4 differential absorption detection comprising the
5 steps of:

6

7 transmitting radiation of two wavelengths through
8 tissue;

9

10 detecting transmitted radiation;

11

12 measuring the absorption of said transmitted
13 radiation; and

14

15 comparing absorption of said two wavelengths.

16

17 2. A method as claimed in Claim 1 wherein the tissue
18 is dental tissue.

19

20 3. A method as claimed in Claim 1 or Claim 2 wherein
21 the tissue to be detected includes first and
22 second tissue types having different relative
23 transmission characteristics and wherein the first
24 and second wavelengths are selected such that the
25 relative transmission characteristics of the first
26 and second tissue types at the first wavelength
27 are significantly different from the relative
28 transmission characteristics of the first and
29 second tissue types at the second wavelength.

30

31 4. A method as claimed in Claim 3 wherein the
32 relative transmission of the first tissue type is
33 greater than the relative transmission of the
34 second tissue type at the first wavelength and the
35 relative transmission of the second tissue type is
36 greater than the relative transmission of the

- 1 first tissue type at the second wavelength.
2
- 3 5. A method as claimed in any preceding claim wherein
4 the radiation is non-ionising and has wavelengths
5 of size sufficient to preclude scattering in
6 tissue.
7
- 8 6. A method as claimed in any preceding claim wherein
9 the radiation is infra-red light.
10
- 11 7. A method as claimed in Claim 6 wherein the light
12 comprises two wavelengths within the range 1.1 to
13 1.6 micrometres.
14
- 15 8. A method as claimed in Claim 7 wherein a first
16 wavelength is within the range 1.4 to 1.5
17 micrometres and shows a dominant change in
18 transmission characteristics.
19
- 20 9. A method as claimed in Claim 7 or Claim 8 wherein
21 a second wavelength is within the range 1.2 to 1.3
22 micrometres and acts as a reference.
23
- 24 10. A method as claimed in any preceding claim wherein
25 the two wavelengths are modulated at the same
26 frequency.
27
- 28 11. A method as claimed in any preceding claim wherein
29 the two wavelengths are in antiphase.
30
- 31 12. A method as claimed in any preceding claim wherein
32 detection of the transmitted radiation is by
33 optical means.
34
- 35 13. A method as claimed in any of Claims 10 to 12
36 including the step of filtering out all

1 transmitted radiation not at the modulated
2 frequency.

3

4 14. A method as claimed in any preceding claim
5 including the step of restricting detection to
6 radiation transmitted in a substantially direct
7 path through the tissue.

8

9 15. A method as claimed in Claim 14 wherein detection
10 is restricted by optical spatial filtering.

11

12 16. A method as claimed in any preceding claim applied
13 to the detection of dental caries.

14

15 17. A method as claimed in any preceding claim applied
16 to distinguish between different dental structures
17 such as enamel, dentine, root canals and pulp
18 chambers.

19

20 18. Apparatus for differential absorption detection
21 comprising:

22

23 two modulated light sources, each source producing
24 light of a different wavelength;

25

26 optic transmission means to train the light from
27 said light sources to illuminate a first side of a
28 tissue sample *in vivo*; and

29

30 optic detection means to detect light transmitted
31 through said tissue sample to a second side of
32 said tissue sample.

33

34 19. Apparatus as claimed in Claim 19 wherein said
35 optic transmission means is adapted to combine the
36 light from said sources to provide a single beam

- 1 of incident light on said tissue.
2
- 3 20. Apparatus as claimed in Claim 18 or Claim 19
4 wherein the optic detection means includes an
5 electronic filter.
6
- 7 21. Apparatus as claimed in any of Claims 18 to 20
8 comprising means to spatially filter transmitted
9 light.
10
- 11 22. Apparatus as claimed in any of Claims 18 to 20
12 comprising means to display detected transmitted
13 light as an image.
14
- 15 23. Apparatus for detection of dental caries and/or
16 visualisation of dental structures by differential
17 absorption detection comprising:
18
19 two modulated light sources;
20
21 optic means to combine light from said sources to
22 train a single beam on a first side of a tooth;
23
24 receiving means on a second side of the tooth for
25 receiving light transmitted through the tooth;
26
27 an optic filter to spatially filter the
28 transmitted light; and
29
30 optic detection means to detect the spatially
31 filtered light.
32
- 33 24. Apparatus as claimed in Claim 23 comprising means
34 to display detected transmitted light as an image
35 of the tooth.
36

1 25. Apparatus as claimed in Claim 24 comprising means
2 to display the image in real time.

3

4 26. Apparatus as claimed in Claim 24 or Claim 25
5 comprising means to electronically enhance the
6 image.

7

8 27. Apparatus as claimed in any of Claims 24 to 26
9 comprising means to produce hard copies of said
10 image.

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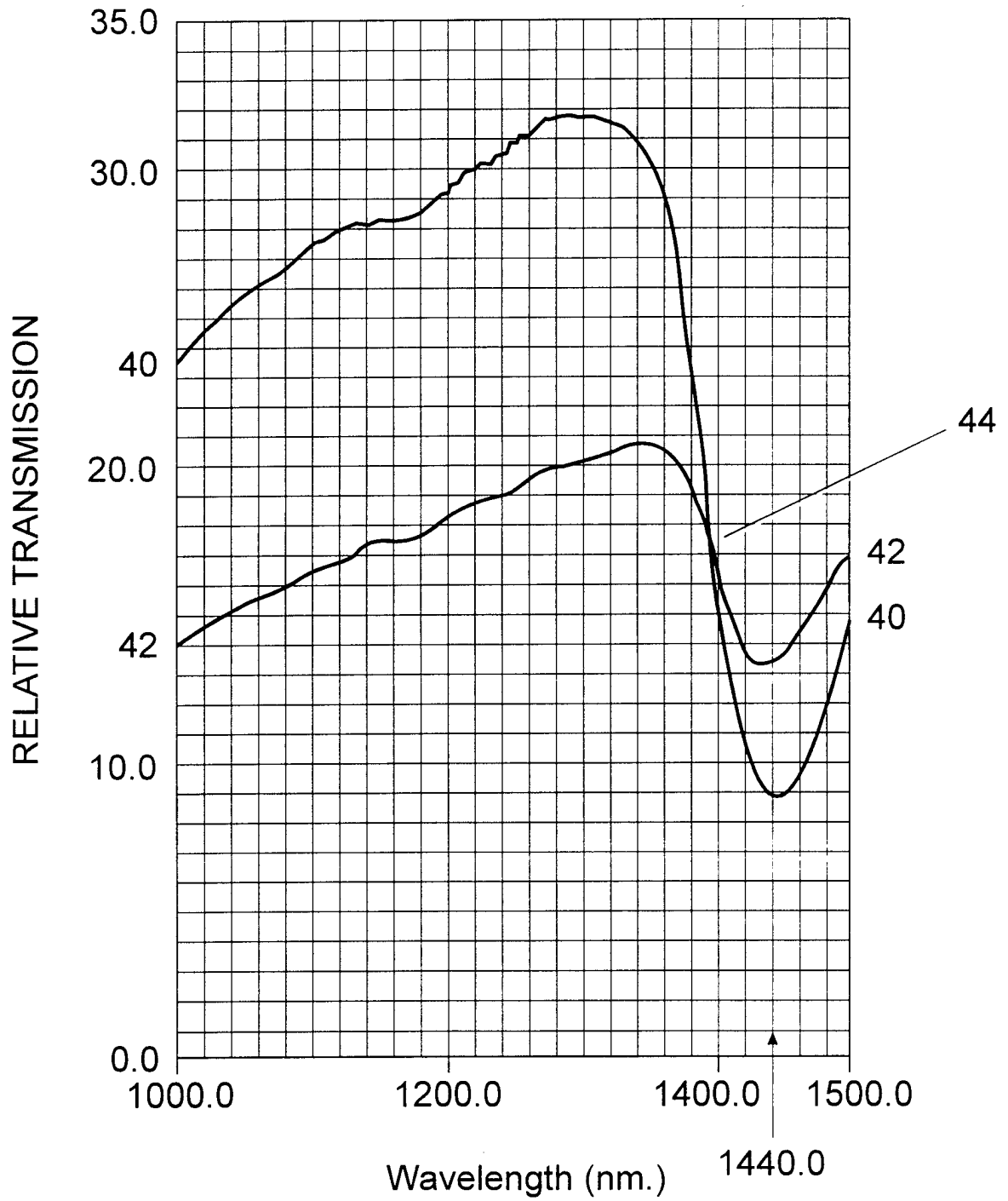


Fig. 1

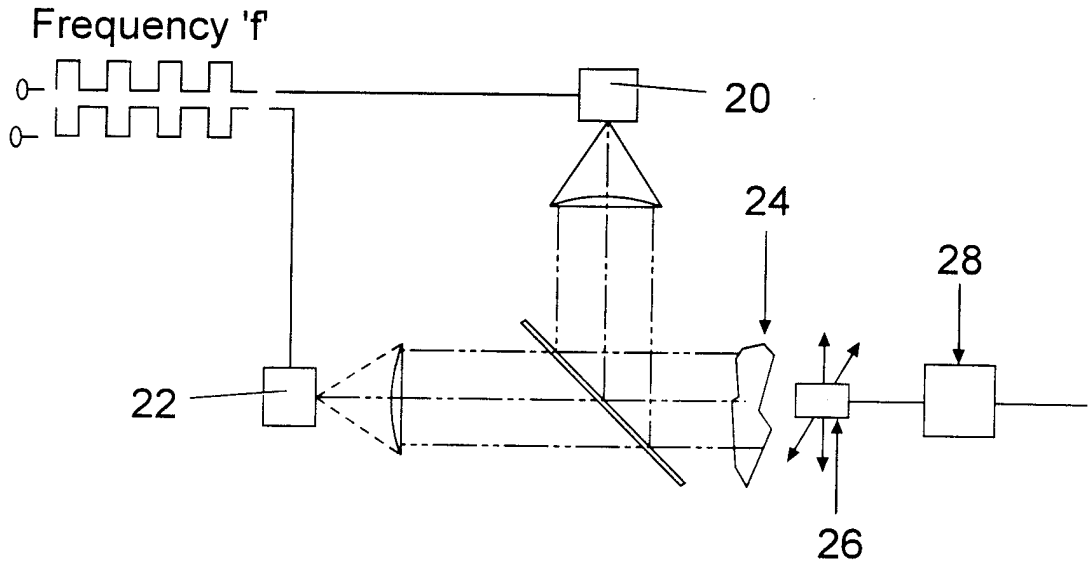


Fig. 2a

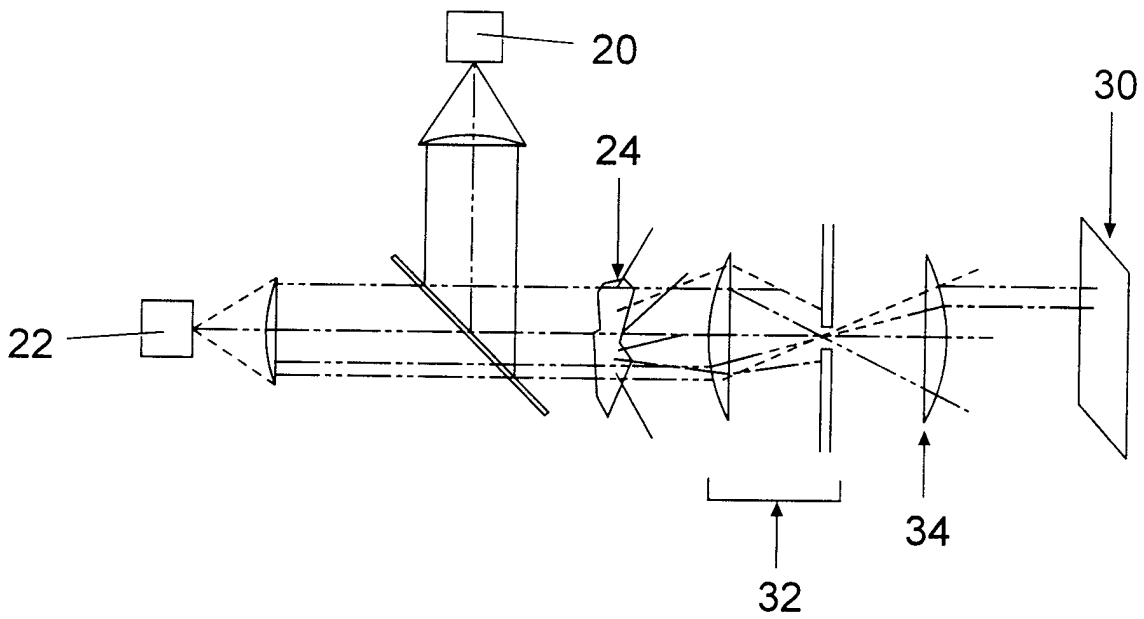


Fig. 2b

3 / 3

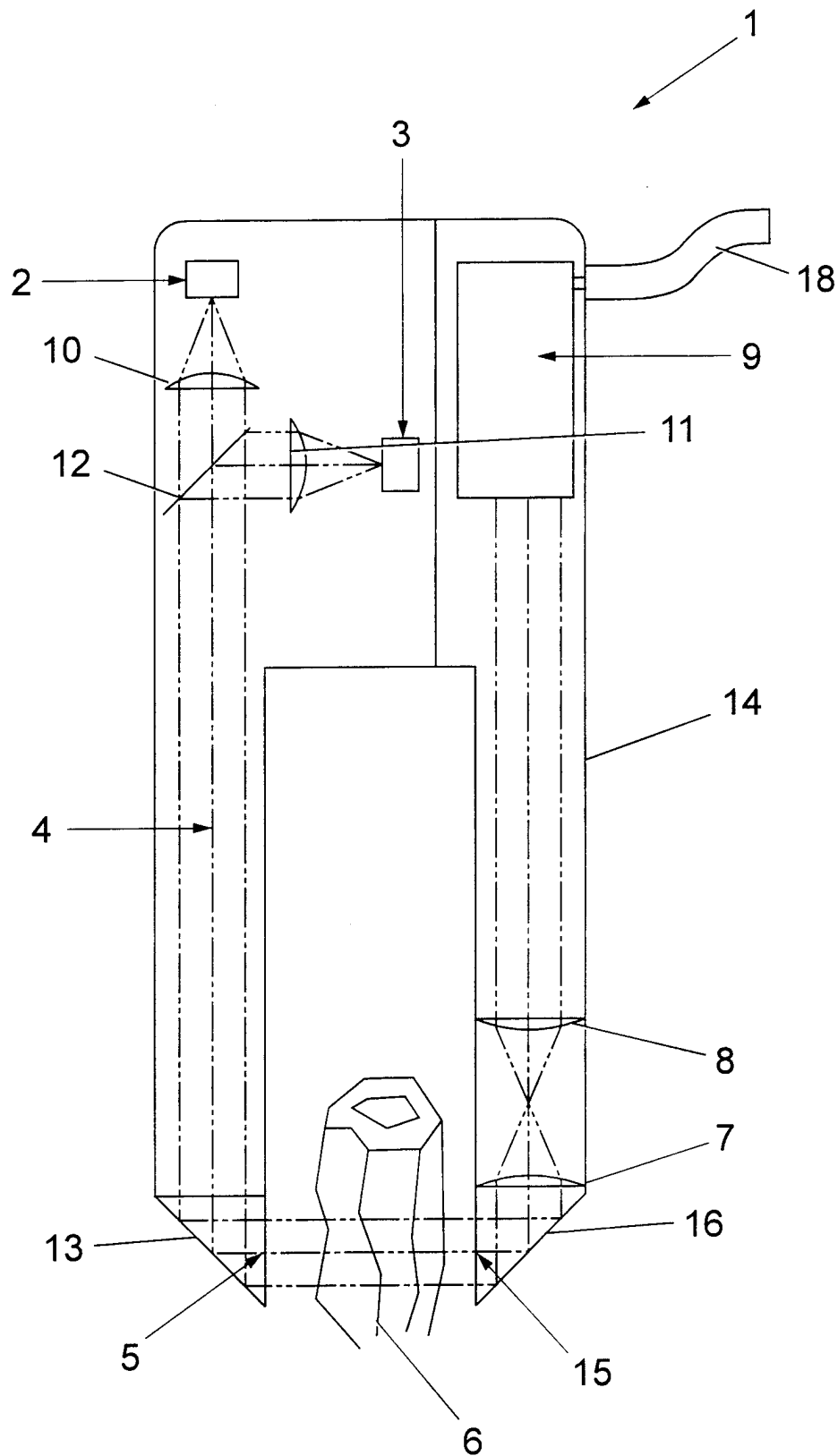


Fig. 3

INTERNATIONAL SEARCH REPORT

International Application No

PCT/GB 98/01524

A. CLASSIFICATION OF SUBJECT MATTER

IPC 6 A61B5/00

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

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 6 A61B

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

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

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 5 259 761 A (J.M. SCHNETTLER ET AL.) 9 November 1993	1-6,12
X	see column 7, line 50 - column 9, line 17	17-19
A	see column 13, line 11 - line 20 ---	23
X	US 5 040 539 A (J.M. SCHMITT ET AL.) 20 August 1991	1-6
X	see column 4, line 4 - line 18	10-14
X	see column 8, line 58 - column 9, line 32 see column 9, line 47 - column 10, line 43 ---	17-19,23
A	DE 43 07 411 A (MIRA G.M.B.H.) 15 September 1994	1,2,5-7
A	see column 2, line 22 - line 63	9,12,14
A	see column 3, line 6 - line 36	16-19
A	see column 5, line 54 - column 7, line 57 ---	22-25
	-/--	

Further documents are listed in the continuation of box C.

Patent family members are listed in annex.

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Date of the actual completion of the international search

4 September 1998

Date of mailing of the international search report

14/09/1998

Name and mailing address of the ISA

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INTERNATIONAL SEARCH REPORT

International Application No
PCT/GB 98/01524

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	US 5 570 182 A (H. NATHEL ET AL.) 29 October 1996	1,2,5-7
A	see column 2, line 54 - column 3, line 14	12,14-19
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	see column 8, line 34 - column 9, line 20 ----	
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