

# On magnetophoretic separation of blood cells using Halbach array of magnets

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Abstract Magnetophoretic separation has gained much attention in recent years due to its easy application and low-cost fabrication compared to other active particle separation techniques. Due to the different properties of white blood cells (WBCs) and red blood cells (RBCs), it is possible to manipulate and separate them using a magnetic field. In this paper, a simple microfluidic device is proposed to fractionate WBCs and RBCs from whole blood using magnetophoretic force applied by Halbach array of three permanent magnets. Plasma streams containing WBCs and RBCs enter a simple microchip fabricated by PDMS. Permanent magnets apply positive and negative magnetophoretic forces to the RBCs and WBCs, respectively. Two cladding streams containing blood plasma are used to concentrate the cells in the magnetophoretic area. A wide range of inlet velocities and different distances of magnets from the channel (d) are investigated. It is demonstrated that the volume flow rate of core, and cladding streams, total flow rate and the distance between magnets and microchannel affect the separation efficiency individually. The results reveal that  $d = 0.1, 0.2, 0.3, 0.4,$  and 0.5 mm may lead to complete separation when core and cladding flow rates are 1 and 7  $\mu$ l/h, respectively.

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## 1 Introduction

Microparticle separation is of great importance in the diagnostic, chemical, and biological analysis, as well as food processing and environmental assessment. The concept of separation can be generalized to genetic engineering, cellular transplantation, and immunology. Many investigations have been performed to improve microfluidic techniques to miniaturize the microfluidic devices. Particle separation based on microfluidic technology offers a simple, reliable, and cost-effective approach to disease detection. A microfluidic device can distinguish dead or cancerous cells from living and normal cells in the blood and separate them. Using unique micro-flow phenomenon features, various methods have been developed for continuous and accurate separation of microparticles. In general, particle separation is performed by two active and passive techniques. Passive methods use interactions between particles, microchannel structure, and fluid flow field to separate particles. These methods are easily measurable and do not depend on external forces, but their operating range is small due to their fixed geometry  $[1-3]$ . Common methods of passive separation include microfiltration [[4–6\]](#page-11-0), inertial separation [[7–10\]](#page-11-0), deterministic lateral displacement (DLD) [\[11](#page-11-0)[–14](#page-12-0)] and compact flow method [[15–17\]](#page-12-0). Active methods are used to separate the particles using an external field such as electric, magnetic, optical, and acoustic fields. Active methods have more control over the separation processes and are more reliable. The separation efficiency of active methods is also greater than the passive ones. Active separation methods include dielectrophoresis [\[18–23](#page-12-0)], acostophoresis [\[24–27](#page-12-0)] and magnetophoresis [\[28](#page-12-0)[–63](#page-13-0)]. Magnetophoresis-based separation has received much attention in recent years due to its easier application and lower cost than other active separation methods [\[58](#page-13-0)]. For this reason, this technique is described in detail.

Magnetophoresis is a phenomenon in which the particle migrates in a magnetic field. This phenomenon can be divided into two general categories: positive and negative magnetophoresis. Positive magnetophoresis is the movement of a magnetic particle in a diamagnetic environment, and negative magnetophoresis is the migration of a diamagnetic particle in a magnetic environment. Magnetophoretic force is created when either the magnetic field gradient or the magnetization gradient of the environment, or a combination of both exist. This technique does not affect the properties of the sample solution, including pH, ion concentration, surface charge, and temperature. Besides, this method is easy to use and inexpensive. The key parameters for magnetophoretic applications are magnetic permeability  $(\mu)$ , magnetic flux density (B), and susceptibility ( $\chi$ ). The magnetic field is provided in the microfluidic devices using an electromagnet or permanent magnet. An electromagnet can produce a higher magnetic field gradient than a permanent one, which increases the separation efficiency. However, the permanent magnet is simpler to use and does not generate heat in the sample. An array of permanent magnets can be put together to increase the magnetic gradient. Magnetic susceptibility of particles and their surroundings is another important parameter in particle separation. Based on the susceptibility, the materials are divided into three categories: ferromagnetic  $(\chi \gg 0)$ , paramagnetic  $(\chi > 0)$ , and diamagnetic ( $\chi$  < 0). The magnetic susceptibility of the particles can be changed by their labeling using magnetic beads. Particle labeling increases the cost and time of the separation process. Negative magnetophoresis is a method without labeling to separate diamagnetic particles such as cells in a paramagnetic medium, for example, ferrous solutions. The diamagnetic particles in the ferrofluids act as magnetic holes. If the system is exposed to an external magnetic field, the imbalance of the particle and fluid susceptibility creates a negative magnetophoretic force, leading to that the particles move away from the magnet. The negative magnetophoretic force is also proportional to the particle size so that the particles can be separated due to their size [[45,](#page-12-0) [55](#page-13-0), [57](#page-13-0)]. Negative magnetophoresis has attracted considerable attention in recent years in the fields of disease diagnosis, therapeutic studies, concentration, and cell fractionation [[58\]](#page-13-0).

Pamme and Mans [[29\]](#page-12-0) proposed a free-flow magnetophoresis for continues flow separation of magnetic particles. In this method, the sample enters the channel and is driven by a cladding flow toward the channel wall. Then, a magnetic field perpendicular to the direction of fluid flow is applied to the magnetic particles. Non-magnetic particles move directly to the channel outlet, while magnetic ones move transverse and are separated based on their size and magnetization. Adams et al. [[34\]](#page-12-0) developed a method for particle separation using multi-target magnetic cell separator. They separated magnetic particles with an output power of 10<sup>9</sup> cells per hour and reached a purity of about 90%. Mizuno et al. [\[41](#page-12-0)] used a combination of hydrodynamic filtration and magnetophoresis to separate lymphocyte cells. In this method, the cells are first divided into two groups by a size-based hydrodynamic filter, and then each group is separated due to their magnetic labels. Nam et al. [\[42](#page-12-0)] used a magnetic field to separate malaria-infected red blood cells and achieved 98.3% separation efficiency. Zhu et al. [\[43\]](#page-12-0) used a combination of positive and negative magnetophoretic forces to separate the magnetic particles from non-magnet ones and particles with different magnetizations. Zhang et al. [\[49](#page-12-0)] proposed a microfluidic device based on negative magnetophoresis to separate 4- and 13-um polystyrene particles using a 4-mm neodymium magnet at a distance of 1 mm from the channel. Hejazian and Nguyen [[47\]](#page-12-0) experimentally separated non-magnet polystyrene particles of sizes 3.1 and 4.8 µm suspended in a paramagnetic solution flowing through a straight microchannel of 1 mm width and 500 µm height. They employed an array of permanent magnets on either side of the channel. Tarn et al.  $[52]$  $[52]$  separated 5- and 10- $\mu$ m diamagnetic particles based on their size and magnetization in a paramagnetic solution. They reported that the deviation of particles concerning the channel center increases along the microchannel by increasing their size and magnetization. Munaz et al. [[58\]](#page-13-0) designed a negative magnetophoresis-based microfluidic device for the separation of fluorescent polystyrene microparticles with diameters of 3.2 and 4.8  $\mu$ m using two different geometries using two-stream and threestream structures. Experimental results showed that the highest separation efficiency corresponds to the three-stream structure. It was 78% and 75% for the 3.2- and 4.8-µm particles, respectively. Wu et al.  $[60]$  $[60]$ investigated a microfluidic device based on negative magnetophoresis for separating of  $5-$  and  $12-\mu m$ fluorescent polystyrene particles experimentally and numerically. They concluded that the Halbach array results in better performance than the other configuration. They also reported that the deflection angle increases with flow rate, magnetic field, and particle size.

Magnetic separation of blood cells using microfluidic devices has been previously reported [\[28](#page-12-0), [30–33](#page-12-0), [36](#page-12-0), [37](#page-12-0), [39](#page-12-0), [62](#page-13-0), [63](#page-13-0)]. Due to the different properties of WBCs and RBCs, it is possible to manipulate and separate them using a magnetic field [\[39](#page-12-0)]. Han and Frazier [\[30,](#page-12-0) [31](#page-12-0)] modeled a magnetophoretic separator to separate WBCs and RBCs based on their intrinsic magnetic properties. Furlani [\[33](#page-12-0)] proposed a continuous and label-free method for separating of WBCs and RBCs in plasma using magnetic force. Seo et al. [[36,](#page-12-0) [37\]](#page-12-0) used a combination of hydrodynamic forces and magnetophoresis to increase the output power of blood cell separation. Their experimental results showed that separation efficiency depends on the magnetophoretic force.

In the present paper, the separation of WBCs and RBCs from whole blood is simulated numerically using a magnetophoretic force using Halbach array of three permanent magnets. Two cladding flows are used to concentrate the particles in the area under the magnetic field. To the best of our knowledge, the separation of bold cells has not been carried out using Halbach arrays of magnets. On the other hand, the proposed microdevice can be easily fabricated, and its separation efficiency reaches 100% for a relatively wide range of throughputs. The numerical results are analyzed by changing the magnet array distance from the channel and the ratio of inlet streams.

#### 2 Governing equations

Figure 1 shows a schematic of the microfluidic device. WBCs and RBCs suspended in the blood plasma enter the microfluidic device and are separated by a magnetophoretic mechanism. The main microchannel has a rectangular cross-section with the length of  $L = 20$  mm, the width of  $W = 200$  µm, and depth of  $H = 20 \mu m$ . Whole blood containing plasma and WBCs and RBCs is injected into the microfluidic chip using a syringe pump with  $Q = 5 \mu L/h$ . Two cladding flows containing plasma without WBCs and RBCs are injected into other entrance channels to concentrate blood cells in the central area of the main channel. The described symmetric microchannel can be easily fabricated using Polydimethylsiloxane (PDMS) [\[67](#page-13-0)]. To produce a non-uniform magnetic field, an array of three NdFeB permanent magnets with dimensions of  $5 \times 5 \times 5$  mm<sup>3</sup> is placed in the distance d from the microchannel wall, as shown in Fig. [2.](#page-3-0) In this study, the magnets are positioned alongside the microchannel in the Halbach array. This arrangement strengthens the magnetic field on one side of the magnet [[60\]](#page-13-0).

The magnetophoretic separation of cells is a combination of electromagnetic, fluidic, and particle dynamics [[39\]](#page-12-0). To solve the magnetic field statically (magnetostatic equations), the Maxwell-Ampere's law is employed for the magnetic field [[44,](#page-12-0) [49](#page-12-0), [58,](#page-13-0) [64](#page-13-0)]:



Fig. 1 Magnetophoretic separation of WBCs and RBCs within blood plasma: a Schematic of the microfluidic device, and b structure of microchip along with its inlets and outlets

<span id="page-3-0"></span>

Fig. 2 Geometrical parameters of the problem (yellow arrows indicate the direction of magnetization)

$$
\vec{H} = -\nabla V_m \tag{1}
$$

where H is the magnetic field strength, and  $V_m$  is scalar magnetic potential. Using Gauss's law for magnetic flux density B, the following equation is concluded [\[44](#page-12-0), [49](#page-12-0), [58,](#page-13-0) [64\]](#page-13-0):

$$
\nabla \cdot \vec{B} = 0 \tag{2}
$$

The relationship between  $B$  and  $H$  in the fluid and in the magnet domain is shown in Eqs. 3 and 4, respectively [\[44](#page-12-0), [49,](#page-12-0) [58,](#page-13-0) [64](#page-13-0)]:

$$
\vec{B} = \mu_0 \mu_r \vec{H} \tag{3}
$$

$$
\vec{B} = \mu_0 \left( \vec{H} + \vec{M} \right) \tag{4}
$$

where  $\mu_0 = 4\pi \times 10^{-7}$  (T m/A) (T m/A) is the air permeability coefficient,  $\mu_r$  is the relative magnetic permeability of the fluid or environment, and  $M(A/m)$ is the magnetization of the magnet. It should be noted that the relative magnetic permeability can be calculated as follows [\[44](#page-12-0), [49,](#page-12-0) [58,](#page-13-0) [64](#page-13-0)]:

$$
\mu_r = 1 + \chi \tag{5}
$$

The governing equations of the fluid flow are the continuum and Navier–Stokes equations:

$$
\nabla \cdot \vec{u}_f = 0 \tag{6}
$$

$$
\rho_f \vec{\mathbf{u}}_f \cdot \nabla \vec{\mathbf{u}}_f = -\nabla p + \mu_f \nabla^2 \vec{\mathbf{u}}_f \tag{7}
$$

where  $u_f$  is the fluid velocity (blood plasma),  $\mu_f$  is the dynamic viscosity,  $\rho_f$  is the fluid density, and p is the fluid pressure. Plasma is considered to be incompressible Newtonian fluid. The properties used for plasma and blood cells are presented in Table [1.](#page-4-0)

Since the density of cells is close to the cladding solution (the blood plasma), the dominant force

governing the particles is the hydrodynamic drag force  $(F_D)$  [[29,](#page-12-0) [55\]](#page-13-0) and magnetophoretic force  $(F_M)$ . The particle trajectory is described using Newton's second law:

$$
m_P\left(d\vec{u}_p/dt\right) = \vec{F}_D + \vec{F}_M\tag{8}
$$

where  $m_P$  is the particle mass and  $u_p$  is its velocity. Since the flow rate in microfluidic devices is very low, the flow is assumed to be laminar. The Reynolds number for the present work is  $Re = 0.05$ , which confirms that the flow regime is laminar.

$$
Re = \rho_f U_{in} D_h / \mu_f \tag{9}
$$

where  $U_{in} = Q/WH$  is inlet velocity and  $D_h$  is hydraulic diameter defined as follows:

$$
D_h = 2WH/(W+H) \tag{10}
$$

Hydrodynamic drag force is calculated using Stokes' law by assuming blood cells as spherical microparticles for low Reynolds numbers [\[33](#page-12-0), [35\]](#page-12-0):

$$
\vec{F}_D = 3\pi d_p \mu \left(\vec{u}_f - \vec{u}_p\right) \tag{11}
$$

The magnetophoretic force is calculated using Eq. 12 [[35,](#page-12-0) [45](#page-12-0), [52\]](#page-13-0) by neglecting the magnetoviscosity effects for microparticle concentrations less than 1% [\[65](#page-13-0)]:

$$
\vec{F}_M = \frac{V_p}{\mu_0} \left( \chi_p - \chi_f \right) \left( \vec{B} . \nabla \right) \vec{B} \tag{12}
$$

where  $V_P = \frac{1}{6}\pi d_P^3$  is the volume of the particle. By placing Eqs. 3 and 5 in Eq. 12, the magnetophoretic force is [\[49](#page-12-0), [58](#page-13-0)]:

$$
\vec{F}_M = 2\pi r_p^3 \mu_0 \mu_{rf} K \nabla \vec{H}^2 \tag{13}
$$

	$\rho$ (kg/m <sup>3</sup> )	$\rho$ (kg/m <sup>3</sup> )	$\mu$ (Pa s)	X	$d_{\rm P}$ (µm)
Plasma	1000	1000	0.001	$-7.7 \times 10^{-6}$	$\qquad \qquad -$
<b>WBCs</b>	1070	1070	-	$-9.9 \times 10^{-6}$	10
<b>RBCs</b>	1100	1100	-	$-3.9 \times 10^{-6}$	-

<span id="page-4-0"></span>Table 1 Physical properties of blood contents [[33](#page-12-0), [62](#page-13-0), [63\]](#page-13-0)

$$
K = \frac{\mu_{r,p} - \mu_{rf}}{\mu_{r,p} + 2\mu_{rf}}\tag{14}
$$

where  $r_P$  is the radius of the particle.  $\mu_{r,f}$ , and  $\mu_{r,p}$  are the relative magnetic permeability of the fluid and the particle, respectively. Since the direction of the magnetic field is perpendicular to the direction of the fluid flow in microfluidic devices, the particle velocity  $u_p$  can be obtained by equating drag and magnetophoretic forces under equilibrium position [[45,](#page-12-0) [52](#page-13-0)]:

$$
\vec{F}_D + \vec{F}_M = 0\tag{15}
$$

By placing the definition of  $\vec{F}_D$  from Eq. [11](#page-3-0) in Eq. 15,

$$
3\pi d_p \mu \left(\vec{u}_f - \vec{u}_p\right) + F_M = 0, \qquad (16)
$$

the particle velocity can be calculated as follows:

$$
\vec{u}_p = \vec{u}_f + \left(\vec{F}_M/3\pi d_p\mu_f\right) \tag{17}
$$

The rate of cell collection or cell separation efficiency can also be calculated using Eq. 16 [\[36](#page-12-0), [39](#page-12-0), [58\]](#page-13-0):

$$
SE(\%) = \frac{N_O}{N_T} \times 100\tag{18}
$$

where  $N_O$  and  $N_T$  are numbers of cells passing through the outlet and the total number of cells entering the microchannel, respectively.

The governing equations are solved by the finite element method using COMSOL Multiphysics 5.5 software. The relative magnetic permeability of the space around the microchannel and the magnets is set to one, and the magnetization of the magnets is  $M = 9.9 \times 10^5$  A/m [[60\]](#page-13-0). No-slip boundary condition is imposed on the microchannel walls, and constant pressure condition is used for the output. WBCs and RBCs enter the microchannel at an initial velocity equal to the fluid velocity through the central inlet channel.

## 3 Validation

To validate the numerical method, the results of the present simulation are compared with those of He et al. [\[66](#page-13-0)], who investigated the effect of magnetic manipulation on label-free nonmagnetic particles (Figs. 3 and [4](#page-5-0)). Figure 3 shows the magnetic field's strength in terms of channel length along the centreline of a straight microchannel containing diamagnetic microparticles within a magnetic fluid exposed to a non-uniform magnetic field generated by a permanent magnet. Figure [4](#page-5-0) presents the corresponding magnetic force. The distribution of negative magnetophoretic force along and perpendicular to the fluid stream, i.e.,  $F_{mx}$  and  $F_{my}$ , respectively, is compared with the reference results, indicating very good agreement.

For more verification, the separation of diamagnetic microparticles suspended in a conventional ferrofluid with poly ethylene oxide (PEO) flowing in a straight microchannel exposed to a permanent magnet is compared with numerical and experimental results of Zhang et al. [[49\]](#page-12-0). The ratio of lateral velocity to the horizontal velocity of microparticles along the microchannel is shown in Fig. [5](#page-5-0) for different inlet flow rates. It should be pointed out that the PEO-based



Fig. 3 Magnetic field strength along the channel centerline

<span id="page-5-0"></span>

Fig. 4 Magnetophoretic force along the channel centerline



Fig. 5 The ratio of lateral velocity to horizontal velocity of microparticles along the microchannel

ferrofluid exhibits viscoelastic behavior. However, as Zhang et al. [[49\]](#page-12-0) reported, the effect of elastic force can be neglected in the present simulations due to the low channel aspect ratio. Besides, in Fig. 6, the separation of  $5-$  and  $13-$ µm magnetic particles within the PEO-based ferrofluid is compared with the numerical results of Zhang et al. [\[49](#page-12-0)]. They carried out the numerical modeling of magnetophoretic separation of microbeads using COMSOL Multiphysics 5.1. In this figure, the colored contours show the distribution of the magnetic field strength. It is demonstrated that larger microbeads are repelled more quickly toward the opposite wall compared to the smaller ones. As can be seen, there is a very good agreement between the present results and those of Zhang et al. [\[49](#page-12-0)].



Fig. 6 Numerical simulation of magnetophoretic separation of 5- and 13-um microparticles suspended in the PEO-based ferrofluid (color contour indicates the magnetic field strength, A/m): **a** present work, and **b** the results of Zhang et al. [\[49\]](#page-12-0)

#### 4 Grid study

The structured grid is used for the microchannels, and a triangular one is employed for the surrounding space (Fig. [7](#page-6-0)). The average magnetic field strength along the microchannel centerline, lateral velocity, and lateral position of the WBCs at the output of the main channel are presented in Table [2](#page-6-0) for the grids with 25,000, 90,000, 200,000, and 350,000 elements. The results demonstrate that the grid resolution of 200,000 can be selected for further simulations.

## 5 Results

The distribution of the magnetic field for the magnet with the distance of  $d = 1$  mm to the microchannel is shown in Fig. [8.](#page-6-0) In this figure, three magnets with a Halbach arrangement are positioned at different distances from the microchannel wall. This arrangement extends the magnetic field from one side to the farther distances. The red lines represent the distribution of the induced magnetic field. In the Halbach array of magnets, the magnets' polarization angles rotate as much as 90°. The yellow arrows show the direction of the magnetization.

In Fig. [9,](#page-6-0) the distribution of the magnetic field density along the microchannel centreline is plotted for different distances of the magnet array from the channel. As can be seen, the magnitude of the magnetic field decreases by increasing the distance from the microchannel. The magnetic field peak points

<span id="page-6-0"></span>

Fig. 7 The grid used for present simulations

Table 2 The results of the grid study

Number of elements	Averaged magnetic strength field along the centerline H $\times$ 10 <sup>5</sup> (A/m)	Lateral velocity of RBCs in the Lateral position of RBCs in outlet $(\mu m/s)$	the outlet $(\mu m)$
25,000	4.5196	21	153
90,000	4.5196	45	147
200,000	4.5196	45	146
325,000	4.5196	46	146



Fig. 8 The density distribution of the magnetic field where the magnet array has a distance of  $d = 1$  mm from the microchannel. The red lines represent the induced magnetic field, and the yellow arrows indicate the direction of the magnetization

are created where the induced magnetic field of central magnet reaches those of side magnets. As the magnet array distance from the microchannel decreases, the value of these peak points becomes larger.

A strong magnetic field is required to separate these cells due to the slight difference in the relative magnetic permeability coefficient of plasma and blood cells. Thus, the distance between the array of magnets



Fig. 9 Magnetic field density  $B(T)$  along the microchannel centreline for different distances of magnet array from the microchannel (d)

and main channel is assumed to be  $0 \le d \le 1$  (mm). The relative permeability coefficient of RBCs is greater than that of plasma. Hence, a positive magnetophoretic force is exerted on RBCs under a nonuniform magnetic field, leading to their migration toward higher magnetization region, i.e. magnets. For

the case of WBCs, a negative magnetophoretic force pushes them toward the farther microchannel outlet due to their lower relative magnetic permeability coefficient than plasma. Figure 10 presents the vertical and horizontal components of the magnetophoretic force applied to the RBCs and WBCs along the centreline of the microchannel for different values of d. As the magnet distance from the microchannel increases, the magnetophoretic force decreases. It is demonstrated that the vertical component of the magnetic force applied to WBCs is higher than that exerted on RBCs. For example,  $|F_{my}|$  is 0.154 and 0.25 pN for RBCs and WBCs, respectively, for  $x = 7.51$  mm and  $d = 0.6$  mm.

In order to concentrate suspending blood cells in the main microchannel that is under the influence of magnetophoretic force (to increase the separation efficiency), two cladding streams containing only blood plasma enters through two inlets (Fig. [2](#page-3-0)). Another important factor affecting the separation of particles is the inlet flow rate. The high inlet velocity

leads to that the hydrodynamic force (the drag Stokes force) overcomes the magnetophoretic force, resulting in a reduction in the separation efficiency. On the other hand, very low velocity of the fluid containing particles causes the magnetophoretic force to overcome the drag force, leading to that the particles are trapped within the main channel and accumulated in the vicinity of the channel walls. In addition, the ratio of the cladding flow rate to the core flow rate may cause the particle to be concentrated in the centreline of the main channel. Therefore, the optimal value of this ratio is a key factor of particle separation effectively. In this paper, a wide range of different ratios of cladding flow and inlet core flow is investigated. In general, total flow rates of 7.5, 15, and 30  $\mu$ L/ h are considered. Figure [11](#page-8-0) shows an example of a complete separation of RBCs and WBCs for  $d = 0.2$  mm and a total flow rate of  $Q = 15$  µL/h. The time-dependent Eulerian approach is used to trace particles. The convergence criterion is set to  $10^{-5}$  and the time step is 1 ms. Steady fluid flow in the



Fig. 10 a, c The vertical, and b, d horizontal components of the magnetophoretic force applied to the RBCs and WBCs along the centreline of the microchannel for different values of d

<span id="page-8-0"></span>

Fig. 11 Particle trajectories and separation of WBCs (blue) and RBCs (red) for core flow rate of  $3 \mu L/h$ , cladding flow rate of 6  $\mu$ L/h and  $d = 0.2$  mm

microchannel is solved in the absence of blood cells. Figure 12 shows velocity contours throughout the microchannel for core flow rate of  $3 \mu L/h$ , cladding flow rate of 6  $\mu$ L/h and  $d = 0.2$  mm and demonstrates that the fluid flow becomes fully developed along the channel. As indicated in this figure, the parabolic velocity profile is formed at all channel cross sections. The maximum velocity remains constant with the channel length. The obtained velocity field solution is used when the particles enter the channel. Seven RBCs and seven WBCs are injected into the microchannel at each time step. Section A shows the main microchannel inlet. As can be seen, the particle focusing is generated in the vicinity of the channel centerline for exposure to the magnetic field. The highest cell separation rate occurs in sections B and C with the magnetization of 0.375 W and 0.625 W, respectively.

For cladding and core flow rates of  $6$  and  $3 \mu L$ h, respectively, and  $d = 0.2$  mm, the complete separation of RBCs and WBCs can be achieved.

Particle trajectories of blood cells in the main channel is plotted in Fig. [13](#page-9-0) for two volume flow rates of 7.5 and 15  $\mu$ l/h and  $d = 0.2$  mm. Three different values of cladding and core flow rates are considered as follows: (cladding flow rate, core flow rate, cladding flow rate) =  $(2.5, 2.5, 2.5), (3, 1.5, 3),$  and  $(3.5, 0.5, 3.5)$ for  $Q = 7.5$  µl/h and  $(5, 5, 5)$ ,  $(6, 3, 6)$ ,  $(7, 1, 7)$  for  $Q = 15$  µl/h. As the cladding flow rate increases relative to the core flow, particle motion becomes more concentrated, but the magnetophoretic force overcomes the hydrodynamic drag force causing more deviation of the cells from the microchannel walls due to the deceleration of the particles. In this case, it is possible to trap the cells in the channel. In other words, the blood cells can be directed to the outlet by increasing the distance between the magnets and the microchannel (reducing the magnetophoretic force).

Figure [14](#page-10-0) shows the particle trajectories of blood cells for volume flow rates of  $(3.5, 0.5, 3.5)$  and  $(7, 1, 7)$  $\mu$ l/h and different values of d. This figure confirms that, for  $Q = 7.5$  µl/h, the particle separation is performed better as the magnetic force decreases. It is shown that the best performance of the chip for the volume flow rate of 7.5  $\mu$ l/h is achieved when  $d = 0.8$  and 1 mm. For  $Q = 15$  µl/h, as the distance between the magnets and microchannel decreases, the particle deviation from the channel centreline increases. It can be observed that  $d = 0.2, 0.4,$  and 0.6 mm may result in more appropriate positions of magnet array. It can be concluded that the



Fig. 12 Velocity contour throughout the microchannel for core flow rate of 3  $\mu L/h$ , cladding flow rate of 6  $\mu L/h$  and  $d = 0.2$  mm

<span id="page-9-0"></span>

Fig. 13 Particle trajectories of blood cells for  $d = 0.2$  mm and: (cladding flow rate  $\mu$ I/h, core flow rate  $\mu$ I/h, cladding flow rate  $\mu$ I/h, h) = a (2.5, 2.5, 2.5), b (3, 1.5, 3), c (3.5, 0.5, 3.5), d (5, 5, 5), e (6, 3, 6), f (7, 1, 7)

volume flow rate of core and cladding channels, total flow rate, and the distance between magnets and microchannel affect the separation efficiency individually. In other words, there are optimal conditions in which the separation efficiency becomes 100% according to different magnitudes of these parameters.

Figure [15](#page-11-0) shows separation efficiency for different values of  $d$  and various volume flow rates of cladding and core channels. This figure demonstrates that the volume flow rates of (3.5, 0.5, 3.5) is more efficient for  $Q = 7.5$  µl/h, where separation efficiency is 100% for  $d = 0.8, 0.9,$  and 1 mm. For the case of  $(3, 1.5, 3)$ ,  $d = 0.9$  and 1 mm results in SE = 100%, and complete separation does not occur when the volume flow rates are (2.5, 2.5, 2.5). Different results are obtained for the case of  $Q = 7.5$  µl/h. While SE = 100% does not occur for volume flow rates of (5, 5, 5), there are several magnet array positions, i.e.,  $d = 0.1, 0.2, 0.3, 0.4$  and 0.5 mm, in which complete particle separation happens when core and cladding flow rates are (7, 1, 7).

## 6 Conclusions

In this work, a simple microfluidic device was proposed to fractionate WBCs and RBCs from whole

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Fig. 14 Particle trajectories of blood cells for (cladding flow rate  $\mu l/h$ , core flow rate  $\mu l/h$ ) cladding flow rate  $\mu l/h$ ) = a (3.5, 0.5, 3.5), **b** (7, 1, 7) and different values of  $d$ 

blood using a Halbach array of three permanent magnets. In order to concentrate the particles, two cladding flows were used. The numerical results were analyzed by changing the magnet array distance from the channel and the ratio of core and cladding streams. Total flow rates of 7.5  $\mu$ l/h and 15  $\mu$ l/h were considered to evaluate the influence of the relative flow rate of cladding streams. In addition, different distances between the Halbach array of magnets and main channel were examined. The results demonstrated that the volume flow rate of core and cladding channels, total flow rate and the distance between magnets and microchannel affect the separation efficiency individually. It was revealed that the volume flow rates of  $(3.5, 0.5, 3.5)$  is more efficient for  $Q = 7.5$  µl/h, where separation efficiency is  $100\%$  for  $d = 0.8$ , 0.9, and 1 mm. the results showed that complete particle

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Fig. 15 Separation efficiency for (cladding flow rate  $\mu l/h$ , core flow rate  $\mu l/h$ , cladding flow rate  $\mu l/h$ ) = a (2.5, 2.5, 2.5), b (3, 1.5, 3), c (3.5, 0.5, 3.5), **d** (5, 5, 5), **e** (6, 3, 6), **f** (7, 1, 7), and different values of d

separation happens when core and cladding flow rates are  $(7, 1, 7)$  and  $d = 0.1, 0.2, 0.3, 0.4$  and 0.5 mm.

### Compliance with ethical standards

Conflict of interest The authors declare that they have no conflict of interest.

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