

US 20050145785Al

## (19) **United States**  (12) **Patent Application Publication**  (10) **Pub. No.: US 2005/0145785 Al**

# ( 43) **Pub. Date: Jul. 7, 2005**

### (54) **OPTICAL PERISTALTIC PUMPING WITH OPTICAL TRAPS**

**Grier et al.** 

(75) Inventors: **David G. Grier,** Chicago, IL (US); **Sven Holger Behrens,** Mannheim (DE)

> Correspondence Address: **FOLEY** & **LARDNER 321 NORTH CLARK STREET SUITE 2800 CHICAGO, IL 60610-4764 (US)**

- (73) Assignee: **University of Chicago**
- (21) Appl. No.: **11/041,640**
- (22) Filed: Jan. 24, 2005

#### **Related U.S. Application Data**

(63) Continuation of application No. 10/651,370, filed on Aug. 29, 2003, now Pat. No. 6,847,032, which is a continuation of application No. 09/875,812, filed on Jun. 6, 2001, now Pat. No. 6,639,208.

#### **Publication Classification**

- (51) **Int. Cl.7** ............................... **H05H 3/02;** HOlS 3/00
- (52) **U.S. Cl.** .. **250/251**

#### (57) **ABSTRACT**

A method of use for holographic optical traps or gradients in which repetitive cycling of a small number of appropriately designed arrays of traps are used for general and very complex manipulations of particles and volumes of matter. Material transport results from a process resembling peristaltic pumping, with the sequence of holographically-defined trapping or holding manifolds resembling the states of a physical peristaltic pump.













![](_page_3_Figure_4.jpeg)

![](_page_3_Figure_5.jpeg)

![](_page_3_Figure_7.jpeg)

![](_page_3_Figure_8.jpeg)

![](_page_3_Figure_9.jpeg)

![](_page_3_Figure_10.jpeg)

![](_page_4_Figure_2.jpeg)

![](_page_5_Figure_2.jpeg)

![](_page_5_Figure_3.jpeg)

![](_page_5_Figure_4.jpeg)

FIG. 15

#### **OPTICAL PERISTALTIC PUMPING WITH OPTICAL TRAPS**

**[0001]** This invention was made with U.S. Government support under Contract No. DMR-9730189 awarded by the National Science Foundation, and through the MRSEC Program of the National Science Foundation under Award No. DMR-9880595. The U.S. Government also has certain rights to the invention pursuant to these contracts and awards. This application is a continuation of U.S. patent application Ser. No. 10/651,370, filed Aug. 29, 2003, which is a continuation of U.S. patent application Ser. No. 09/875, 812, filed Jun. 6, 2001, now U.S. Pat. No. 6,639,208.

#### FIELD OF THE INVENTION

**[0002]** The present invention is directed generally to a method and apparatus for controlling and manipulating small particles, a movable mass or a deformable structure. More particularly, the present invention is directed to a method and apparatus for using holographic optical traps to control and manipulate particles and volumes of matter in both general and complex ways.

#### BACKGROUND OF THE INVENTION

**[0003]** Optical traps use optical gradient forces to trap, most preferably, micrometer-scale volumes of matter in both two and three dimensions. A holographic form of optical trap can use a computer-generated diffractive optical element to create large numbers of optical traps from a single laser beam. These traps can be arranged in any desired configuration dependent on the need at hand.

**[0004]** Although systems are known to move particles precisely and with a relatively high degree of confidence, conventional systems require a separate hologram to be projected for each discrete step of a particle's motion. Computing multiple holograms can be very time consuming and requires substantial computational effort. Furthermore, computer-addressable projection systems required to implement such computer-generated optical traps or other dynamic optical trap systems, such as scanned optical tweezers, tend to be prohibitively expensive.

#### SUMMARY OF THE INVENTION

**[0005]** It is therefore an object of the invention to provide an improved method for manipulating particles and volumes of matter in both general and complex methods.

**[0006]** It is an additional object of the invention to provide an improved method for moving particles along a predetermined path with a high degree of accuracy and confidence.

**[0007]** It is still another object of the invention to provide a method for manipulating particles and volumes of matter which removes the computational burden of achieving complex rearrangements.

[0008] In accordance with the above objects, projecting a time varying sequence of such trap patterns makes possible dynamic reconfiguration of traps, with each new pattern updating the position of each trap by a distance small enough that particles trapped in the original pattern naturally fall into a corresponding trap in the next. The present invention therefore offers a method for accomplishing complex rearrangements of matter by cycling through a small number of precalculated holographic optical trap patterns. The cycling can be performed mechanically, removing both computational complexity and the expense of a fully general holographic optical trap system.

#### BRIEF DESCRIPTION OF THE DRAWINGS

**[0009] FIG. 1** depicts an individual particle being trapped in an optical trap within a manifold of optical traps, wherein the manifold's position is represented by a dashed line;

**[0010] FIG. 2** shows the transfer of an individual particle from a manifold of traps in a first pattern to a manifold of traps in a second pattern;

**[0011] FIGS. 3A-3D** shows the operative action of an optical peristalsis method;

**[0012] FIG. 4** shows the use of parallel linear manifolds of optical traps for transferring particles along a linear trajectory normal to the manifolds;

**[0013] FIG. SA** shows curved manifolds directing particles from the periphery of the pattern towards the centers of curvature; and **FIG. 5B** schematically shows how the pattern described in **FIG. SA** can sweep particles into a channel;

**[0014] FIG. 6A** shows nonuniformly curved manifolds used to divide a flow of particles into two separate flows; and **FIG. 6B** shows nonuniformly curved manifolds to mix two separate flows into a single, larger flow;

**[0015] FIG. 7A** shows a plurality of concentric manifolds transporting particles out of a region; and **FIG. 7B** shows a plurality of concentric manifolds transporting particles into a region;

**[0016] FIG. 8** is a representation of two particles moving in response to an externally applied field and an optical peristalsis pattern;

**[0017] FIG. 9** shows two stages of optical fractionation, with particles of a first type transported to the right and particles of a second type are transported to the left;

**[0018] FIG. 10** is a representation of the implementation of optical peristalsis using dynamical holographic optical traps;

**[0019] FIG. 11** shows a dynamic holographic optical trap system using a transmission-mode computer-addressed spatial light modulator in an optical train;

**[0020] FIG. 12** shows the mechanical cycling of a sequence of static computer-generated diffractive optical elements;

**[0021] FIG. 13** is a representation of a mechanically cycled optical peristalsis system using transmissive computer-generated diffractive optical elements arranged on the periphery of a disc;

**[0022] FIG. 14** shows a plurality of manifolds of optical traps trapping an extended object and rotating the object; and

**[0023] FIG. 15** shows the use of manifolds of optical traps trapping an extended deformable object.

#### DETAILED DESCRIPTION OF THE PREFERRED EMBODIMENTS

**[0024]** Optical peristalsis involves the use of a sequence of pre-calculated holograms projected over time to implement complex redistributions of large numbers of particles over large or selected areas. A key aspect of the invention of optical peristalsis is the non-specific transfer of particles from one manifold of optical traps in a given pattern to the next pattern through the intercession or application of at least two intermediate patterns. The term "pattern" is meant to encompass at least one manifold. **FIG. 1** shows a typical manifold **20** of optical traps **24** arranged in a straight line. Each of the traps **24** is capable of trapping a particle **22** of interest, and the traps **24** are spaced relative to each other so that the particle **22** is unlikely to pass through the manifold **20** without falling into an available one of the traps **24** or being blocked by particles already in the trap **24.** The particle **22** is drawn as a sphere, but could just as easily be irregularly shaped, or even much larger than the separation between the traps **24.** 

**[0025]** Operation of the optical peristalsis method proceeds by extinguishing the manifold **20** of the traps **24** which frees the particle **22** to move. If another pattern of the traps **24** is illuminated sufficiently nearby, then the particle **22** will be trapped by one (or more) of the traps 24 in the new pattern. In the illustrated case of **FIGS. 3A-3D** a pattern includes two of the manifolds **20** at line **23** and **25.** However, the next pattern could include only one of the manifolds, along line **27** for example. In effect, the particle **22** is thereby transferred from one of the manifolds **20** of the traps **24** in the first pattern **26** to another one of the manifolds **20** in a second pattern **28.** This process is in its simplist form depicted in **FIG. 2,** and shown more generally in **FIGS. 3A-3D.** To effect the transfer of the particle **22,** the first pattern **26** can be extinguished first; and then the second pattern **28** is illuminated, provided the interval between the two patterns **26** and **28** is short enough to prevent the trapped particle **22** from "wandering off'' ( out of the optical gradient) before it can be captured by the next, nearest available trap **24.** Illuminating the second pattern **28** before extinguishing the first pattern **26** also is another operative embodiment, albeit, more complicated to implement.

**[0026] A** pattern of the traps can therefore include one or more of the manifolds **20** of discrete the traps **24,** such as discrete tweezers in one embodiment of the invention. Each of the manifolds **20** can include several of the traps **24**  arranged along a one-dimensional curve or line, as shown schematically in **FIG. 1,** or also on a two-dimensional surface, or within a three-dimensional volume. The notion of a trapping pattern consisting of a collection of the manifolds **20** is useful for visualizing the process of optical peristalsis.

**[0027] FIG. 3A** shows in further detail one of the particles **22** trapped on one manifold **20** of a particular pattern, labeled as the first pattern **26.** The first pattern comprises two manifolds **50** and **56.** The positions of trapping the manifolds **52** and **54** in the second extinguished pattern **28** (only one manifold for this pattern) and a third extinguished pattern **30** ( only one manifold) are also shown. In the first time step, only the first pattern **26** is illuminated. In the next time step represented in **FIG. 3B,** the first pattern **26** is extinguished and the second pattern **28** is illuminated. This action transfers the particle **22** from the first manifold **50** of the first pattern **26** to the nearby manifold **52** of the second pattern **28.** In the next time incremental step shown in **FIG. 3C,** the second pattern **28** is extinguished and the third pattern **30** is illuminated, thereby transferring the particle **22**  again and this time to a manifold **54** on the third pattern **30.**  In the final time step as shown in **FIG. 3D,** the third pattern **30** is extinguished and the first pattern **26** is illuminated once again. This transfers the particle **22** to the first pattern **26** on the next manifold **56.** Optical peristalsis therefore arises from deterministically transferring the particle **22** from one of the manifolds **20** on a pattern of the optical traps to another of the manifolds **20** on the same second pattern **28**  by cycling through a sequence of intermediate patterns.

**[0028]** In a most preferred embodiment of the invention, a minimum of three of the patterns **26, 28** and **30** are needed to advance the particle **22** deterministically from the one manifold **50** on a trapping pattern to the next manifold **52.**  If only two of the equally spaced patterns **26** and **28** were used, the particle **22** could have a substantial probability of either advancing to the next manifold **52** or returning to the initial manifold **50.** In other embodiments, more than the three patterns **26, 28** and **30** can be used to transfer a particle **22** in a particular direction. Methods for illuminating and extinguishing the individual manifolds **20** of optical traps **24**  are well understood in the art.

**[0029]** Repeatedly cycling through the first, second and third patterns **26, 28** and **30,** respectively, tends to move the particles **22** from left to right in the arrangement described in **FIG. 3.** Reversing the sequence would drive them right to left. More extensive patterns consisting of more of the manifolds **20** thus can be used to transfer the particles **22**  back and forth over the entire field of view of the holographic optical trap system.

**[0030]** There are a variety of ways in which optical peristalsis can be used to effect useful rearrangements of collections of the particles **22.** These methods include modifying the shapes of the manifolds **20** within a pattern of the traps **24** by continuous curves. Although a single pattern is described in detail herein, additional intermediate patterns required for transfer between the manifolds **20** would be easily understood and recognized by those skilled in the art. In the examples described herein, the direction of particle flow will be indicated by overlaid arrows.

**[0031] FIG. 4** shows one of the patterns **26** from a linear optical peristaltic pump **33.** Two or more patterns (not shown) interleaved between the manifolds **20** of this pattern **26** can be activated in sequence to drive one or more trapped particles **22** from left to right. Reversing the sequence transfers the particles **22** from right to left. This pattern, and all of the patterns to be described herein, can be oriented in any desired direction

**[0032] FIGS. SA and 5B** show that patterns consisting of the curved manifolds **20** can be used to concentrate a flow of particles. Conversely, running the same sequence backwards disperses the particles **22.** This capability would be useful for directing the particles **22** out of an open region and into a confined region, such as a reservoir. It is not necessary that the individual manifolds **20** have equal curvature, and varying the curvature can be useful in particular situations. For instance, a linear pumping pattern can be used to sweep the particles **22** into a focusing pattern. The individual spacings between the manifolds **20** also do not have to be

equal. Regions of a pattern with more closely spaced forms of the manifolds **20** tend to transfer particles **22** more slowly than regions with more widely spaced ones of the manifolds **20.** The densely packed manifolds **20** tend to concentrate the particles **22** along the direction of motion, while widely spaced manifolds **20** can be used to spread them out. This approach could be particularly beneficial in a focusing pattern to avoid overcrowding the particles **22** as they are concentrated.

**[0033]** The distribution and density of the traps **24** along a manifold also can be used to control the flow of the particles **22** between the manifolds **20.** For instance, the traps **24** may be evenly spaced along each of the manifolds **20** and aligned simply from the one manifold **20** to the next and from one pattern to the next. In other embodiments, more complicated arrangements of the traps **24** along the manifolds **20** and between patterns can have uses for controlling the flow of particles **22** along a sequence of patterns. Similarly, varying the intensity, as well as the spacing, of individual traps **24** along the manifolds **20** in a pattern can have useful applications for controlling transport of the particles **22.** 

**[0034]** The tendency of the shaped manifolds **20** to direct the flow of the particles **22** can also be used to direct the particles **22** into any desired complicated pattern. The example shown in **FIG. 6A** shows the shaped manifolds **20**  directing one flow of the particles **22** into two. When run in reverse, such a pattern could be used to combine two (or more) flows into one. Although this may not be as efficient, because the particles **22** from one flow will remain near others from the same flow once the manifolds **20** merge, the methodology can still be used to advantage.

**[0035]** The example shown in FIG. of 6B shows one way to induce mixing of the particles **22** from combined flows. This example shows that the manifolds **20** in a pattern need not be disjoint. The patterns in this systems include a crossed form of the manifolds **20** in the mixing regions. Such crossings can be useful for exchanging the particles **22**  between the initially distinct flows. Crossing or otherwise intersecting the simple manifolds **20** to form more complex manifolds **20** introduces a probabilistic element into optical peristalsis. The particles **22** are given a choice of directions to travel near each crossing. Which direction the individual particles **22** choose to follow is determined by random thermal forces at the hand-off from one pattern to the next in a sequence. Hence, the crossings shown in **FIG. 6B** can lead to a certain degree of mixing.

**[0036] A** pattern of closed form of the manifolds **20,** such as the example shown in **FIGS. 7Aand 7B,** can transport the particles **22** into or out of a region. Whether the pattern compacts or rarefies the region depends on the order in which the sequence of patterns is projected. The example in **FIG. 7A** is useful for clearing the particles **22** out of a region, such as to facilitate tests on the suspending fluid or measurements on isolated particles **22.** Such patterns need not be circular, nor need they be confined to the plane. In principle, two-dimensional forms of the manifolds **20** in three-dimensional patterns can be useful for drawing material into a volume, or pushing material out of a volume.

**[0037]** Additionally, it should be noted that competition between optical trapping and other external forces can have useful applications. For example, competition between optical trapping and other external forces could be particularly useful in fractionating the particles **22** from a distribution. As an example, it is helpful to consider the particles **22**  entrained in a flow of surrounding fluid. Each of the particles **22** is transported by viscous drag in the local flow field  $\overline{u(r)}$ with a force  $\bar{f}$ = $\gamma u$  determined by its drag coefficient  $\gamma$ . For a sphere of radius  $\alpha$  in a fluid of viscosity  $\eta$ , the drag coefficient is given by  $\gamma = 6\pi\eta\alpha$  and increases linearly with the particle's radius. **A** larger particle feels a greater force when held stationary against a flow than a smaller particle. While the force due to viscous drag is one example of an external force, others such as those due to electric or magnetic fields also would pertain in this embodiment described herein.

**[0038]** If the external force is weaker than the optical gradient force of a given one of the optical traps **24,** then the particle **22** being transported by optical peristalsis will move much as described hereinbefore. If the external force is greater than the optical gradient force of the optical trap **24,**  then optical peristalsis may only perturb the motion of the particle **22** in the external field. In the idealized example shown in **FIG. 8,** one type of the particle **22** is more strongly attracted to the optical traps **24** than it is driven by the external field. In the example shown in **FIG. 8,** a first particle **60** is more amenable to trapping than a second particle **62** or is less strongly influenced by the external field than the second particle **62.** The first particle **60** is therefore transported by optical peristalsis and can be collected. The second particle **62** is more strongly driven by the external field and passes through the pattern of the traps **24,** perhaps being diverted to a certain extent from its initial course.

**[0039]** The two types of the particles **60** and **62** in the example embodiment shown in **FIG. 8** are distinguished either by their affinity for the optical traps **24,** by their response to the external field, or both. Choosing the spatial distribution, strength, and other characteristics of the optical traps **24** in such a pattern makes fractionation of particles possible, with the selectivity determined by the particles' differing physical characteristics.

**[0040]** The optical fractionation technique has a number of significant advantages. Fractionation occurs along the direction of the applied field in electrophoresis. Optical fractionation can transport the selected fraction laterally. This means that optical fractionation can operate continuously, rather than on one batch at a time. Because optical fractionation relies on holographic optical trap technology, it can be adapted readily to different fractionation problems.

**[0041]** For example, multiple stages of optical fractionation can be applied one after another using the same method and apparatus. Tuning each stage to extract a particular fraction of an initially mixed multicomponent sample then will separate the sample into each of its components, conveniently displacing the sorted components laterally away from the flow, and perhaps transporting them to channels or reservoirs using techniques previously described.

**[0042]** The example embodiment shown in **FIG. 9** builds on a single fractionation stage by including a second stage of optical fractionation. The external force driving the particles **22** through the region is directed downward. **A** first pattern, labeled **80** in **FIG. 9,** selects particles of first type **84**  and moves them to the right, diverting, but not collecting particles of second type **86.** The second stage of fractionation, labeled portion **82,** can feature more intense or more closely spaced examples of the traps **24** with the ability to divert particles **22** of the second type **86** away from the external force. As shown, this second stage pattern **82**  transports to the left, still further enhancing the separation between the fractions **84** and **86.** Although the two stages of fractionation are presented as conceptually separate, they could be implemented as a single pattern of the optical trap manifolds **20.** This process can also be generalized to include more stages and to incorporate transferring fractionated particles for collection.

**[0043]** As discussed above, optical peristalsis works by repetitively cycling through a sequence of trapping patterns. The dynamic holographic systems represented schematically in **FIGS. 10 and 11** are a fully general implementation. In this case, a computer-addressed spatial light modulator **102** creates the configuration of laser beams **104** needed to implement a given pattern of optical traps **114** by encoded the necessary phase modulation onto the wavefront of an input laser beam **100.** In principle, such a system can implement any sequence of trapping patterns, and thus any variant of optical peristalsis. In practice, however, the spatial light modulator **102** has physical limitations such as spatial resolution which limit the complexity of the patterns which they encode. Also, such spatial light modulators **102** tend to be costly.

**[0044]** In the embodiment shown in **FIG. 10,** optical peristalsis can be performed with the dynamical holographic optical traps **114,** a typical implementation of which is shown. An input laser beam **100** is reflected off the surface of the computer-addressed spatial light modulator (SLM) **102.** The SLM **102** encodes a computer-generated pattern of phase shifts onto the wavefront of the beam **100,** thereby splitting it into one or more separate laser beams **104,** each emanating from point **107** in the center of the face of the **SLM 102.** Lenses **108** and **110** relay each of these laser beams **104** to the conjugate point **112** at the center of the back aperture of a high NA objective lens **112.** This objective lens **112** focuses each of the laser beams **104** into a separate optical trap **114,** only one of which is shown in **FIG. 10** for clarity. **A** dichroic mirror **116** reflects trapping light into the objective lens **112** while allowing imaging illumination to pass through, thereby permitting images to be formed of the particles being trapped. Updating the phase modulation encoded by the SLM **102** causes a new pattern of the traps **114** to appear. Cycling through a sequence of optical peristalsis patterns in this manner implements the corresponding optical peristalsis process. Because this system can be reconfigured in software, it represents a general implementation of optical peristalsis. In another embodiment shown in **FIG. 11,** the dynamic holographic optical trap system uses a transmission-mode computer-addressed spatial light modulator **200** in an optical train otherwise similar to that in **FIG. 10.** This system also can be used to implement optical peristalsis by cycling through a sequence of trapping patterns.

**[0045]** Implementing optical peristalsis does not necessarily require the generality and reconfigurability offered by a dynamic holographic optical trap system. Instead, implementing optical peristalsis preferably uses a holographic optical trap system capable of projecting a (small) sequence of otherwise static patterns. In its simplest preferred form, optical peristalsis can be implemented by mechanically cycling through a sequence of phase patterns to implement a corresponding sequence of holographic optical trapping patterns. One particularly useful embodiment appears in **FIG. 12.** As shown in **FIG. 12,** the phase patterns needed to implement a particular optical peristalsis process are encoded in the surface relief of reflective diffractive optical elements **304, 306** and **308.** These elements **304, 306,** and **308** are mounted on the face of a prism **300,** and each is rotated into place by a motor **302.** Reversing the motor's rotation reverses the sequence of patterns and thus the direction of optical peristalsis. Rotating the prism **300** with the motor **302** orients each of the patterns in the input laser beam so that the diffracted beams created by the aligned diffractive optical elements **304, 306** and **308** all create optical traps **114.** Stepping the motor **302** through each of the patterns in sequence implements optical peristalsis. Prisms with more than three patterns can be employed, if desired or necessary.

**[0046]** Mounting a sequence of fixed reflective diffractive optical elements **304, 306** and **308** on the face of a rotating prism **300** can have other uses in holographic optical trap methodologies. Similarly, transmissive diffractive optical elements **404, 406, 408** and **410** can be located on the periphery of a disk **312** and rotated into the beam **100,** as shown in **FIG. 13,** or into a reflective optical train in sequence. This also has potential applications beyond optical peristalsis. In **FIG. 13,** for example, each of the diffractive optical elements **404, 406, 408** and **410** is rotated into the optical train to project one pattern of the optical peristalsis sequence.

**[0047]** Static reflective or transmissive diffractive optical elements can be fabricated with feature sizes down to the diffraction limit, can have essentially continuous phase encoding, and thus can implement a wider variety of more complicated trapping patterns than can spatial light modulators. Such elements can be produced much more cheaply and do not require a computer to operate. The sequence of patterns in such a system can be changed by changing the prism or disk of diffractive optical elements. In this sense, this implementation is less general than that based on computer-addressed spatial light modulators.

**[0048]** Because only a small number of precalculated diffractive optical elements are required to implement optical peristalsis, switchable phase gratings also can be used. The benefits of such an approach include, for example: freedom from moving parts which can drift out of alignment and wear out, the absence of motors which cause vibration and radiate stray electric and magnetic fields, reduction in power requirements and improved compactness.

**[0049]** Encoding high-quality phase holograms on film media will allow optical peristalsis to be implemented with the equivalent of film loops. By offering high-speed cycling through large numbers of diffractive optical elements, filmbased implementations of holographic optical traps will have applications beyond optical peristalsis.

**[0050]** Optical peristalsis also can be useful for particles and other materials such as biological cells which are larger than the physical separation between the traps in an optical peristalsis pattern. Similarly, materials such as proteins, **DNA,** or molecules could also be manipulated using optical peristalsis. **A** large object trapped on a "bed of nails" optical trapping pattern still can be moved by translating the bed of nails. Rather than defining a single trapping region, however, an optical peristalsis pattern can establish a large field of traps suitable for immobilizing a large object wherever it is found. Updating the pattern with small displacements, as described above, then will displace the entire object. Potential applications include translating an extended sample into a region where it can undergo tests, rotating the object for examination, or controllably deforming the object. For example, in the embodiment of **FIG. 14,** the manifolds **20** of included optical traps are shown trapping an extended object **80.** Updating the pattern with the manifolds **20** will tend to rotate the extended object **80.** Similarly, **FIG. 15** shows the manifolds **20** of optical traps trapping an extended deformable object **82.** The object **82** is more strongly trapped by denser regions of traps, and moving these regions outward in subsequent patterns tends to stretch the object **82.** 

**[0051]** Each optical peristalsis sequence performs one specific operation. In some applications, it can be desirable to perform a series of optical peristalsis operations, with the order of the series perhaps depending on the outcome of the preceding operations. For example, optical peristalsis can be used to move a living cell into the center of a microscope's field of view for reproducible observation. A second sequence then could be engaged to rotate the cell into a desired orientation. Then a third sequence can implement a particular test. Based on the outcome of that test, additional optical peristalsis sequences can be selected to collect the cell or dispose of it. Each of these sequences can be precalculated, thereby removing much of the computational burden from the holographic optical trap system. Similarly, different subsequences of optical peristalsis operations could be incorporated into a single program, wherein a first subsequence could separate particles into two or more distinct flows, a second subsequence could disperse particles from a particular location, a third subsequence could mix two separate streams of particles into a single flow, a fourth subsequence could concentrate a plurality of particles into a particle region, and particles can be "moved" from pattern to pattern in a variety of other ways as well. A variety of combinations of subsequences such as those described herein could be incorporated into a single program, and these subsequences could be used sequentially and/or simultaneously as needed using a variety of types of optical gradients as described herein. Because very few diffractive optical elements are required to implement any one of the sequences, only modest elaboration of the proposed implementations would be needed to select among a collection of available sequences for such multistage operations.

**[0052]** Additionally, it is also possible to practice the present invention without the use of optical traps as conventionally understood to require specific optical gradient conditions to hold a particle. For example, a plurality of deterministic optical gradients can be established and incorporated into a plurality of manifolds and patterns as generally described above. These optical deterministic gradients operate to "hold" or restrain, but not necessarily form an optical trap, for individual particles in a particular position for a sufficient period of time in sequence to generate an optical peristalsis effect. In other words, repeatedly cycling through first, second, and third patterns of deterministic optical gradients will move individual particles along a designated path. The optical gradients are deterministic in a sense that the conditions that are applied are sufficient to achieve the intended result with more than just a mere probability of success.

**[0053]** While preferred embodiments of the invention have been shown and described, it will be clear to those skilled in the art that various changes and modifications can be made without departing from the invention in its broader aspects as set forth in the claims provided hereinafter.

**1.** A system for manipulating a plurality of particles, comprising:

- a plurality of beams of laser light, the beams of laser light establishing a plurality of optical traps;
- a plurality of sequentially spaced patterns with each of the patterns comprising at least one optical trap formed from at least one of the beams of laser light; and
- an optical component sequentially illuminating and extinguishing each of the patterns using the beams of light at intervals close enough after the extinguishing of the previous pattern enabling capture and transfers of a particle from one of the patterns to another causing the particle to travel from one of the patterns to the next one of the patterns.

\* \* \* \* \*