

[0075] Turning now to configurations of the sorting station, certain buffer switching designs will now be described. As explained, buffer switching employs separate streams of buffer and sample that are delivered to a sorting region. The force exerted on magnetic particles in the sorting region guides those particles out of the sample stream and into the buffer stream. Due to the low Reynolds number, the flow streams within the sorting region of a CMACS device are generally uniaxial and laminar (e.g., Re is approximately 2000 or less). Buffer switching regions may be designed such that in the absence of an external magnetic field, only the buffer medium arrives at a collection channel (see FIG. 1). When an external field is applied, the MFG elements become magnetized and the magnetically labeled components are selectively transported across the stream boundary, from the inlet stream into the buffer medium, resulting in a high purity of the target species in the collection channel. On the other hand, the unlabeled components (typically the majority of the components in the sample) are not deflected by the magnetic field and will enter the waste channels.

[0076] In buffer switching devices, the separate buffer and sample streams may be provided to the sorting region via one, two or more separate channels. Typically, though not necessarily, a sorting region will have at least one inlet channel dedicated to delivering buffer and another channel dedicated to delivering sample. In certain embodiments, at least one inlet channel to the separation region provides both the sample stream and the buffer stream. Because these streams are provided as laminar flows, they can be combined into an inlet channel upstream from the sorting region. They will then flow into the sorting region separated from one another as separate streams.

[0077] FIGS. 4A-4E present various buffer switching configurations. Each figure shows a distinct sorting region with at least one inlet channel and at least one outlet channel. Further each sorting region has at least one MFG, sometimes shown.

[0078] FIG. 4A shows a simple configuration in which a buffer switching configuration 401 includes one inlet channel 405, one outlet channel 407 and a sorting region 403. The inlet channel 405 includes two laminar streams, a buffer stream shown in the lower portion of channel 405 and a sample stream shown in the upper portion of channel 405. The streams maintain their uniaxial flow trajectory through sorting region 403. At least one MFG (not shown) within region 403 imparts a "rightward" velocity component to magnetic particles entering through the sample stream in inlet channel 405. As a consequence, the magnetic particles are directed into the buffer stream and then exit through a buffer stream portion 411 of outlet channel 407. The non-magnetic components of the input sample remain in the sample stream and exit the sorting region 403 via a sample stream portion 409 of outlet channel 407.

[0079] FIG. 4B shows a similar buffer switching configuration 413 but with two separate outlet channels: a dedicated waste or sample channel 419 and a dedicated buffer channel 421. In this embodiment, separate sample and buffer streams enter the lower and upper portions of an inlet channel 417. Together they flow into a sorting region 415 where at least one MFG (not shown) diverts any magnetic particles in the sample stream "rightward" into the buffer stream. The buffer stream, with diverted magnetic particles, exits the sorting region via a dedicated buffer exit channel 421. The sample stream, which has been depleted of magnetic particles, exits the sorting region via a dedicated waste exit channel 419.

[0080] FIG. 4C shows another buffer switching configuration 423, this time with two inlet channels and a single outlet channel. In this embodiment, a sample stream is provided via a dedicated sample inlet channel 427. Similarly, a buffer stream is provided via a dedicated buffer inlet channel 429. The sample and buffer streams from the two inlets enter a sorting region 425 together, but remain as distinct uniaxial streams. Within the sorting region, at least one MFG imparts a downward velocity component to the magnetic particles in the sample stream. As a consequence, some or all magnetic particles in the sample stream cross the boundary between the sample and buffer streams and enter the buffer stream. The two streams exit the sorting region 425 together through an outlet channel 431. Within this channel a waste stream portion 433 carries the remainder of the sample stream, which has been depleted of magnetic particles. Also within outlet channel 431, a buffer stream portion 435 carries the buffer with the magnetic particles. As should be apparent, the contents of the outlet channels in the embodiments of FIGS. 4A and 4C must be separately treated downstream from the depicted sorting configurations. This may be accomplished, for example, via a downstream divide in the flow path or a chamber having separate treatment regions for the sample and buffer streams (e.g., a capture region for magnetic particles in the buffer stream).

[0081] FIG. 4D shows a microfluidic sorting channel configuration 437 having three separate inlet channels and two outlet channels. Sample is provided via an inlet channel 441, while buffer is provided via two separate inlet channels 443 and 445 located below the sample inlet channel. Sample enters a sorting region 439 having two separate MFGs 447 and 449 arranged in series. Other MFG arrangements are possible in similar multi-channel sorting configurations. The MFGs in FIG. 4D divert magnetic particles from the sample stream (provided via inlet channel 441) toward and into the buffer stream(s) from inlet channels 443 and 445. The buffer stream(s) exit the sorting region via an outlet channel 453, while the sample stream (with a diminished or depleted concentration of magnetic particles) exits via a waste outlet channel 451. Note that the embodiment depicted in FIG. 4D allows for multiple buffer streams having potentially different compositions. In other embodiments, more buffer and/or sample inlet channels may be employed; e.g., three buffer inlet channels and two sample inlet channels may be employed. Further, in some embodiments, the number of sample inlet channels may be the same or greater than the number of buffer inlet channels; e.g., three buffer inlet channels and three sample inlet channels. Similar variations on the arrangement of outlet channels may be realized.

[0082] FIG. 4E depicts yet another arrangement 455 of channels for sorting. This embodiment provides a generic version of the sorting station depicted in FIG. 1. In the arrangement of FIG. 4E, two sample streams enter a sorting region 457 via inlet sample channels 459 and 461. These inlet channels straddle a buffer inlet channel 463. Thus, a buffer stream flows uniaxially through the sorting region along with sample streams on either side. In this embodiment, a first MFG 465 exerts a magnetophoretic force on magnetic particles entering from sample inlet channel 459, directing them downward into the central buffer stream provided via inlet 463. A second MFG 467 exerts a magnetophoretic force in the opposite direction, deflecting magnetic particles entering from sample inlet channel 461 upward into the central buffer stream. Thus, two parallel acting MFGs provide a high