Mathematical Modelling and Numerical Simulation of Fluid-Magnetic Particle Flow in a Small Vessel

BENCHAWAN WIWATANAPATAPHEE

Mahidol University Department of Mathematics Bangkok 10400 THAILAND scbww@mahidol.ac.th KITTISAK CHAYANTRAKOM Curtin University of Technology Department of Mathematics and Statistics Perth, Western Australia 6845 AUSTRALIA kittisak.chayantrakom@postgrad.curtin.edu.au

YONG-HONG WU Curtin University of Technology Department of Mathematics and Statistics Perth, Western Australia 6845 AUSTRALIA yhwu@maths.curtin.edu.au

Abstract: Fluid-solid flow phenomena is an interdisciplinary research area with great technological, commercial and medical importance. One particular application is related to the drug delivery system in which magnetic targeting offers the ability to target a specific site, such as a tumor. This paper presents a mathematical model and a finite element method, based on the Arbitrary Lagrangian Eulerian approach, for studying blood-magnetic particle flow in small vessels. Four models with one, three, five and nine particles are used to analyze the flow pattern and pressure distribution along the flow direction. Effects of magnetic force on the blood-particle flow are investigated.

Key-Words: Fluid-particle flow, magnetic fluids, Arbitrary Lagrangian Eulerian, finite element method.

1 Introduction

Cancer is one of the most insidious and potentially fatal diseases in human being. Many evidences indicate that progressive tumor growth is dependent on angiogenesis which is the process in which new blood vessels develop from an existing vasculature through endothelial cell sprouting, proliferation and fusion [1]. New blood vessels provide nutrients to proliferating cancer cells, which is in favor of tumor growth. Tumor cells need an adequate blood supply in order to perform vital cellular functions. The degree of disturbance of blood flow is thus a good predictor of the course of the disease, and hence regional blood flow measure can permit earlier cancer detection. The modern-day approach to cancer treatment is a multidisciplinary one involving varying combination of surgery, radiation therapy, chemotherapy and targeted therapies (a new weapon). In targeted therapies, a medication or drug is controlled to target a specific pathway in the growth and development of a tumor. Although most of the drugs used to date have proven to be successful on small animals such as mice [8, 9], their efficiency in humans remains highly variable from one patient to another. Understanding the flow

of blood and drug in the capillary bed is very important for investigating the efficiency of drug treatment as they pass from parent blood vessel to tumor surface via an associated capillary bed. Over the last 15 years, a number of mathematical models for blood vessel formation [2, 3], blood flow and/or particle flow in capillary networks [4, 5, 6, 7] in the area of tumorinduced angiogenesis have been developed. One of these, magnetically targeted drug delivery, involves binding a drug to small biocompatible magnetic particles with diameters less than $5\mu m$.

Driscoll *et al.* [11] had studied magnetically targeted drug delivery by tracking each individual particle under the influence of Stokes drag force and magnetic force. Grief and Richardson [7] conducted a theoretical analysis of targeted drug delivery using magnetic particles and proposed a two-dimensional network model. In their model, the motion of fluid is described by Poiseuille flow, while the motion of a magnetic particle, due to balancing hydrodynamic and magnetic force, is governed by an advection-diffusion equation for the particle concentration. They found that drug targeting can be achieved by pulling magnetic particles to the edge of the vessel, and that the use of magnetically targeted drug delivery with an externally applied magnetic field is appropriate only for targets close to the surface of the body.

However, most existing models do not take into account of the real 3D effect and the interaction between blood flow and magnetic particles, Non-Newtonian behavior of blood and the effect of magnetic forces. Therefore in order to fully understand and control the local flow behavior of blood and particles through the area of tumor-induced angiogenesis, it is essential to develop a sophisticated model and computational technique for the flow analysis. Hence, based on the current development in the field, the objective of this paper is to propose a sophisticated model and computational technique for analyzing the complex flow (blood-particle flow) behavior in tumorinduced capillary networks using the current state-ofthe-art computational fluid dynamic (CFD) technology. Our model couple the interaction of blood flow with the particle flow using the Arbitrary Lagrangian Eulerian (ALE) formulation. The governing equations for blood flow are the continuity equation and the Navier-Stokes equations. For particle movement, Newton's law is applied. The complete model includes the governing equations for the blood flow, the governing equations for the motion of fine particles, the interaction conditions between blood and particle at the interfaces, and boundary conditions.

2 Governing Equations

To study the motion of solid particles immersed in a fluid, we assume that the fluid-solid particle system occupies a bounded domain $\overline{\Omega}$ in \mathbf{R}^3 . At a typical instant of time t, Q particles occupy Q closed connected subsets $\sum_{q=1}^{Q} \Omega_q \subset \mathbf{R}^3$ which is surrounded by a viscous homogeneous fluid filling the domain $\overline{\Omega} - \sum_{q=1}^{Q} \Omega_q$ called the flow-channel area.

In this study we use two coordinate systems: a reference system, Ω , where the model is drawn and the particle movement is solved, and a moving mesh system, Ω_{def} , corresponding to the deformed mesh of the flow channel, where we simulate the fluid flow. The time evolution of the domain Ω_{def} is determined by means of an Arbitrary Lagrangian-Eulerian (ALE) mapping $\mathbf{x} : \Omega \times \mathbf{R}^+ \mapsto \Omega_{def}$ which maps any point (\mathbf{X}, t) to its image $\mathbf{x}(\mathbf{X}, t)$.

2.1 Transformation

In the flow-channel area, the two coordinate systems, $(X, Y, Z) \in \Omega$ and $(x, y, z) \in \Omega_{def}$, are connected through a transformation T. At the initial state at t = 0, the two mesh systems are assumed to coincide. The

transformation T maps the point initially located at (X, Y, Z) to the point (x, y, z) at time t:

$$\begin{array}{rcl} x &= x(X,Y,Z,t) \\ T &: & y &= y(X,Y,Z,t) \\ & z &= z(X,Y,Z,t). \end{array}$$

Suppose that the functions x, y and z are continuous differentiable with respect to X, Y, Z. Then the infinitesimals dX, dY, dZ transform into dx, dy, dz according to

$$dx = x_{,X}dX + x_{,Y}dY + x_{,Z}dZ,$$

$$dy = y_{,X}dX + y_{,Y}dY + y_{,Z}dZ,$$

$$dz = z_{,X}dX + z_{,Y}dY + z_{,Z}dZ,$$

(1)

where $(\cdot)_{,X}$ denotes differentiation with respect to *X*. System (1) can be written in matrix form as

$$\begin{bmatrix} dx \\ dy \\ dz \end{bmatrix} = \begin{bmatrix} x_{,X} & x_{,Y} & x_{,Z} \\ y_{,X} & y_{,Y} & y_{,Z} \\ z_{,X} & z_{,Y} & z_{,Z} \end{bmatrix} \begin{bmatrix} dX \\ dY \\ dZ \end{bmatrix}.$$
 (2)

The 3×3 matrix of partial derivatives in (2) is called the Jacobian matrix of the transformation. Denote the matrix by **J**, then

$$\begin{aligned} |\mathbf{J}| &= x_{,X}(y_{,Y}z_{,Z} - y_{,Z}z_{,Y}) - x_{,Y}(y_{,X}z_{,Z} - y_{,Z}z_{,X}) \\ &+ x_{,Z}(y_{,X}z_{,Y} - y_{,Y}z_{,X}). \end{aligned}$$

For $|\mathbf{J}| \neq 0$, the transformation is invertible and there exists an inverse transformation at time *t*, i.e.,

$$\begin{array}{rcl} X &= X(x,y,z) \\ T^{-1} &: & Y &= Y(x,y,z) \\ & & Z &= Z(x,y,z). \end{array}$$

As in (2), we have

$$\begin{bmatrix} dX \\ dY \\ dZ \end{bmatrix} = \begin{bmatrix} X_{,x} & X_{,y} & X_{,z} \\ Y_{,x} & Y_{,y} & Y_{,z} \\ Z_{,x} & Z_{,y} & Z_{,z} \end{bmatrix} \begin{bmatrix} dx \\ dy \\ dz \end{bmatrix}.$$
 (3)

From (2), we also have

$$\begin{bmatrix} dX \\ dY \\ dZ \end{bmatrix} = \mathbf{J}^{-1} \begin{bmatrix} dx \\ dy \\ dz \end{bmatrix}, \qquad (4)$$

where

$$\mathbf{J}^{-1} = \begin{bmatrix} I_{Xx} & I_{Xy} & I_{Xz} \\ I_{Yx} & I_{Yy} & I_{Yz} \\ I_{Zx} & I_{Zy} & I_{Zz} \end{bmatrix} = \frac{1}{|\mathbf{J}|} \mathbf{M}, \quad (5)$$

$$\mathbf{M} = \begin{bmatrix} y_{,Y}z_{,Z} - y_{,Z}z_{,Y} & x_{,Z}z_{,Y} - x_{,Y}z_{,Z} & x_{,Y}y_{,Z} - x_{,Z}y_{,Y} \\ y_{,Z}z_{,X} - y_{,X}z_{,Z} & x_{,X}z_{,Z} - x_{,Z}z_{,X} & y_{,X}x_{,Z} - y_{,Z}x_{,X} \\ y_{,X}z_{,Y} - y_{,Y}z_{,X} & x_{,Y}z_{,X} - x_{,X}z_{,Y} & x_{,X}y_{,Y} - x_{,Y}y_{,X} \end{bmatrix}$$

in which the I syntax is used to emphasize the computation of the Jacobian of the inverse transformation from the inverse of the original Jacobian.

2.2 Motion of Fluid-Solid Flow in the Deformed Mesh System

To study the motion of magnetic particles in the fluid flow channel, we assume that the gravitational force can be neglected and the particle movement is governed by Newton's second law:

$$m_q \frac{\partial \mathbf{V}_q}{\partial t} = \mathbf{F}_v + \mathbf{F}_q + \mathbf{F}_{mag}, \quad q = 1, 2, 3..., Q$$
$$\mathbf{V}_q|_{t=0} = 0.$$
(6)

The position \mathbf{X}_q of the center of the *q*th particle can be determined by the equation:

$$\frac{d\mathbf{X}_q}{dt} = \mathbf{V}_q, \quad q = 1, 2, 3..., Q
\mathbf{X}_q|_{t=0} = \mathbf{X}_q^0.$$
(7)

In equation (6)₁, \mathbf{V}_q and m_q denote the velocity vector and the mass of the *q*th particle. The three applied loads, drag force \mathbf{F}_v , collision force \mathbf{F}_q and magnetic force \mathbf{F}_{mag} , are defined based on the following assumptions:

All boundaries of particles experience drag force
 F_v from fluid,

$$\mathbf{F}_{v} = -\mathbf{n}_{f} \cdot \left(-p \mathbf{I} + \eta (\nabla \mathbf{v} + (\nabla \mathbf{v})^{T})\right) \quad (8)$$

which consists of the pressure and the viscous drag of the fluid.

• To prevent the collisions among the particles, and the particles and the vessel walls, the particleparticle interaction force $\mathbf{F}_{q,p}$ and the particlewall interaction force $\mathbf{F}_{q,w}$ are applied when the distance between two particles, or between a particle and a wall, is within the order of the element size [10]

$$\mathbf{F}_{q} = \sum_{p=1, p \neq q}^{Q} \mathbf{F}_{q,p} + \sum_{w=1}^{2} \mathbf{F}_{q,w}, \qquad (9)$$

in which

$$\mathbf{F}_{q,p} = \begin{cases} 0, \text{ for } d_{q,p} > R_q + R_p + \alpha \\ \frac{1}{\varepsilon_q} (\mathbf{X}_q - \mathbf{X}_p) (R_q + R_p + \alpha - d_{q,p})^2, \\ \text{ for } d_{q,p} \le R_q + R_p + \alpha \end{cases}$$
(10)

and

$$\mathbf{F}_{q,w} = \begin{cases} 0, \text{ for } d_{q,w} > 2R_q + \alpha \\ \frac{1}{\varepsilon_w} (\mathbf{X}_q - \mathbf{X}_w) (2R_q + \alpha - d_{q,w})^2, \\ \text{ for } d_{q,w} \le 2R_q + \alpha \end{cases}$$
(11)

where $d_{q,p}$ denotes the distance between the centers of the *q*th and *p*th particles, $d_{q,w}$ denotes the distance between the centers of the *q*th particle and the imaginary particle on the other side of the wall, \mathbf{X}_q and R_q are center and radius of the *q*th particle, α is the force range, and ε_q and ε_w are small positive stiffness parameters.

• To trap magnetic particles (drugs) at the target site, an external magnetic field is applied to generate a magnetic force acting on the particle [12]. This force, which is composed of three components, is governed by the equations:

$$\mathbf{F}_{mag} = \frac{1}{\mu_r} (\mathbf{M} \cdot \nabla) \mathbf{B}, \qquad (12)$$

where μ_r is the relative permeability of a magnetic material, $\mathbf{M} = (M_x, M_y, M_z)$ is the magnetic moment of the particle and $\mathbf{B} = (B_x, B_y, B_z)$ is the magnetic flux density.

To determine the drag force \mathbf{F}_v in (8), blood is assumed to be an isotropic, homogeneous incompressible fluid. The motion of the blood is described by the continuity equation and the Navier-Stokes equations

$$\nabla \cdot \mathbf{v} = 0, \tag{13}$$

$$\rho_f \frac{\partial \mathbf{v}}{\partial t} + \rho_f (\mathbf{v} \cdot \nabla) \mathbf{v} - \nabla \cdot \sigma = \mathbf{F}, \qquad (14)$$

for \mathbf{x} in $\Omega_{def}(t)$ where ρ_f denotes the blood density, $\mathbf{v} = [u, v, w]^T$ represents the 3D velocity vector, and \mathbf{F} is the volume force acting on the fluid. For this model, we neglect the effect of gravitational force and thus $\mathbf{F} = \mathbf{0}$. The quantity σ in equation (14) is the stress tensor given by

$$\sigma = -p\mathbf{I} + \eta(\nabla \mathbf{v} + (\nabla \mathbf{v})^T), \quad (15)$$

where η is the blood viscosity and p is the blood pressure.

On the wall, the no-slip condition is applied. On the inflow boundary Γ_{in} , the velocity is assumed to be constant, while on the outflow boundary Γ_{out} , the stress-free condition is used:

$$\mathbf{v} = \mathbf{v}_0 \quad \text{on } \Gamma_{in} \\ \sigma \cdot \mathbf{n} = 0 \quad \text{on } \Gamma_{out}.$$
 (16)

For static condition in stationary bodies, the magnetic flux density \mathbf{B} is governed by Maxwell's equations:

$$\nabla \cdot \mathbf{B} = 0$$

$$\nabla \times \mathbf{H} = \mathbf{0}$$
(17)

where the magnetic flux density \mathbf{B} and the magnetic field strength \mathbf{H} are related through the constitutive relation

$$\mathbf{B} = \mu_0 \mu_r \mathbf{H} + \mathbf{B}_r, \tag{18}$$

in which $\mathbf{B}_r = \mu_0 \mu_r \mathbf{M}$ denotes a residual flux density, μ_0 is the permeability in vacuum.

From the first equation of (17), the magnetic flux density can be determined from a vector potential by $\mathbf{B} = \nabla \times \mathbf{A}$ which identically satisfies the first equation of (17). Using the identity

$$\nabla \times (\nabla \times \mathbf{A}) = \nabla (\nabla \cdot \mathbf{A}) - \triangle \mathbf{A},$$

and the Coulomb gauge $\nabla \cdot \mathbf{A} = 0$, the second equation of (17) takes the form

$$\nabla \times (\mu_0^{-1} \mu_r^{-1} \nabla \times \mathbf{A} - \mathbf{M}) = \mathbf{0}, \forall \mathbf{x} \in \Omega_{def}$$

or
$$\triangle \mathbf{A} = -\nabla \times (\mu_0 \mu_r \mathbf{M}),$$
(19)

which is the vector-valued Poisson equation for the magnetic potential **A**.

Due to the movement of the coordinate system, the mesh velocity $\Psi = (\Psi_x, \Psi_y, \Psi_z)$ is introduced in the deformed domain Ω_{def} . To guarantee a smoothly varying distribution of the nodes, we assume that the nodes on $\partial \Omega_q$ move with the particle (no slip) and that each component of the mesh velocity in the fluid channel is governed by a Laplace equation:

$$\nabla^2 \Psi = \mathbf{0}, \quad \forall \mathbf{x} \in \Omega_{def}. \tag{20}$$

The above equation is to smooth gradient of the mesh velocity over the domain so as to reduce mesh distortion. Once the mesh velocity components are determined, we can determine the smoothed deformed mesh for the flow channel at each time instant by updating the coordinates of the nodes according to the following formulae

$$\begin{aligned}
x &= X + \int_{0}^{t} \Psi_{x} dt, \\
y &= Y + \int_{0}^{t} \Psi_{y} dt, \\
z &= Z + \int_{0}^{t} \Psi_{z} dt.
\end{aligned}$$
(21)

Another condition that needs to be specified is that the fluid, particle and mesh move with the same velocity on the particle boundaries, i.e.,

$$\Psi = \mathbf{v} = \mathbf{V}_q \quad \text{on } \partial \Omega_q. \tag{22}$$

We now have the strong coupled problem for the fluid-particle flow in the drug delivery system. These equations are solved to yield \mathbf{V}_q in Ω and $\mathbf{v}, p, \mathbf{A}, \Psi$, in Ω_{def} .

3 Finite Element Formulations

The weak formulation of the fluid flow problem is to find $(\mathbf{v}, p, \mathbf{A}, \Psi) \in \mathfrak{T} \equiv [\mathrm{H}^1(\Omega_{def})]^3 \times \mathrm{H}^1(\Omega_{def}) \times [\mathrm{H}^1(\Omega_{def})]^3 \times [\mathrm{H}^1(\Omega_{def})]^3$ in the deformed mesh system at each time instant such that all the Dirichlet boundary conditions are satisfied and $\forall (\hat{\mathbf{v}}, \hat{p}, \hat{\mathbf{A}}, \hat{\Psi}) \in \mathfrak{T}_0 \equiv \{(\hat{\mathbf{v}}, \hat{p}, \hat{\mathbf{A}}, \hat{\Psi}) \in \mathfrak{T} \mid \hat{\mathbf{v}} = \mathbf{0} \text{ on } \partial\Omega_{def_{\mathbf{v}}}, \hat{p} = 0 \text{ on } \partial\Omega_{def_p}, \hat{\mathbf{A}} = \mathbf{0} \text{ on } \partial\Omega_{def_{\mathbf{A}}} \text{ and } \hat{\Psi} = \mathbf{0} \text{ on } \partial\Omega_{def_{\mathbf{Y}}} \},$

$$\int_{\Omega_{def}} \hat{p} \left(\nabla \cdot \mathbf{v} \right) d\Omega = 0, \qquad (23)$$

$$\int_{\Omega_{def}} \left(\rho_f \hat{\mathbf{v}} \cdot \frac{\partial}{\partial t} \mathbf{v} + \eta \nabla \hat{\mathbf{v}} : \nabla \mathbf{v} + \rho_f \hat{\mathbf{v}} \cdot (\mathbf{v} \cdot \nabla) \mathbf{v} \right)$$

$$-p\nabla\cdot\hat{\mathbf{v}}\right)d\Omega = \int_{\partial\Omega_{def}}\hat{\mathbf{v}}\cdot(\boldsymbol{\sigma}\cdot\mathbf{n})\,ds \qquad (24)$$

$$\int_{\Omega_{def}} \left(\nabla \hat{\mathbf{A}} : \nabla \mathbf{A} - \hat{\mathbf{A}} \cdot \nabla \times (\mu_0 \mu_r \mathbf{M}) \right) d\Omega = 0,$$
(25)

and

$$\int_{\Omega_{def}} \nabla \hat{\Psi} : \nabla \Psi \ d\Omega = 0, \qquad (26)$$

where $\partial\Omega_{def_{\Psi}}$, $\partial\Omega_{def_{P}}$, $\partial\Omega_{def_{A}}$ and $\partial\Omega_{def_{\Psi}}$ are the parts of boundary where the velocity, the pressure, the magnetic potential and the mesh velocity are specified. It should also be addressed that various surface integral terms, arising in the formulation, vanish as the test functions involved in the terms are zero on the boundary.

Since the computations are conducted in the reference coordinates, Ω , we need to transform equations (23) - (26) in the deformed coordinates to those equations in the reference coordinates. Through this and using (16)₂, we obtain

$$\int_{\Omega} \hat{p} \left(\nabla \cdot \mathbf{v} \right) \left| \mathbf{J} \right| \, d\Omega = 0, \qquad (27)$$

$$\int_{\Omega} \left(\rho_f \hat{\mathbf{v}} \cdot \frac{\partial}{\partial t} \mathbf{v} + \eta \nabla \hat{\mathbf{v}} : \nabla \mathbf{v} + \rho_f \hat{\mathbf{v}} \cdot (\mathbf{v} \cdot \nabla) \mathbf{v} - p \nabla \cdot \hat{\mathbf{v}} \right) |\mathbf{J}| \, d\Omega = 0, \quad (28)$$

$$\int_{\Omega} \left(\nabla \hat{\mathbf{A}} : \nabla \mathbf{A} - \hat{\mathbf{A}} \cdot \nabla \times (\mu_0 \mu_r \mathbf{M}) \right) |\mathbf{J}| \, d\Omega = 0,$$
(29)

and

$$\int_{\Omega} \nabla \hat{\Psi} : \nabla \Psi |\mathbf{J}| \, d\Omega = 0, \qquad (30)$$

where the derivatives of the unknown functions Ψ_i (i = x, y, z) are determined by the following expressions:

$$\Psi_{i,x} = \Psi_{i,X}I_{X,x} + \Psi_{i,Y}I_{Yx} + \Psi_{i,Z}I_{Z,x},
\Psi_{i,y} = \Psi_{i,X}I_{X,y} + \Psi_{i,Y}I_{Yy} + \Psi_{i,Z}I_{Z,y},
\Psi_{i,z} = \Psi_{i,X}I_{X,z} + \Psi_{i,Y}I_{Yz} + \Psi_{i,Z}I_{Z,z}.$$
(31)

The derivatives of other unknown functions u, v, w, A_x, A_y and A_z are defined in the same way as those of the Ψ_i functions.

4 Numerical Results and Discussion

In a two dimension case, the magnetic potential is assumed to have a nonzero component only in the direction perpendicular to the plane, i.e., $\mathbf{A} = (0, 0, A_z)$. On $\partial \Omega_q$ and $\partial \Omega$, the magnetic potential is set to zero, that is, $A_z = 0$.

The magnetization $\mathbf{M} = (M_x, M_y)$ for the magnetic source is given by $M_x = 0, M_y = 5 \times 10^4 A \cdot m^{-1}$, and for the magnetic particles

$$M_x = a \arctan\left(\frac{b}{\mu_0 \mu_r} A_{z,y}\right),$$

$$M_y = a \arctan\left(-\frac{b}{\mu_0 \mu_r} A_{z,x}\right),$$
(32)

where a and b are two material parameters.

To understand the blood-particle flow in a small vessel, a 2D domain with one, three, five and nine particles are used. The computation domain is a horizontal channel with height of 6.2 μm and length of 45 μm . The particles are circular with diameter of 0.5 μm . Blood is assumed to flow into the channel with speed 1.85 cm/s from the left to the right. The fluid properties are typical of human blood with the viscosity η of 0.0035 $Pa \cdot s$ and the density ρ_f of 1060 $kg \cdot m^{-3}$. All particles are assumed to be solid with the density of $1112 kg \cdot m^{-3}$. The relative permeability μ_r is 5×10^3 for the magnet particles and 0.99998 for the tissue in the blood vessel. The material parameters a and b are 1×10^{-4} and 3×10^{-5} , respectively.

The Arbitrary Lagrangian Eulerian approach is used to handle the dynamics of deforming geometry and the moving boundaries. New mesh coordinates on the channel area are calculated based on the movement of the particles. The Navier-Stokes equations are formulated in the moving coordinate system. Particle interactions and particle collisions are neglected. Via the simulations of the model we can describe the flow pattern and pressure distribution in the particle-fluid system.



Figure 1: The 2D geometry of the blood vessel with a magnetic source at the middle of the vessel and its finite element mesh



Figure 2: Velocity profiles at various instants of time for the case with three particles



Figure 3: Velocity profiles at various instants of time for the case with five particles

Figure 1 shows the finite element mesh and the external magnetic field applied to the system. The computation domain consists of 3519 elements with 1791 nodes.

Figures 2-3 show the velocity distributions for the models with three and five particles, respectively, at the absence of the external magnetic field. In these cases, the particles flow in the axial direction. Figure 4 shows the pressure distribution along the axis of the tube at t = 0 for various cases with different number of particles in the fluids. It is noted that with the increase of particle number, the pressure required on the entry of the tube increases significantly. Figure 5 shows the velocity profile of fluid and the particle motion for the case with four particles at the presence of an external magnetic field. It is clearly noted that the



Figure 4: Pressure profiles along the flow direction at t = 0s



(c) t = 9.96 micro sec.

Figure 5: Velocity profiles at various instants of time for the case with four magnetic particles

model can simulate the flow of particles toward the targeted region.

Acknowledgements: The first author gratefully acknowledges the support of the Thailand Research Fund (TRF) and the support of the Australian government under the 2007 Endeavor Australian Research fellowship. The research is done in collaboration with Mahidol University, Bangkok, Thailand and Curtin University of Technology, Perth, WA.

References:

- Risau W, Mechanisms of angiogenesis, *Nature* 386:671–74, 1997.
- [2] Byrne HM, Chaplain MAJ, Mathematical models for tumour angiogenesis: numerical simulations and nonlinear wave solutions, *Bull.Math.Biol.* 57:461–86, 1995.
- [3] Levine HA, Pamuk S, Sleeman BD, Nielsen-Hamilton M, Mathematical modeling of capillary formation and development in tumour angiogenesis: penetration into the stroma, *Bull. Math. Biol.* 63:801-63, 2001.
- [4] Stephanou A. McDougall SR, Anderson ARA, Chaplain Maj, Mathematical modeling of flow in 2D and 3D vascular networks: applications to antiangiogenic and chemotherapeutic drug strategies, *Math. Comput. Model.* **41**:1137–56, 2005.
- [5] Stephanou A, McDougall SR, Anderson ARA, Chaplain MAJ, Mathematical modeling of the influence of blood rheological properties upon adaptive tumourinduced angiogenesis, *Math. Compt. Model.* In press.
- [6] Ruuge EK, Rusetski AN, Magnetic fluids as drug carriers: targeted transport of drugs by a magnetic field, *Journal of Magnetism and Magnetic Materials* 122:335–339, 1993.
- [7] Grief AD and Richardson G, Mathematical modelling of magnetically targeted drug delivery, *Journal of Magnetism and Magnetic Materials* 293(1):455–463, 2005.
- [8] Goodwin S, Peterson C, Hoh C, Bittner C, Targeting and retention of magnetic targeted carriers (MTCs) enhancing intra-arterial chemotherapy, *J. of Magnetism and Magnetic Materials* **194**:132–139, 1999.
- [9] Widder KJ, Morris RM, Poore G, Howard DP Jr and Senyei AE, Tumor remission in Yoshida sarcoma-bearing rates by selective targeting of magnetic albumin microspheres containing doxorubicin, *Pro.Natl.Acad.Sci. USA* **78**(1):579–581, 1981.
- [10] Singh P, Joseph DD, Hesla TI, Glowinski R, Pan T-W, A distributed Lagrange multiplier/fictitious domain method for viscoelastic particulate flows, *J. Non-Newtonian Fluid Mechanic* **91**:165–188, 2000.
- [11] Driscoll CF, Morris RM, Senyei AE, Widder KJ and Heller GS, Magnetic targeting of microspheres in blood flow, *Microvasc. Res.* **27**:353–369, 1984.
- [12] Felix AH and Sylvain M, Suggested Shape for a First Generation Endovascular Untethered Microdevice Prototype, Proceedings of the 2005 IEEE Engineering in Medicine and Biology 27th Annual Conference Shanghai, China, September 1-4, 2005, p.1286–1288.