(11) EP 1 757 357 B1

(12)

EUROPEAN PATENT SPECIFICATION

(45) Date of publication and mention of the grant of the patent: 24.04.2013 Bulletin 2013/17

(21) Application number: 05720779.7

(22) Date of filing: 15.03.2005

(51) Int Cl.:

B01F 3/08 (2006.01) B01F 13/00 (2006.01) B01J 13/04 (2006.01) B01F 5/04 (2006.01) B01F 15/00 (2006.01)

(86) International application number: **PCT/JP2005/004522**

(87) International publication number:WO 2005/089921 (29.09.2005 Gazette 2005/39)

(54) METHOD AND DEVICE FOR PRODUCING MICRO-DROPLETS

VERFAHREN UND VORRICHTUNG ZUR HERSTELLUNG VON MIKROTRÖPFCHEN PROCEDE ET DISPOSITIF DE FABRICATION DE MICROGOUTTELETTES

(84) Designated Contracting States: CH DE FR GB LI NL SE

(30) Priority: 23.03.2004 JP 2004083802

(43) Date of publication of application: 28.02.2007 Bulletin 2007/09

(73) Proprietor: Japan Science and Technology Agency Kawaguchi-shi, Saitama 332-0012 (JP)

(72) Inventors:

 TORII, Toru 1670051 (JP)

HIGUCHI, Toshiro;
Yokohama-shi, Kanagawa 2240006 (JP)

 NISHISAKO, Takashi 1100008 (JP)

 OKUSHIMA, Shingo 1580098 (JP)

(74) Representative: Hoarton, Lloyd Douglas Charles

Forresters Skygarden

Erika-Mann-Strasse 11 80636 München (DE)

(56) References cited:

EP-A1- 1 362 634 EP-A2- 1 358 931 WO-A1-02/068104 JP-A- 2001 517 545 JP-A- 2004 122 107 US-A- 6 146 103 US-A1- 2003 106 596

US-B1- 6 342 142

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).

15

25

30

Technical Field

[0001] The present invention relates to a method and an apparatus for producing microdroplets, and more particularly, relates to a method and an apparatus for producing double emulsion-microcapsules.

1

Background Art

[0002] The inventors of the present invention have already filed a patent application relating to a method for producing emulsions and microcapsules and an apparatus therefor as the following Patent Document 1.

Patent Document 1: WO 02/068104 A1.

[0003] EP 1 362 634, which is a family member of Patent Document 1, discloses a method according to the preamble of claim 1, and an apparatus according to the preamble of claim 4, for rapidly producing an emulsion and microcapsules.

Disclosure of Invention

[0004] The above prior art on production of microdroplets was further improved, and an object of the present invention is to provide a method and an apparatus for producing microdroplets with various embodiments.

[0005] The present invention provides a method and an apparatus for producing microdroplets as claimed.

Brief Description of the Drawings

[0006]

Fig. 1 includes schematic views each showing the state in which microdroplets are produced at a long period using a cross microchannel when a flow rate ratio between a dispersion phase and a continuous phase is small, according to a first embodiment of the present invention.

Fig. 2 is a view showing the state in which the droplets are alternately produced, as shown in Fig. 1.

Fig. 3 is a photograph taken by a high-speed camera showing the state in which microdroplets having uniform sizes and different components are produced. Fig. 4 is a view showing the state in which microdroplets are sequentially produced, according to a second embodiment which is not part of the present invention.

Fig. 5 includes schematic views of an apparatus showing the state in which microdroplets are produced at a short period using a cross microchannel.when a flow rate ratio between a dispersion phase and a continuous phase is large, according to a third embodiment of the present invention.

Fig. 6 is a schematic view showing the state of a fourth embodiment according to the present invention in which double emulsion-microcapsules are produced by using microdroplets which have uniform sizes and have different components and which are alternately produced at a regular period in the third embodiment.

Fig. 7 is a view showing the state of production of a W/O/W type emulsion encapsulating two types of microdroplets, according to the present invention. Fig. 8 is a schematic view showing a modified em-

bodiment according to the first embodiment of the present invention.

Fig. 9 is a schematic view showing the state of separation of satellite droplets, according to a fifth embodiment which is not part of the present invention. Fig. 10 is a view showing a modified embodiment according to the fifth embodiment shown in Fig. 9. Fig. 11 is a view showing the state of separation of satellite droplets.

Fig. 12 is a view showing the shape of an acrylic resin-made microchannel shown in Fig. 11.

Fig. 13 is a schematic view showing the state of separation of satellite droplets, according to a sixth embodiment which is not part of the present invention. Fig. 14 is a schematic view showing the state of production of a double emulsion using satellite droplets, according to a seventh embodiment which is not part of the present invention.

Best Mode for Carrying Out the Invention

[0007] In a method for producing microdroplets, at a cross intersection portion at which a first continuous phase and a first and a second dispersion phase intersect with each other, different microdroplets are alternately produced by allowing the first dispersion phase and the second dispersion phase to alternately act on the first continuous phase. In addition, various types of microdroplets, in particular, double emulsion-microcapsules can be easily produced in a simple manner.

[0008] In addition, primary droplets and satellite droplets are easily separated and then can be recovered separately. In addition, by using the satellite droplets, high-quality and highly precise double emulsion-microcapsules can be manufactured.

Embodiments

[0009] Hereinafter, embodiments of the present invention will be described in detail.

[0010] Fig. 1 includes schematic views each showing the state in which microdroplets are produced at a long period using a cross microchannel when a flow rate ratio between a dispersion phase and a continuous phase is small, according to a first embodiment of the present in-

[0011] An embodiment will be described in which

30

45

microdroplets having uniform sizes and different components are alternately produced at a regular period.

[0012] In Fig. 1, reference numeral 1 indicates a first microchannel, reference numeral 2 indicates a continuous phase supplied from the first microchannel 1, reference numeral 3 indicates a second microchannel, reference numeral 4 indicates a first dispersion phase supplied from the second microchannel 3, reference numeral 5 indicates a third microchannel, reference numeral 6 indicates a second dispersion phase supplied from the third microchannel 5, reference numeral 7 indicates an intersection portion having a cross structure, reference numeral 8 indicates a fourth microchannel, reference numeral 9 indicates a first microdroplet sent through the fourth microchannel 8, reference numeral 10 indicates a second microdroplet alternately produced with the first microdroplet 9 at a regular period, reference numeral 11 indicates a control device controlling supply of the first dispersion phase 4 and the second dispersion phase 6 in the microchannels, reference numeral 12 indicates a syringe pump (flow rate changeable liquid feed device) which is connected to the control device 11 and which is used as a first liquid feed device supplying the first dispersion phase 4, and reference numeral 13 indicates a syringe pump (flow rate changeable liquid feed device) which is connected to the control device 11 and which is used as a second liquid feed device supplying the second dispersion phase 6.

[0013] In this embodiment, using the cross intersection portion 7 of the microchannels, the microdroplets 9 and 10 having uniform sizes are produced at predetermined intervals, so that a line of the microdroplets is formed. That is, when the first syringe pump (flow rate changeable liquid feed device) 12 for the first dispersion phase 4 and the second syringe pump (flow rate changeable liquid feed device) 13 for the second dispersion phase 6 are alternately operated so as to feed the first dispersion phase 4 and the second dispersion phase 6 at the same flow rate, at an appropriate flow rate of the continuous phase 2, the microdroplets 9 and 10 having uniform sizes and different components are alternately produced at the cross intersection portion 7 at a regular period.

[0014] Fig. 2 is a view showing the state in which the microdroplets are alternately produced, and Fig. 3 is a photograph taken by a high-speed camera showing the state of production of microdroplets having uniform sizes and different components.

[0015] In this embodiment, a glass-made microchannel was used which had a channel width of $80~\mu m$ and a depth of $40~\mu m$ and which was processed by a hydrophobic treatment. As the dispersion phases shown in Fig. 2, a red ink a and a blue ink b each diluted with water were used as the first and the second dispersion phases, respectively, and each dispersion phase was supplied at a flow rate of 0.01 ml/h. A corn oil (viscosity: $58.5~mPa\cdot s$, surface tension: 33.2~mN/m (both being measured at $20^{\circ}C$)) used as the continuous phase was supplied at a flow rate of 0.10 ml/h.

[0016] Fig. 4 is a view showing the state in which microdroplets are sequentially produced, according to a second embodiment which is not part of the present invention.

[0017] In this embodiment, at a cross intersection portion at which a first continuous phase, a first dispersion phase, and a second dispersion phase intersect with each other, the first dispersion phase and the second dispersion phase are allowed to act on the first continuous phase so as to produce different microdroplets, and in this case, by the control using the control device 11, when one droplet 9 is formed from the first dispersion phase, two droplets 10 are then continuously formed from the second dispersion phase.

[0018] Fig. 5 includes schematic views of an apparatus showing the state in which microdroplets are produced at a short period using a cross microchannel when a flow rate ratio between a dispersion phase and a continuous phase is large, according to a third embodiment of the present invention.

[0019] In this figure, reference numeral 21 indicates a first microchannel, reference numeral 22 indicates a continuous phase supplied from the first microchannel 21, reference numeral 23 indicates a second microchannel, reference numeral 24 indicates a first dispersion phase supplied from the second microchannel 23, reference numeral 25 indicates a third microchannel, reference numeral 26 indicates a second dispersion phase supplied from the third microchannel 25, reference numeral 27 indicates an intersection portion having a cross structure, reference numeral 28 indicates a fourth microchannel, reference numeral 29 indicates a first microdroplet sent through the fourth microchannel 28, reference numeral 30 indicates a second microdroplet alternately produced with the first microdroplet 29 at a regular period, reference numeral 31 indicates a control device controlling supply of the first dispersion phase 24 and the second dispersion phase 26 in the microchannels, reference numeral 32 indicates a first syringe pump (flow rate changeable liquid feed device) which is connected to the control device 31 and which is used as a first liquid feed device supplying the first dispersion phase 24, and reference numeral 33 indicates a second syringe pump (flow rate changeable liquid feed device) which is connected to the control device 31 and which is used as a second liquid feed device supplying the second dispersion phase 26.

[0020] In this third embodiment, the microdroplets 29 and 30 are produced at a shorter period compared to the case in which the microdroplets 9 and 10 are alternately produced in the first embodiment shown in Fig. 1.

[0021] Fig. 6 is a schematic view showing the state of a fourth embodiment according to the present invention in which double emulsion-microcapsules are produced by using microdroplets which have uniform sizes and have different components and which are alternately produced at a regular period in the third embodiment.

[0022] In this embodiment, reference numeral 40 indicates an outlet for ejecting two types of microdroplets,

that is, the microdroplets 29 and 30 which are alternately produced at a regular period and which have uniform sizes and different components, reference numeral 41 indicates an intersection portion having a cross structure, reference numeral 42 indicates a fifth microchannel, reference numeral 43 indicates a continuous phase supplied from the fifth microchannel 42, reference numeral 44 indicates a sixth microchannel, reference numeral 45 indicates a continuous phase supplied from the sixth microchannel 44, reference numeral 46 indicates a produced microcapsule (double emulsion), reference numeral 47 indicates a microcapsule (double emulsion) recovery channel recovering the microcapsules (double emulsion) 46, and reference numeral 48 indicates a continuous phase sending the microcapsule (double emulsion) 46.

[0023] As described above, the microdroplets 29 and 30 alternately produced at a regular period and having different components are further encapsulated, so that the microcapsules (double emulsion) 46 containing the same numbers of two types of microdroplets are obtained.

[0024] Fig. 7 is a view showing the state of production of a W/O/W type emulsion encapsulating two types of microdroplets, according to the present invention.

[0025] Next, as for a method for producing microdroplets using microchannels, a method for obtaining a monodispersion emulsion will be described in which satellite droplets are separated and removed from microdroplets produced in the microchannels.

[0026] Fig. 8 is a schematic view showing a modified embodiment according to the first embodiment of the present invention.

[0027] In this embodiment, the structure is formed in which different droplets are ejected from a T-shaped intersection portion 27-1 and a T-shaped intersection portion 27-2 which is located at a position shifted therefrom, and the first microdroplets 29 sent through the fourth microchannel 28 and the second microdroplets 30 alternately produced with the first microdroplets 29 at a regular period are sequentially produced. The rest of the structure is equivalent to that in the first embodiment.

[0028] Fig. 9 is a schematic view showing the state of separation of satellite droplets, according to a fifth embodiment which is not part of the present invention.

[0029] In this figure, reference numeral 51 indicates a first microchannel (continuous phase supply channel), reference numeral 52 indicates a continuous phase supplied from the first microchannel (continuous phase supply channel) 51, reference numeral 53 indicates an intersection portion having a T-shaped structure, reference numeral 54 indicates a second microchannel (dispersion phase supply channel), reference numeral 55 indicates a dispersion phase supplied from the second microchannel (dispersion phase supply channel) 54, reference numeral 56 indicates a third microchannel, reference numeral 57 indicates a primary droplet produced at the intersection portion 53 having a T-shaped structure and

sent through the third microchannel 56, reference numeral 58 indicates a satellite droplet produced together with the primary droplet 57, reference numeral 59 indicates an outlet of the third microchannel 56, reference numeral 60 indicates a junction portion of a microchannel connected to the outlet 59, reference numeral 61 indicates an expansion portion (tapered portion) of the microchannel, reference numeral 62 indicates a branching portion, reference numeral 63 indicates a primary droplet recovery channel recovering the primary droplets 57, reference numeral 64 indicates a liquid carrying the primary droplets, and reference numeral 65 indicates a satellite droplet recovery channel recovering the satellite droplets 58. [0030] As shown in Fig. 9, when the primary droplets 57 are produced at the intersection portion 53 having a T-shaped structure, the fine satellite droplets 58 are simultaneously produced. In the case in which microcapsules (double emulsion) are produced using the primary droplets 57, it is often unfavorable if the satellite droplets 58 are encapsulated in the microcapsules (double emulsion) together with the primary droplets 57.

[0031] Accordingly, in order to avoid the case described above, at the expansion portion 61, the primary droplets 57 are designed to be sent to the right side in the same manner as that before, and the satellite droplets 58 are designed to be sent downward, that is, the primary droplets 57 are sent toward the primary droplet recovery channel 63, and the satellite droplets 58 are sent downward to the satellite droplet recovery channel 65 recovering the satellite droplets 58.

[0032] According to this embodiment, the expansion portion (tapered portion) 61 and the branching microchannels 63 and 65 are provided downstream of the microdroplet production portion (intersection portion having a T-shaped structure) 53, and the satellite droplets 58 can be continuously separated from the primary droplets 57 thus formed.

[0033] Instead of the expansion portion 61 having a taper described above, an expansion portion 66 having a curved surface may be used, as shown in Fig. 10.

[0034] Fig. 11 is a view showing the state of separation of satellite droplets.

[0035] As shown in this figure, primary droplets (diameter: 70 μ m) 71 are separated from satellite droplets 72 (diameters: 1, 3, and 5 μ m).

[0036] Fig. 12 is a view showing the shape of an acrylic resin-made microchannel shown in Fig. 11, this microchannel is formed of a first microchannel (continuous phase supply channel) 73 of 200 μm wide and 100 μm deep, a second microchannel (dispersion phase supply channel) 74 of 120 μm wide and 100 μm deep, a primary droplet recovery channel 75 of 800 μm wide and 100 μm deep, and a satellite droplet recovery channel 76 of 200 μm wide and 100 μm deep, and in addition, the branching angle θ of the satellite droplet recovery channel 76 to the primary droplet recovery channel 75 is 30°.

[0037] In this embodiment, as the dispersion phase and the continuous phase, pure water and a corn oil (vis-

cosity: 58.5 mPa·s, surface tension: 33.2 mN/m, both being measured at 20°C) were used, respectively, and the flow rate control was performed for each phase by a liquid feed device (syringe pump).

[0038] As for the flow rate conditions in Fig. 11, the dispersion phase flow rate and the continuous phase flow rate were set to 1.0 ml/h and 15.0 ml/h, respectively, and it was confirmed that the primary droplets 71 (diameter: approximately 70 μ m) were separated from the satellite droplets 72 having three different sizes (diameters: 1, 3, and 5 μ m). It was observed that the satellite droplets having different sizes form respective lines and flow into the branching channel. When the flow rate of the dispersion phase and that of the continuous phase are increased, the size of the satellite droplets and the production number thereof both tend to be increased.

[0039] Fig. 13 is a schematic view showing the state of separation of satellite droplets, according to a sixth embodiment which is not part of the present invention.

[0040] In this figure, reference numeral 81 indicates a first microchannel (continuous phase supply channel), reference numeral 82 indicates a continuous phase supplied from the first microchannel (continuous phase supply channel), reference numeral 83 indicates an intersection portion having a cross structure, reference numeral 84 indicates a second microchannel (first dispersion phase supply channel), reference numeral 85 indicates a first dispersion phase supplied from the second microchannel (first dispersion phase supply channel) 84, reference numeral 86 indicates a third microchannel (second dispersion phase supply channel), reference numeral 87 indicates a second dispersion phase supplied from the third microchannel (second dispersion phase supply channel) 86, reference numeral 88 indicates a fourth microchannel, reference numeral 89 indicates a first primary droplet produced at the intersection portion 83 having a cross structure, reference numeral 90 indicates a first satellite droplet produced simultaneously with the first primary droplet 89, reference numeral 91 indicates a second primary droplet produced at the intersection portion 83 having a cross structure, reference numeral 92 indicates a second satellite droplet produced simultaneously with the second primary droplet 91, reference numeral 93 indicates a junction portion of a microchannel connected to an outlet ejecting the droplets 89 to 92, reference numeral 94 indicates an expansion portion (tapered portion) of the microchannel, reference numeral 95 indicates a branching portion, reference numeral 96 indicates a primary droplet recovery channel recovering the primary droplets 89 and 91, reference numeral 97 indicates a liquid carrying the primary droplets 89 and 91, reference numeral 98 indicates a first satellite droplet recovery channel recovering the first satellite droplets 90, and reference numeral 99 indicates a second satellite droplet recovery channel recovering the second satellite droplets 92.

[0041] In this embodiment, the expansion portion (tapered portion) 94 and the first and the second satellite

droplet recovery channels 98 and 99 are provided downstream of the microdroplet production portion (intersection portion having a cross structure) 83, and the first and the second satellite droplets 90 and 92 can be independently separated from the primary droplets 89 and 91.

[0042] According to the structure described above, the droplet production and the classification operation can be simultaneously performed in the microchannels, and monodispersion droplets/fine particles can be advantageously obtained without classification operation performed outside the apparatus.

[0043] The satellite droplets thus separated and recovered are very fine, and hence these droplets can be used for production of a double emulsion.

[0044] Fig. 14 is a schematic view showing the state of production of a double emulsion using satellite droplets, according to a seventh embodiment which is not part of the present invention.

[0045] In this figure, reference numeral 101 indicates a first microchannel (satellite droplet supply channel), reference numeral 102 indicates a liquid carrying satellite droplets, reference numeral 103 indicates a satellite droplet, reference numeral 104 indicates an outlet for the satellite droplets, reference numeral 105 indicates a second microchannel (continuous phase supply channel), reference numeral 106 indicates a first continuous phase supplied from the second microchannel (continuous phase supply channel) 105, reference numeral 107 indicates a third microchannel (continuous phase supply channel), reference numeral 108 indicates a second continuous phase supplied from the third microchannel (continuous phase supply channel) 107, reference numeral 109 indicates a recovery channel for a double emulsion using satellite droplets, reference numeral 110 indicates a liquid carrying the double emulsion using satellite droplets, and reference numeral 111 indicates the double emulsion using satellite droplets.

[0046] According to this embodiment, as shown in Fig. 14, the double emulsion 111 encapsulating the satellite droplets 103 can be produced.

[0047] Incidentally, the present invention is not limited to the above examples, and various modifications may be carried out without departing from the scope of the present invention as defined in the claims and may not be excluded therefrom.

[0048] According to the present invention, using intersecting microchannels in combination, various types of microdroplets, in particular, double emulsion-microcapsules can be easily produced in a simple manner.

[0049] In addition, the primary droplets and the satellite droplets can be easily separated and recovered separately. Accordingly, high-quality and highly precise double emulsion-microcapsules can be manufactured.

Industrial Applicability

[0050] The method for producing microdroplets and the apparatus therefor, according to the present inven-

15

30

35

45

50

tion, can be used as a tool producing microcapsules in the fields of gene and pharmaceutical technologies.

Claims

- 1. A method for producing microdroplets, comprising the step of allowing a first dispersion phase (4) and a second dispersion phase (6) to act on a first continuous phase (2) at an intersection portion (7) among the first continuous phase (2), the first dispersion phase (4), and the second dispersion phase (6), wherein the intersection portion (7) is a cross intersection portion or the intersection portion (7) includes T-shaped intersection portions, the T-shaped intersection portions being located at positions shifted from each other, whereby the microdroplets are sequentially produced and wherein microdroplets formed of the first dispersion phase (4) are different from microdroplets formed of the second dispersion phase (6), characterised in that the first dispersion phase (4) and the second dispersion phase (6) are allowed to alternately act on at predetermined time intervals so that microdroplets having uniform sizes and different components are alternately produced at a regular period.
- The method of claim 1, wherein the period is changeable
- The method for producing microdroplets of any preceding claim, further comprising:

sequentially producing different microdroplets;

supplying a liquid containing the different microdroplets to another cross intersection portion to which the first continuous phase (22) and a second continuous phase (43, 45) are supplied, whereby double emulsion-microcapsules are produced.

An apparatus for producing microdroplets, comprising:

(a) an intersection portion (7) at which a first continuous phase (2), a first dispersion phase (4) and a second dispersion phase (6) intersect with each other, wherein the intersection portion (7) is a cross intersection portion, or the intersection portion (7) includes T-shaped intersection portions, the T-shaped intersection portions being located at positions shifted from each other; (b) a first liquid feed device controlling the first

- (b) a first liquid feed device controlling the first dispersion phase (4);
- (c) a second liquid feed device controlling the second dispersion phase (6); and
- (d) a control device connected to the first liquid

feed device and the second liquid feed device, wherein

- (e) the first liquid feed device and the second liquid feed device are controlled by a signal from the control device so that microdroplets formed of the first dispersion phase (4) and microdroplets formed of the second dispersion phase (6) are sequentially produced, **characterised in that**, by the signal from the control device, microdroplets having uniform sizes and different components are alternately produced at a regular period.
- **5.** The apparatus of claim 4, wherein by the signal from the control device, the period is changeable.

Patentansprüche

- Verfahren zur Herstellung von Mikrotröpfchen, umfassend den Schritt des Zulassens, dass eine erste Dispersionsphase (4) und eine zweite Dispersionsphase (6) auf eine erste kontinuierliche Phase (2) an einem Schnittbereich (7) unter der ersten kontinuierlichen Phase (2), der ersten Dispersionsphase (4) und der zweiten Dispersionsphase (6) einwirken, wobei der Schnittbereich (7) ein Querschnittsbereich ist oder der Schnittbereich (7) T-förmige Schnittbereiche einschließt, wobei sich die T-förmigen Schnittbereiche an Positionen befinden, die voneinander verschoben sind, wodurch die Mikrotröpfchen nacheinander hergestellt werden und wobei sich aus der ersten Dispersionsphase (4) ausgebildete Mikrotröpfchen von aus der zweiten Dispersionsphase (6) ausgebildeten Mikrotröpfchen unterscheiden, dadurch gekennzeichnet, dass es der ersten Dispersionsphase (4) und der zweiten Dispersionsphase (6) erlaubt ist, abwechselnd zu vorgegebenen Zeitintervallen einzuwirken, so dass Mikrotröpfchen von gleichförmiger Größe und unterschiedlichen Bestandteilen in einem regelmäßigen Zeitabschnitt abwechselnd hergestellt werden.
- 2. Verfahren nach Anspruch 1, wobei der Zeitabschnitt veränderbar ist.
- Verfahren zur Herstellung von Mikrotröpfchen nach irgendeinem vorhergehenden Anspruch, ferner umfassend:

aufeinanderfolgendes Herstellen verschiedener Mikrotröpfchen; und

Zuführen einer Flüssigkeit, die die verschiedenen Mikrotröpfchen enthält, zu einem weiteren Querschnittsbereich, dem die erste kontinuierliche Phase (22) und eine zweite kontinuierliche Phase (43, 45) zugeführt werden, wodurch Doppelemulsions-Mikrokapseln hergestellt werden.

35

40

45

- 4. Vorrichtung zur Herstellung von Mikrotröpfchen, umfassend:
 - (a) einen Schnittbereich (7), in dem sich eine erste kontinuierliche Phase (2), eine erste Dispersionsphase (4) und eine zweite Dispersionsphase (6) schneiden, wobei der Schnittbereich (7) ein Querschnittsbereich ist oder der Schnittbereich (7) T-förmige Schnittbereiche einschließt, wobei sich die T-förmigen Schnittbereiche an Positionen befinden, die voneinander verschoben sind;
 - (b) eine erste Flüssigkeitseinspeisungsvorrichtung, die die erste Dispersionsphase (4) steuert; (c) eine zweite Flüssigkeitseinspeisungsvorrichtung, die die zweite Dispersionsphase (6) steuert; und
 - (d) eine Steuervorrichtung, die mit der ersten Flüssigkeitseinspeisungsvorrichtung und der zweiten Flüssigkeitseinspeisungsvorrichtung verbunden ist, wobei
 - (e) die erste Flüssigkeitseinspeisungsvorrichtung und die zweite Flüssigkeitseinspeisungsvorrichtung durch ein Signal von der Steuervorrichtung gesteuert werden, so dass aus der ersten Dispersionsphase (4) ausgebildete Mikrotröpfchen und aus der zweiten Dispersionsphase (6) ausgebildete Mikrotröpfchen nacheinander hergestellt werden, dadurch gekennzeichnet, dass durch das Signal von der Steuervorrichtung Mikrotröpfchen von gleichförmiger Größe und unterschiedlichen Bestandteilen in einem regelmäßigen Zeitabschnitt abwechselnd hergestellt werden.
- Vorrichtung nach Anspruch 4, wobei der Zeitabschnitt durch das Signal von der Steuervorrichtung veränderbar ist.

Revendications

1. Un procédé de fabrication de microgouttelettes, consistant à permettre à une première phase de dispersion (4) et à une deuxième phase de dispersion (6) d'agir sur une première phase continue (2) à une intersection (7) entre la première phase continue (2), la première phase de dispersion (4) et la deuxième phase de dispersion (6), l'intersection (7) étant une intersection croisée ou l'intersection (7) comprenant des intersections en forme de T, les intersections en forme de T se trouvant à des endroits décalés les uns des autres, les microgouttelettes étant produites de manière séquentielle et les microgouttelettes de la première phase de dispersion (4) étant différentes de celles formées de la deuxième phase de dispersion (6), se caractérisant par le fait que la première phase de dispersion (4) et la deuxième phase de

- dispersion (6) sont autorisées à agir tour à tour à intervalles prédéterminés, de telle sorte que les microgouttelettes possèdent des tailles uniformes et que des composants différents soient produits tour à tour à une période régulière.
- Le procédé de la revendication 1, dans lequel la période est modifiable.
- 3. Le procédé de production de microgouttelettes de n'importe laquelle des revendications précédentes, supposant également de :

produire de manière séquentielle des microgouttelettes différentes ; et

fournir un liquide contenant les différentes microgouttelettes à une autre intersection croisée à laquelle la première phase continue (22) et une deuxième phase continue (43, 45) sont fournies, ce qui permet de produire des microcapsules d'émulsion doubles.

- **4.** Un appareil de production de microgouttelettes, se composant de ce qui suit :
 - (a) une intersection (7) à laquelle une première phase continue (2), une première phase de dispersion (4) et une deuxième phase de dispersion (6) se croisent, l'intersection (7) étant une intersection croisée ou l'intersection (7) comprenant des intersections en forme de T, les intersections en forme de T se trouvant à des endroits décalés les uns des autres ;
 - (b) un premier dispositif d'injection de liquide commandant la première phase de dispersion (4).
 - (c) un second dispositif d'injection de liquide commandant la deuxième phase de dispersion(6); et
 - (d) un dispositif de commande connecté au premier dispositif d'injection de liquide et au second dispositif d'injection de liquide, dans lequel
 - (e) les premier et second dispositifs d'injection de liquide sont commandés par un signal provenant du dispositif de commande pour produire séquentiellement des microgouttelettes formées de la première phase de dispersion (4) et des microgouttelettes formées de la deuxième phase de dispersion (6), se caractérisant par le fait que des microgouttelettes de taille uniforme et de composants différents sont tour à tour produites à une période régulière par le signal émanant du dispositif de commande.
- 55 5. L'appareil de la revendication 4, dans lequel la période peut être modifiée par le signal émanant du dispositif de commande.

FIG. 1

(a) .13 SECOND CONTROL DEVICE SYRINGE PUMP -6 10 -12 **FIRST** SYRINGE PUMP (b) .13 SECOND CONTROL DEVICE SYRINGE PUMP _3 -12 **FIRST SYRINGE PUMP**

FIG. 2

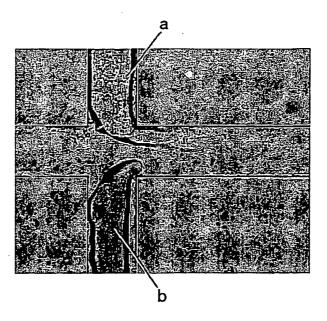


FIG. 3

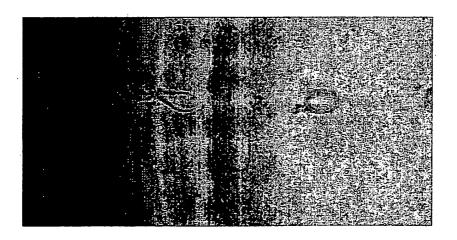


FIG. 4

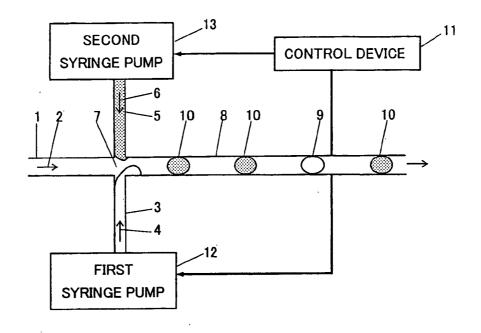
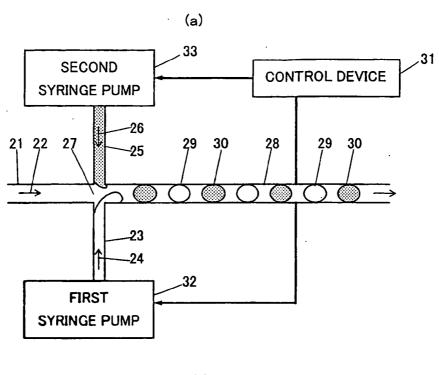


FIG. 5



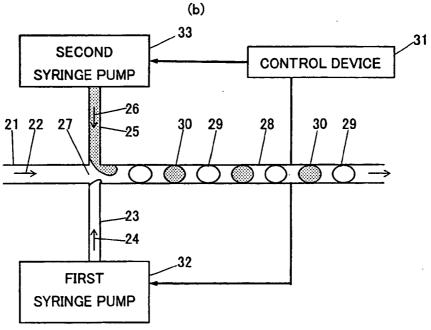


FIG. 6

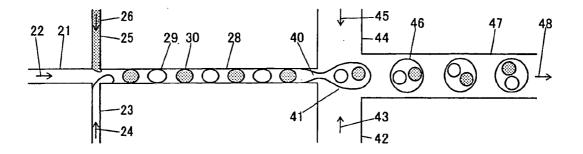


FIG. 7

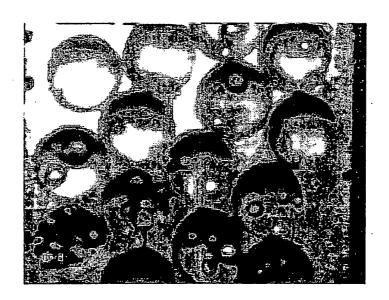


FIG. 8

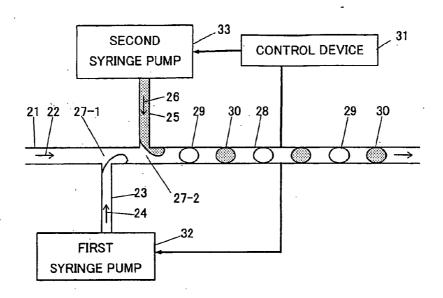


FIG. 9

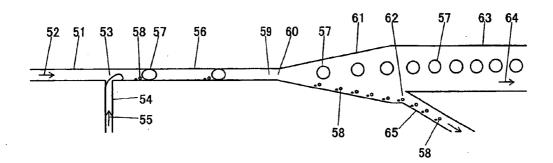


FIG. 10

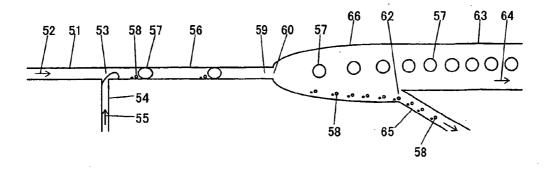


FIG. 11

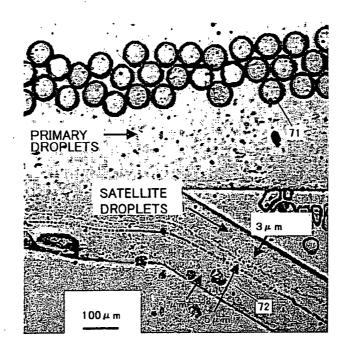


FIG. 12

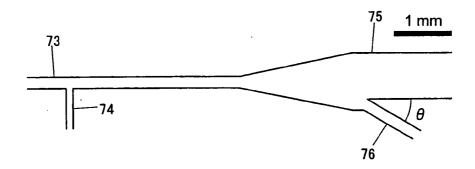


FIG. 13

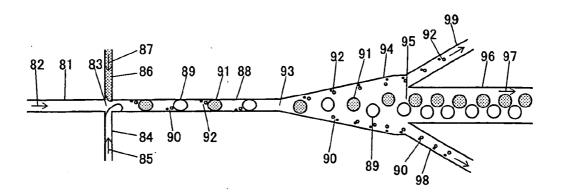
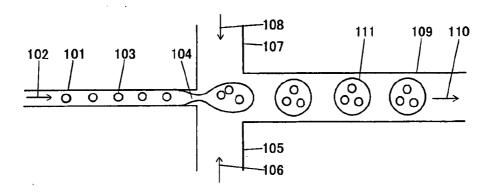


FIG. 14



EP 1 757 357 B1

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

• WO 02068104 A1 [0002]

• EP 1362634 A [0003]