Decompressive craniectomy in massive cerebral infarction

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ABSTRACT
Twenty one patients were submitted to decompressive craniectomy for massive cerebral infarct. Ten patients (47.6%) presented a good outcome at the 6 months evaluation, eight had a poor outcome (38%) and three died (14.2%). There was no outcome statistical difference between surgery before and after 24 hours of ictus, dominant and non-dominant stroke groups. Patients older than 60 years and those who had a Glasgow Coma Scale (GCS)<8 in the pre-surgical exam presented worst outcome at six months (p<0.05). Decompressive craniectomy for space-occupying large hemispheric infarction increases the probability of survival. Age lower than 60 years, GCS ≥ 8 at pre-surgical exam and decompressive craniectomy before signs of brain herniation represent the main factors related to a better outcome. Dominant hemispheric infarction does not represent exclusion criteria. Key words: cerebral infarction, decompressive hemicraniectomy, surgical decompression.

Ischemic stroke is a medical emergency and the most common affection of the central nervous system (CNS). This is the second-leading cause of death worldwide and the first cause of morbidity¹. Ischemic stroke correspond to 85% of all strokes with a mortality of 10-50%. Large space-occupying infarction accounts for 1-10% of all supratentorial infarction with signs of elevated intracranial pressure (ICP) and brain herniation usually in the second to the fifth day leading to a mortality rate of 53% to 89%²⁵. The high mortality rate makes some authors call this stroke as...
“malignant” infarct and create the term malignant middle cerebral artery syndrome to describe the rapid development of fatal brain swelling. Large space-occupying infarction is generally secondary to an occlusion of the carotid artery or the M1 segment of the middle cerebral artery (MCA), including or not the anterior cerebral artery (ACA) or the posterior cerebral artery (PCA). Neuroimaging criteria varies between the authors: infarct volume on diffusion-weighted magnetic resonance imaging (MRI) of more than 145 cm³; brain computed tomography (CT) ischemic changes affecting more than two-thirds of the MCA territory and including the basal ganglia; brain CT ischemic changes affecting at least two-thirds of the MCA territory with space-occupying edema; signs on CT of an infarct of at least 50% of the MCA territory, with or without additional infarction in the territory of the anterior or posterior cerebral artery on the same side.

Many studies have suggested that decompressive surgery, consisting of a hemicraniectomy and duraplasty, reduces mortality and improves outcome in patients with massive brain infarctions.

We report our series of 21 patients treated with decompressive craniectomy.

**METHOD**

**Study design**

In this retrospectively designed study, we describe the results of decompressive hemicraniectomy in 21 patients with large hemispheric infarctions at UNICAMP Medical School Hospital from March 2003 to September 2007. We have considered massive cerebral infarction as brain CT ischemic changes affecting at least two-thirds of the MCA territory with space-occupying edema or both MCA and ACA or PCA infarctions leading to a minimum of 50% hemispherical volume compromised. We analyzed gender, age, Glasgow Coma Scale (GCS) on admission and pre-surgical evaluation, clinical status on pre-surgical exam, time from initial symptoms to decompressive craniectomy, length of stay in the hospital and Glasgow Outcome Scale (GOS) 6 months after discharge. Data was subsequently analyzed for comparative study of patients with good to moderate outcome (GOS≥4) and patients with poor outcome (GOS≤3).

**Patient selection**

The decision to perform decompressive craniectomy was based on the presence of a space-occupying large

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**Fig 1. Approach to ischemic stroke.**

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hemispheric infarction on CT scan and the clinical status of the patients. Patients with GCS >13 and no midline shift or basal cistern compression at initial evaluation were managed in the intensive care unit. Neurological deterioration or development of brain herniation signs were indications to decompressive craniectomy as patients initially presented with GCS ≤8 and cistern compression or midline shift at CT scan. Figure 1 shows our approach to ischemic stroke. The patients presented with GCS between 9 and 13 were individually managed.

Surgical technique
A question mark-shaped skin flap based on the ear and a wide craniotomy was performed on the affected side with partial removal of the frontal, temporal, and parietal bones, so that the floor of the middle fossa could be exposed and the bone flap have a minimum of 12 cm diameter. The dura was opened in a “C” shape all over and 1cm distant to the border of the craniotomy. Homologous temporal fascia was placed into the incision for volume-enlarged dural repair (Fig 2). The bone flap was placed in a subcutaneous pocket overlying the abdomen for preservation until subsequent cranioplasty.

Data analysis
To make possible the comparison between the different studies, the outcomes were classified into 4 specific categories (Table 1): grade 1 (G1) functionally independent; grade 2 (G2) mild to moderate disability; grade 3 (G3) severely disabled; and grade 4 (G4) death. Good outcomes were defined as functionally independent or mild to moderate disability. Poor outcomes were defined as severe disability or death. It was not necessary that the study fulfill all the criteria listed, for instance, a G1 outcome could be based only on a GOS of 5, or only on a Barthel index >90, or on a mRS 0-1, not been necessary all three classifications.

Statistical analysis
All data are expressed as mean ± standard deviation (SD). Mann-Whitney U non-parametric tests, t tests and Fisher’s exact test were used for analysis of statistical evidence, with p<0.05 considered significant. Statistical software, SPSS 14.0 (SPSS Inc., Chicago, IL) was used for statistical analysis.

RESULTS
A total of 21 patients (16 males and 5 females) were submitted to decompressive craniectomy during the period analyzed by the study (Table 2). The mean age was 50.09±14.29 years. On admission, the mean GCS was 12±2.42 points (range from 6 to 14).

The mean GCS on immediate pre-surgical evaluation was 8±2.19 points. Nine patients (42.85%) presented with pupillary changes on pre-surgical evaluation; afasia occurred in six cases (28.5%) and hemiplegia presented in all patients.

Seventeen patients (80.95%) had “malignant” MCA infarction and 4 (19.04%) had associated ACA territory infarction. The dominant hemisphere was affected in 6 cases (28.5%) and the non-dominant hemisphere in 15 cases (71.4%).
Table 2. Characteristics of 21 patients included in the study.

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Pct: patient; GCS: Glasgow Coma Scale; GOS: Glasgow Outcome Scale.

Time between onset of symptoms and decompressive craniectomy was less than 24 hours in 10 (47.61%), 24-48 hours in 5 (23.8%), 48-72 hours in 4 (19%) and 72-96 hours in 2 cases (9.52%). We have considered that early surgery was a decompressive craniectomy in the absence of brain herniation signs and in patients with GCS >8 despite the length of hospital stay. It was carried out in 12 cases and late surgery in other 9 patients. Length of stay in the hospital was of 25.76±23.3 days. Surgery improved significantly the GCS of patients comparing the immediate pre-operative scores (8.0±2.19) and immediate post-operative GCS (11.62±4.41), (p<0.05). Occurrence of GCS <9 in the pre-surgical evaluation was associated to a higher length of hospital stay (30.06±25.3 versus 12.0±3.46, p<0.05).

There was no statistical significance in the outcome between men and women, surgery before and after 24 hours of ictus, left and right side stroke groups (p>0.05). Patients older than 60 years presented worst outcome at 6 months (1.75±0.957 versus 3.41±0.87 points in GOS, p<0.05). The presence of brain herniation signs represented a variable associated to poor prognosis, but statistical significance was not reached in this study (2.59±1.33 versus 3.5±0.67 points in GOS, in patients with and without pupillary changes, respectively; p=0.09). According to the Mann-Whitney test, the group of patients who had a poor GCS in the pre-surgical evaluation presented a trend toward poor prognosis, however statistical significance was not demonstrated (2.88±1.14 versus 3.8±0.44 points in GOS, p=0.08). However, when GCS ≤7 is used to predict poor prognosis, we reached statistical evidence (2.72±1.27 × 3.54±0.68 points in GOS, p<0.05).

Ten patients (47.61%) presented a good outcome at the 6 months evaluation. Eight patients who survived had a poor outcome (38%). Three patients of our series died (14.2%) after the surgical procedure, secondary to the presented brain lesion and hemodynamic failure.

**DISCUSSION**

Patients with massive space-occupying hemispheric infarction have a poor prognosis, as mass effect usually develops rapidly with occurrence of clinical deterioration in the first 2 to 4 days. Decompressive surgery has been studied as a way to relieve the intracranial hypertension and tissue shifts related to mass lesions. Bendzus et al. in a case report study, analyzing perfusion CT before and after decompressive craniectomy showed the value of this procedure to spare the ischemic but not infarcted area.

Non-randomized studies suggest that late and early decompressive surgery reduces mortality and increases the number of patients with a favorable functional outcome after massive hemispheric infarction compared to the conservative treatment. Indeed, early decompressive surgery with duraplasty is related with even a better outcome.

Several conservative measures have been proposed to limit brain tissue shifts and reduce intracranial pressure,
including intensive care therapy, mild bed elevation, sedation, hyperventilation, osmotic therapy, hypothermia and others. However, conservative treatment for massive brain infarction has been reported (Table 3) with a high mortality rate and poor outcome despite all those measures, suggesting that they are of limited value.  

The three randomized trials, DECIMAL, DESTINY and HAMLET confirm these findings. DECIMAL and DESTINY were interrupted because of a significant difference in mortality favoring decompressive surgery and HAMLET is still ongoing and aims to include 112 patients.  

Several trials (Table 4) have described the effects of decompressive surgery on functional outcome after space-occupying infarction. As shown in Tables 3 and 4, the decompressive surgery increases the probability of survival from 25% to nearly 70% and the probability of a good outcome (G1 and G2) from 13% to 28%; however, the probability of surviving in a condition requiring as-
sistance and severe disability (G3) increases from 11% to 43%, probably due to the higher number of survivors. Therefore, information about quality of life of survivors is essential for guiding the therapeutic decision. In spite of previous reports in the literature differ with respect to the functional outcome and quality of life after decompressive surgery for space-occupying infarction, even patients with aphasia may improve significantly.

It is unclear which groups of patients benefit most from the procedure. Vahedi et al. demonstrated that surgery was beneficial (p<0.01) independently of age (above and below 50 years), presence of aphasia, and time to randomization (above and below 24 h) when compared to conservative treatment. Kuroki et al. describe that the decompressive surgery outcome is better than the conservative treatment even in patient with more than 70 years old. Patients with the larger infarctions as found in the internal carotid artery (ICA) infarct were more likely to have a poorer prognosis as expected and according to Kilincer et al. surgery for an ICA infarction is not beneficial, unless exceptional cases as very young age, non-dominant hemisphere, and good clinical condition.

Identification of patients at high risk of malignant edema based on radiographic and clinical criteria might allow early hemicraniectomy, defined as a surgery performed before signs of brain stem herniation, as a mean of improving mortality and patient outcome. According to Schwab et al., early hemicraniectomy also reduces the time of critical care therapy from 13.3 to 7.4 days. Although no statistical significance was reached, we observed an important trend toward poor prognosis in the group of patients that had pupillary changes (p=0.09).

Radiographic signs such as early hypodensity of >50% of the MCA territory and/or additional vascular territories (ACA or PCA), ICA infarct 9, midline shift ≥10 mm, effacement of subarachnoid space 19,20, attenuation of corticomedullary differentiation, presence of hydrocephalus may predict which patients will develop malignant edema or bad outcome. Infarct volume of more than 200 cm³ has 91% accuracy to predict malignant hemispheric infarction and the extent of infarct of more than two-thirds of MCA territory has a sensitivity of 93% and specificity of 95% and they are the two most sensitive and specific single explanatory variable for prediction of mortality. On the other hand, brain edema is maximized after 24-72 hours, so an early CT examination should not be considered sensitive enough to predict the final outcome.

Clinical signs such as early clinical deterioration, early nausea or vomiting, and a National Institute of Health Stroke Scale (NIHSS) score ≥20 for left or ≥15 for right hemisphere infarction, pre-operative GCS score ≤7, hypertension or heart failure, and increased peripheral white blood cell count, also may predict which patients will develop malignant edema or have a poor outcome. Lam et al. indicate that a NIHSS >22 is predictive of high mortality. In our study, patients who presented GCS <8 in the pre-surgical exam demonstrated a tendency toward poor outcome, which in our view indicates that the surgical approach should not be delayed until neurological deteriorations occurs.

Although elevated ICP was correlated with higher mortality, ICP monitoring should not be the only parameter in the determination of surgical timing as clinical signs of deterioration or herniation can precede the increase in ICP.

Time from stroke to surgery has also been studied before. Non-randomized series have suggested that outcome is substantially improved if surgical treatment is initiated within 24 h of stroke onset as compared with longer time windows for treatment. Schwab et al. presented benefits of decompression before 24 hours after stroke. In a group of 31 patients, 26 (84%) had a BI >60 at follow up in their study. Gupta et al., however, did not show benefit to surgery<24 hours, probably due to a greater proportion of patients (64%) with signs of herniation before surgery in his group. Vahedi et al. in a systematic review, conclude that the timing of surgery did not affect outcome. We did not observe difference in outcome between patients submitted to surgery before or after 24 hours. We believe the reason for the no significant difference in timing of surgery was the poor general clinical status of patients who presented early to our department. In our view, timing between clinical deterioration and surgery and immediate pre-operative GCS are both more relevant factors than timing from stroke to surgery.

The age has been demonstrated to be an important predictor of outcome in decompressive hemicraniectomy. There are reports of poor functional outcomes and increased mortality in older patients compared to younger. The cut-off point age to predict a good outcome is uncertain. Wijdicks and Diringer studied the natural history of 42 patients with MCA territory infarction, 3 of 11 patients (28%) <45 years died, whereas 22 patients ≥45 years, 90.9% died. Important studies suggest that the optimal recovery occurs in patients less than 50 years and Kilincer et al. use a cut-off point of 55 years and Holtkamp et al. use a cut-off point of 60 years and Kilincer et al. when selected 60 years as a cut-off point, provided one of the strongest predictors of outcome. In our series, there was no statistical difference when used a cut-off age of 50 years. However, patients older than 60 years presented worst outcome at six months follow up.

Offering life-saving treatment for large dominant hemisphere infarcts is controversial, mainly because surgery may leave patients with an unacceptable poor qualify.
nty of life because of hemiplegia and aphasia. The side of the infarct did not have prognostic relevance in our study, as demonstrated by other series. In Gupta et al.'s review, the 27 patients who had decompression of the dominant hemisphere had functional outcome similar to the 111 patients who had non-dominant infarcts. In Kilincer et al., half of the patients had dominant hemispheric infarction with global aphasia preoperatively, 6/7 patients in the good outcome group had a dominant hemispheric infarction and most of the patients showed considerable improvements in their aphasia, a finding confirmed by other authors. Therefore, we believe infarction side should not be exclusion criteria for surgery. We agree with some authors that language deficits may be of small consequence in patients who are severely disabled by hemiplegia; also, non-dominant hemisphere strokes can lead to severe depressive, abulic, or neglect states that may interfere with rehabilitation efforts and are as disabling as aphasia. On the other side, global disability scales such as the BI, mRS, and GOS may emphasize mobility as opposed to language dysfunction.

It is not clear which patients may avoid severe disability after the procedure. A large number of patients or relatives (70%) stated that they would undergo the procedure again if faced to the same situation. Even 79% of the patients and their family are satisfied with the surgical results.

In conclusion, decompressive craniectomy for space-occupying large hemispheric infarction increases the probability of survival that can yield good functional outcomes in some cases. Careful patient selection, made on an individual basis, and early operation may improve the functional outcome for large hemispheric infarction. Information about quality of life of survivors is essential for guiding such decisions because most patients require extensive rehabilitative therapy and lifelong assistance.

There are limitations in our study. Although we present important data about decompressive surgery, it is a non-randomized retrospective study with results that need confirmation by larger randomized trials. Although there is not a consensus for the surgical treatment of massive hemispheric infarction, we recommend: (1) in patients under 60 years old; (2) in patients with CT scan evidence of massive cerebral infarction with GCS ≥8; (3) decompressive craniectomy before signs of brain herniation if possible. Dominant hemispheric infarction does not represent an exclusion criteria.

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