# **DCE-CT Perfusion Parametric Coloured Image Using Steepest Gradient Approach Based on Time Frame**

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## Abstract

Technological advances in medical imaging modality, counting higher special and temporal resolution, have been utilized to aid radiologist in diagnosing, prognostic and, hence, management of brain tumour. After tracer injection, CT scanning along with pharmacokinetic models are applied to measure several hemodynamic parameters used to describe microvasculature within a targeted region of interest (ROI). These parameters are a choice for diagnostic and prognostic and the likelihood of preferable treatment for brain tumour. The perfusion pixel-wise perfusion map image is a fingerprint of abnormality of ROI within brains that aid radiologist to separate between necrotic and neoplastic of tumour territory and to avoid arterial partial-volume averaging. This work aims to generate pixel-wise perfusion map colour image of brain by using the steepest gradient approach based on the time frame of tracer passage.

Keywords: perfusion image, DCE-CT, ROI, time frame, steepest gradient approach.

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# 1. Introduction

Angiogenesis is the procedure of proliferation of tumour cells, at which new vessels are formed to feed nutrients and oxygen for tumour growth where blood-brain barrier (BBB) is disrupted. On the other hand, angiogenesis plays an important role in forming and regulating the microvasculature and hence greatly influences tumour treatment [22] [3] [18] [4] [23].

Computed Tomography Dynamic Contrast-Enhanced (CT-DCE) technique has been used to measure certain parameters that describe the microvasculature [11] [14] [6]. Among these parameters is the blood perfusion which is measured by using the gradient approach.

Follow the injection of the tracer, normally iodinated contract agent, cine images of the area of interesting is generated to monitor the behaviour of the tracer first pass. By utilizing these cine images, hence, maps of perfusion parameters are generated, interpreted, and thereby aid in deciding whether a patient needs immediate thrombolytic treatment. This interpretation is critical in tailoring treatment to each patient, and thus saving lives and reducing the possibility of severe disability.

The parametric maps are distant with regards to a certain diagnostic accuracy and further refinement of the techniques and methods in use are desired [24] [17] [19].

Many well-established tracer kinetic models are available that can provide useful estimates of microvasculature functional parameters. These models can be roughly classified into two streams [15]: the compartmental models and distributed parameter models developed by Lawrence and Lee [20]. Unlike the compartmental models, the distributed parameters models define the contrast agent within the vascular space as a function of time and distance along the length of a capillary.

Applying these models requires specific knowledge including the vascular and compartments, the volumes of the compartments constituting the model and the tracer permeability through the semi-membrane compartment barriers. Complexity, of the above models, leads researchers to use simpler and more straightforward model, such as gradient model [11] [21] [2].



The gradient model states that the maximum slope of the parenchyma curve has to be reached before venous outflow starts. The most advantage of this model is that there is no need to use the gamma variate fitting to smooth the AIF data [10].

Further work was done by Ashoor [1] to include the time frame of the bolus passage. In this work, based on pixel-wise analysis, colour map of cerebral blood flow based on Ashoor approach is established.

#### 2. Theoretical Background

The Fick's principle, which is based on the basic law of conservation of mass, describes the detection of the transit of tracer bolus through a region as:

$$\frac{\partial m}{\partial t} = FC_a(t) - FC_v(t) - m_t \tag{1}$$

Where m is the amount of the tracer, F is the flow, Ca(t) is the feeding artery concentration, Cv(t) is the vein concentration and mt is the amount of tracer within the tissue of interest.

Assuming that the complete system is at systemic tracer steady state, the above equation can be expressed as:

$$\frac{\partial m}{\partial t} = 0 = FC_a(t) - FC_v(t) - m_t$$
or
$$F = \frac{m_t}{C_a(t) - C_v(t)}$$
(2)

The above equation can be modified to make its application similar to the microsphere assumption, i.e. no outflow of the tracer from the tissue of interest (Cv(t)=0). The equation can be simplified as:

$$F = \frac{m_t}{C_a(t)}$$
or
$$F = \frac{C_t(t)V_t}{C_a(t)}; \quad m_t = C_t(t)V_t$$
(3)

Where Ct(t) is the tracer concentration that occupied Vt, volume of the tissue of interest.

Hence, the perfusion within the tissue of interest can be written as:

$$\frac{F}{V_t} = perfusion = \frac{C_t(t)}{C_a(t)}$$
(4)

By using the stimulus-response theorem in which the tissue concentration can be written as:

$$C_t(t) = \int_{t_0}^{T_c} C_a(t) dt$$

(5)

Tc is the mean transit time of the tissue of interest, equation (4) can be written as:

$$\frac{F}{V_t} = perfusion = \frac{\int_{t_0}^{T_c} C_a(t)dt}{C_a(t)}$$
(6)

The optimum value of the ratio of the right-hand side occurs at  $t \leq Tc$  at which the steepest gradient must be determined before washout of the tracer takes place from the tissue of interest.

Since the steepest gradient is located at the maximum of the feeding artery curve, usually sagittal sinus, the equation can be expressed in a simple form as:

$$\frac{F}{V_{t}} = perfusion = \frac{max \ slope \int_{t_{0}}^{T_{c}} C_{a}(t) dt}{peak \ C_{a}(t)}$$
or
$$\frac{F}{V_{t}} = perfusion = \frac{max \ slope \ C_{t}(t)}{peak \ C_{a}(t)}$$
(7)



The advantage of using the proposed method by Ashoor [2] is that the initial stage data of the tracer passage is enough to generate colour map image of cerebral blood flow (CBF). For instance, utilising the data from the starting of the enhancement of CT scanner signal (starting point of the tracer's wash in) to the peak of AIF (starting point of the tracer's washout), where the tracer is confined, almost totally, within the voxel of interest (Fig. 1).



Fig.1. The voxel of interest



Fig.2. A sample of DCE-CT series images

Figure 1 showing a sample of DCE-CT series images acquired from the patient brain.

The perfusion map image is constructed by estimating the ratio between the maximum slope of the uptake curve and the peak of arterial input function [16] projecting individual pixel using equation (7).

### 3. Results

Based on the model developed in the previous section (steepest gradient method based on the time frame proposed by Ashoor [1], the perfusion image of blood flow is constructed (Fig.2). The active region of the tumour, as highlighted in the left image, represents the highest blood flow.

This area surrounds a dead area of less blood flow, the necrotic area. In the blood flow map image developed using the gradient method, the right one, indicates higher blood flow within the active area of the tumour (red). Clearly, it shows the enactive area as low blood flow (blue).





Fig.3. Compares the perfusion image HU

Figure 3 compares the perfusion image HU (right) generated using the TTP time frame steepest gradient approach based on the tracer passage time frame with CT grey image (left) HU.

#### 4. Discussion

Nowadays dynamic enhanced contrast CT and computeraided decision CAD become valuable tools in the clinical environment aiding decision-makers (DM) in quantitative and diagnoses of abnormalities within the human brain. Consequently, this enhances favourable treatment, imageguided intervention, and surgical planning. Furthermore, the perfusion technique has been widely spread in the field of oncology as it provides valuable details about tumour grading and angiogenesis "in vivo [13]. Due to the heterogeneity of tumour cells, manual identification of the ROIs is not applicable since essential details about tumour morphology may be lost.

Hence, several attempts have been made to utilise the kinetic analysis of individual pixel data to generate a distribution map based on measured parameters that describe the microvasculature such as CBF [4] [12] [2] [7]. Measurement of CBF within the ROI, i.e. tumour, provides valuable details about the separation between necrotic area, low flow, and neoplastic area, high flow [5].

There exist several models used to establish blood flow coloured map image, however, the steepest gradient method is simple, least error and straight forward [11] [14] [9]. To generate the coloured map image of CBF by using the steepest gradient approach, tissue density curve (TDC) and arterial input function (AIF) should be determined. AIF is the feeding artery of the tissue of interest that is programmatic to identify because of the complexity of the microvasculature [9]. Hence, to avoid arterial partial-volume averaging, superior sagittal sinus [10] is used to establish AIF.

Furthermore, the steepest gradient approach based on the initial time frame of the tracer passage, proposed by Ashoor [2], is utilized to generate pixel-wise profusion coloured map image. After injection of iodinated tracer, intravenously, cine images (Figure 2) of targeted slices of brain are generated to monitor the behaviour of the tracer. Among the existing imaging modalities, CT is favourable, where the signal intensity is linearly proportional to the amount of the tracer within the ROI.

The main goal of image remapping and segmentation are to enhance images by dividing them into meaningful, homogeneous, and nonoverlapping or texture. The outcome of segmentation either an image of labels identifying each homogeneous region or a set of counters which describe the region boundaries. In this work, the map image is divided into colour to specify the more blood flow that can be identified by red colour (Figure 3). This assists decision-makers in the clinical environment to issue improved diagnoses, and hence, contributes overall treatments [8].

#### 5. Conclusion

This study set out to assess the applicability of the steepest gradient based on the time frame by utilising CT dynamic imaging to generate the blood flow colour map. The most obvious finding to emerge from this study is that the steepest gradient based on time frame is practical less error approach allows to estimate the CBF. Furthermore, it is enough to build pixel-wise colour CBF map by using partial time course of tracer's passage that enabling to distinguish between necrotic area, low blood flow, and neoplastic area, high blood flow.

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