ASSESSMENT OF ACUTE SPONTANEOUS INTRACEREBRAL HEMATOMA BY CT PERFUSION IMAGING

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SUMMARY

A single-section deconvolution-derived computerized tomographic perfusion imaging was performed in 45 patients (22 male and 23 female; mean age = 69.89 ± 10.07 years) with acute supratentorial spontaneous intracerebral hemorrhage. Mean rCBF and rCBV were lower in the hemorrhagic core than in the perihematomal low density area (p<0.001), and in the perihematomal low density area than in normal appearing brain parenchyma (p<0.001). Mean rMTT values were higher in perihematomal low density area than in normal appearing area (p<0.001) and in both hemorrhagic and perihematomal area than in controlateral ROI (p<0.001). There were no differences in rMTT mean values between hemorrhagic core and perihematomal area, as well as between normal appearing and controlateral areas. We found a concentric distribution of all CT perfusion parameters characterized by an improvement from the core to the periphery, with low perihematomal rCBF and rCBV values suggesting edema formation.

Key words: CT-perfusion, intracerebral hematoma.

RÉSUMÉ

Imagerie de perfusion d’hématome intracérébral spontané évaluée par tomodensitométrie hémodynamique

Une coupe unique de scanner de perfusion avec déconvolution était réalisée chez 45 patients (22 hommes et 23 femmes; âge moyenne = 69,89 ± 10,07 years) présentant un hématome intracérébral aigué sustentoriel (HIA S). Le débit sanguin cérébral (DSC) et volume sanguin cérébral (VSC) moyens est plus bas au centre qu’en périphérie de la zone d’hypodensité qui est plus bas qu’en zone isodense (p < 0,001). Les valeurs de temps de transit moyen (TTM) étaient > zones périlésionnelle > zone isodense (p < 0,001). Il n’y a pas eu de différence en TTM entre le centre et la zone périhémorragique. Nous avons trouvé une distribution centrifuge caractérisée par une amélioration des paramètres de perfusion du centre à la périphérie avec les valeurs basses de DSC VSC suggérant une formation d’édème.

Mots-clés : scanner de perfusion, hématome intracérébral.

INTRODUCTION

A better understanding of the pathophysiological mechanisms underlying secondary neuronal damage that can affect the perihematomal tissue early after spontaneous intracerebral hemorrhage (SICH) is crucial for the choice of appropriate strategies of therapy [19].

Several efforts have previously been made to verify whether in the region peripheral to hematoma there exists a zone of ischemic penumbra, defined as severely hypoperfused tissue at risk for infarction which is functionally compromised but structurally intact and, therefore, still viable and salvageable if blood flow is rapidly restored [6]. However, the presence of ischemic penumbra tissue surrounding the hematoma remains controversial in both animal [1, 18] and human studies [7, 12, 15, 20, 21, 27]. Among the different techniques currently available to investigate perfusion abnormalities, dynamic computed tomography (CT) perfusion scanning has recently proven to be a promising tool for the detection of cerebral blood flow (CBF) disorders related to penumbral tissue [4, 25]. In this setting, it has been proposed that ischemic but hypothetically salvageable brain tissue could be identified by CT perfusion parameters [8, 13]. Based on these considerations and in an attempt to provide further insight into the assessment and the distribution of ischemic and penumbral perfusion alterations, we sought to quantify CBF changes within and around acute SICH by using deconvolution-derived CT perfusion hemodynamic imaging.

MATERIALS AND METHODS

 Patients

We recruited in the study 45 patients with acute supratentorial SICH on unenhanced admission CT scans carried out within 24 hours of symptomatic onset. Time of onset was considered as the last time the patient was known to be neurologically normal. Patients with infratentorial hemorrhage, hematoma related to tumor, trauma, coagulopathy, aneurysms,
vascular malformations, hemorrhagic transformation of brain infarction, intraventricular extension of hemorrhage and patients who had undergone surgical hematoma evacuation were excluded. Disease severity was scored in all patients at entry using Glasgow Coma Scale (GCS) [22]. Hematoma location was classified as basal ganglia or lobar. Hematoma volume was calculated using the formula A × B × C/2 [9]. Informed consent was obtained from each patient or from close relatives before the perfusion CT was performed.

**CT perfusion studies**

CT perfusion examinations were performed by using a single-section CT scanner (CT HiSpeed ZX/i; GE Healthcare, Milwaukee, Wis) equipped for CT perfusion imaging (CT Perfusion; GE Healthcare, Milwaukee, Wis). After hematoma localization on unenhanced CT scanning, the imaging protocol for CT perfusion consisted of a series of 45 CT scans acquired in a single slice (10-mm slice thickness, 80kVp; 200mAs; matrix 512×512; FOV 25-cm; total scan time 50sec) located at the hematoma level during the automatic injection of 50ml of non-ionic contrast agent at the rate of 3.5ml/sec, starting 5 seconds before the initial image. The reference image on non-contrast CT scans was selected on the basis of the level containing the largest volume of blood. All CT perfusion scans were assessed with a deconvolution-based algorithm by using an imaging workstation (Advantage Windows; GE Medical System, Milwaukee, Wis) supplied with a commercial dedicated software (CT Perfusion 2, GE Healthcare, Milwaukee, Wis). CBF, CBV and MTT perfusion maps were generated for each patient. As illustrated in figure 1,

**RESULTS**

**Patients characteristics**

Demographic and clinical features of the 45 patients included in the study are listed in the table 1. Overall, mean hematoma volume was 18.38±25.32 (range=0.72-129.25), whereas the mean time from symptom onset and CT perfusion scanning was 10.10±4.90 hours (range=2.44-21.24 hours; median=9.42 hours).

**CT perfusion measurements**

Mean rCBF and rCBV levels were significantly lower in hemorrhagic core (10.30±6.85ml/100g/min and 0.82±0.456ml/100g, respectively) than in perihematomal low density area (25.32±15.78ml/100g/min and 2.25±0.927ml/100g, respectively) (p<0.001), and in perihematomal area than in normal appearing area.

<table>
<thead>
<tr>
<th>TABLE I. – Demographic and clinical characteristics in 27 patients with acute spontaneous intracerebral hemorrhage (SICH).</th>
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<tbody>
<tr>
<td><strong>Sex (male/female)</strong></td>
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<td><strong>Mean age, years</strong></td>
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<tr>
<td><strong>Mean admission Glasgow Coma Scale</strong></td>
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<td><strong>SICH location</strong></td>
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<td>Basal ganglia</td>
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(68.779±52.393ml/100g/min and 3.692±2.349ml/100g, respectively) and contralateral ROI (68.158±35.021ml/100g/min and 3.675±1.651ml/100g, respectively) (p<0.001). rCBF and rCBV mean levels were similar in normal appearing and contralateral area. Mean rMTT values were significantly higher in both hemor-
hagic core and perihematomal area (5.507±1.832 and 5.74±1.245 seconds, respectively) than in normal appearing area (4.542±0.973 seconds) (p<0.01) and contralateral ROI (4.53±0.796 seconds) (p<0.001). There were no differences in rMTT mean values between hemorrhagic core and perihematomal area, as well as between normal appearing and contralateral areas. Interestingly, when we evaluated absolute mean values of each perfusion parameter, rCBF levels approached irreversible ischemic levels in hemorrhagic core, while they were oligemic in perihematomal low density area and hyperperfu-
sional in both normal appearing and contralateral areas. rCBV levels were decreased only in hemor-
hagic core, whereas none of the ROI examined showed rMTT abnormal levels.

Correlations between perfusion parameters and hematoma volume

In acute SICH patients, rCBF and rCBV mean levels and hematoma volume were inversely and significantly correlated (r=–0.500 and r=–0.589, respectively; p<0.001) in hemorrhagic core. A negative significant correlation was also seen between rCBF and rCBV mean levels and hematoma volume (r=–0.453 and r=–0.423, respectively; p<0.01) in peri-
hematomal area.

DISCUSSION

By employing an accurate topographic approach, we explored perfusion changes occurring within 24 hours after the onset of symptoms in a group of patients with SICH using, for the first time, single-
section deconvolution-derived CT hemodynamic imaging. As expected from previous studies [7, 12, 15, 20, 21, 27], we found that perfusional parameters were concentrically distributed and gradually improved from the core to the periphery but we did not observe zero-flow values within the hemorrhagic core having rCBV at severe ischemic levels, but rCBF and rMTT values which were only close to the threshold of irreversible tissue damage. This apparent paradox may be due to the presence of a thin peripheral border with residual flow and relatively preserved CBV and MTT surrounding the central part of the hemorrhagic core with no flow. This outer band, probably reflecting penumbral tissue evolving into infarction [6], is easily visible in figure 1. On the other hand, as opposed to some studies [12, 21], but in agreement with others [7, 15, 20, 27], no evidence of ischemic penumbra was identified in perihematomal low density area where CBF levels were largely indicative of oligemic tissue not at risk for infarction. More precisely, the concomi-
tant decrease of rCBF and rCBV detected in this area seems to suggest the development of brain edema [16]. These findings appear to be concordant with data coming from animal investigations reporting that perihematomal edema formation is medi-
ated by the clotting of intrahematoma blood with liberation of the remaining serum proteins, espe-
ially thrombin, into the surrounding brain paren-
chyma [11, 23] and by the toxic effect of hemoglobin degradation products derived from erythrocyte lysis [26]. In addition, recent MRI studies with diffusion-
weighted imaging [2] seem to confirm the existence of perilesional vasogenic edema associated to hematoma, even if other reports have failed to demonstrate any sign of edema development around intracerebral hemorrhage [7, 20]. Therefore, as more recently proposed [20], the possibility that perihematomal perfusion disturbances due to regional hypometabolism cannot be completely excluded. In this regard, the strong inverse correlation emerging between perihematomal hemodynamic alterations and hematoma size apparently supports the hypo-
thesis that the amount of bleeding may contribute to injury of brain tissue surrounding the hematoma by mechanical compression of small perilesional blood vessels [1]. Nevertheless, more recent data suggest that perihematomal damage is mainly mediated by the release of coagulation factors and red blood cell lysis components [11, 23, 26] rather than by hematoma-related mass effect [18, 23]. The findings obtained from normal appearing and normal contra-
lateral brain in which, in agreement with others [12, 15] we found hyperperfusional levels that were not associated to a decrease in rCBV and rMTT values are of particular relevance as they most likely indi-
cate luxury perfusion or abnormal vasodilatation [14]. The occurrence of this hyperperfusional pattern in brain regions remote to SICH currently remains a poorly understood phenomenon and could be ascribed to compensatory vasodilatation or, alterna-
tively, to impaired autoregulation or inflammatory response [12]. The main limitations of the study were the inclusion of large blood vessels in CBF cal-
culation that could lead to an overestimation of CBF values [10] and the restriction of the brain coverage to a single section. However, the first-pass bolus-
tracking CT perfusion methodology and the accuracy of the deconvolution process have been validated [3, 24], and the advent of multislice CT scanners could overcome the limited extension of the perfusion analysis. Collectively taken, our findings suggest that dynamic CT perfusion scanning with deconvolution analysis is a powerful method for the evaluation of perfusion deficits associated with acute SICH. They further indicate that perfusion parameters are concentric-
tially distributed and gradually improve from the core to the periphery. In addition, no evidence of ischemic penumbra was found in perihematomal area with rCBF and rCBV levels arguing for oligemia and edema formation. Finally, hyperperfusional values were observed in normal appearing brain tissue located both ipsilaterally and contralaterally to hematoma.

REFERENCES


