# Embolization of the Middle Meningeal Artery Effectively Treats Refractory Chronic Subdural Hematoma: A Systematic Review

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#### Abstract

Chronic subdural hematoma (CSDH) formation mechanism is very complex, and has not entirely understood. It represents a frequent type of intracranial hemorrhage, and is very common disease in Neurosurgery practice, especially in older patients. Various surgical treatments have been proposed for the treatment of CSDH. The rate of recurrence in CSDH after surgery ranges from 5% to 30%, repeated surgery must be considered. But in some cases subdural collections are still persistent. Endovascular embolization of the middle meningeal artery (MMA) is an option for treatment of refractory CSDH. We review all cases that were treated with embolization to assess the effect of this intervention. Our review revealed 6 papers with a total enrollment of 14 patients were treated with MMA embolization for refractory chronic subdural hematoma without any postoperative complication or recurrence. In this study we suggest MMA embolization as an alternative for treatment of non-curable CSDH, especially for old people with systematic diseases, who cannot tolerate repeat surgery.

#### Introduction

Chronic subdural hematoma (CSDH) is a common neurological condition, particularly for older patients (1, 2), with an estimated incidence of 7.32 per 100,000 people per year for individuals above 70 years old (3). Levels of mortality associated with CSDH range from 1.5% to 8% (4). The pathogenesis of CSDH formation remains unclear, although new imaging and molecular biology techniques have improved our understanding of these processes (5, 6). The rate of recurrence in CSDH after surgery with any of actual techniques (Burr-hole surgery with irrigation and drainage, twist drill, craniotomy) ranges from 5% to 30% (7, 8, 9). Patients who have high risk factors for recurrence, especially with coagulopathy10 or under anticoagulant therapy (11), hematoma may be recur after the first intervention, and repeated surgery must be considered. The treatment options remains controversial. A number of techniques are used to manage recurring CSDH, including burr hole plus drainage, repeated irrigation, the installation of reservoirs and aspiration

(12), subdural peritoneal shunts (13), capsule removal with large craniotomy, and the endoscopic perforation of the septum (14).

For elderly patients, however, efforts should be made to avoid the repeated application of invasive procedures. Embolization of the middle meningeal artery (MMA) is another treatment option for frequently recurring CSDH. This procedure is often used to treat dural arteriovenous shunts or as a preoperative treatment for meningioma tumors. MMA embolization was first used to treat CSDH in 2000, preventing CSDH recurrence in a patient with cirrhosis of the liver (15).

Here we have evaluated whether MMA embolization effectively prevents the recurrence of hematoma in patients with CSDH.

A search of the literature revealed six reports concerning 14 patients. These patients were 55–87 years old and had follow-up periods of 3–18 months. None of the patients treated with MMA embolization for refractory CSDH experienced any postoperative complications or hematoma recurrence. This analysis suggests that MMA embolization is an effective, alternative treatment for noncurable CSDH, especially for older patients with systematic diseases who cannot tolerate multiple surgeries.

## Methods

To identify relevant case reports in the scientific literature, the Medline, PubMed, Embase, and ISI Web of Science databases were searched using the key words "chronic subdural hematoma" combined with "recurrence" and "embolization" or "middle

meningeal artery". Only patients who received MMA embolization for the treatment of recurrent CSDH were included in the analysis. Articles not available in English were excluded. After identifying relevant articles, all studies referenced by these manuscripts were also reviewed. Our search identified six publications that describe outcomes for 14 patients treated with MMA embolization. Included studies were case series that presented class III evidence.

#### Results

Among the 14 patients who met our inclusion criteria were 12 males and two females, with ages that ranged from 55 to 87 years (Table 1). Seven patients had suffered a previous head trauma; three patients had not experienced a trauma; trauma histories were not available for three patients; and for patient number 9 the history of head trauma was unclear. Patients presented with symptoms that included abnormal behavior, disorientation. speech abnormalities, dementia, motor weakness, hemiparesis, gait disturbances, and loss of consciousness.

Three patients were taking the anticoagulant Warfarin (Coumadin), one patient had coagulopathy from cirrhosis of the liver, one patient was on hemodialysis because of chronic renal failure, one patient was receiving chemotherapy for peritoneal mesothelioma, one patient was receiving treatment dentatorubral-pallidoluysian atrophy, one patient was suffering from alcoholism, and six patients had no previous medical histories. Based on computed tomography scans, six patients had bilateral CSDH, whereas eight patients had onesided CSDH.

TABLE 1
Patients' characteristics

| author                            | Patient<br>No. | Age | sex | Head<br>Trauma? | Pre-treatment<br>Symptoms               | Past medical<br>history                        | CSDH<br>on CT             | pre-embolization<br>treatments        |                     | Embolization<br>Materials     | Follow-up<br>(months) |
|-----------------------------------|----------------|-----|-----|-----------------|---|--|---------------------------|---------------------------------------|---------------------|-------------------------------|-----------------------|
|                                   |                |     |     |                 |   |  |                           | major<br>treatments                   | other<br>treatments | Materiais                     | (montns)              |
| Mandai<br>et al. <sup>15</sup>    | 1              | 59  | М   | no              | LOC,<br>hemiparesis                     | coagulopathy<br>from cirrhosis<br>of the liver | 1)<br>bilateral<br>2) MLS | burr hole,<br>irrigation              | Ommaya<br>reservoir | PVA<br>particles              | 7                     |
| Hirai<br>et al. <sup>27</sup>     | 2              | 81  | М   | yes             | motor<br>weakness, gait<br>disturbance  | warfarin<br>(aortic valve<br>replacement)      | bilateral                 | burr hole,<br>irrigation              | irrigation          | PVA particles, platinum coils | 10                    |
|                                   | 3              | 63  | М   | yes             | abnormal<br>behavior,<br>disorientation | warfarin<br>(cardiomyop-<br>athy)              | right<br>side             | burr hole,<br>irrigation,<br>drainage | irrigation          | platinum<br>coils             | unavailable           |
| Ishihara<br>et al. <sup>28</sup>  | 4              | 78  | М   | yes             | unavailable                             | hemodialysis<br>(chronic renal<br>failure)     | left side                 | burr hole,<br>drainage                | none                | 20% NBCA                      | 16                    |
| Mino<br>et al. <sup>29</sup>      | 5              | 73  | М   | yes             | gait<br>disturbance                     | none   | bilateral                 | burr hole,<br>drainage                | irrigation          | gelatin<br>sponge, GDC        | 6                     |
|                                   | 6              | 79  | М   | no              | speech<br>disturbance,<br>dementia      | none   | right<br>side             | burr hole,<br>drainage                | Craniotomy          | gelatin<br>sponge, GDC        | 6                     |
|                                   | 7              | 65  | M   | yes             | hemiparesis                             | none   | left side                 | burr hole,<br>drainage                | none                | gelatin<br>sponge, GDC        | 6                     |
|                                   | 8              | 75  | M   | yes             | gait<br>disturbance                     | none   | bilateral                 | burr hole,<br>drainage                | none                | gelatin<br>sponge, GDC        | 6                     |
| Tsukamoto<br>et al. <sup>30</sup> | 9              | 64  | M   | unclear         | consciousness<br>disturbance            | DRPLA  | bilateral                 | Burr hole,<br>drainage                | steroids            | PVA particles                 | 18                    |
| Hashimoto<br>et al. <sup>31</sup> | 10             | 79  | M   | yes             | none                                    | peritoneal<br>mesothelioma                     | right<br>side             | Burr hole,<br>irrigation              | none                | 16% NBCA                      | 3                     |
|                                   | 11             | 55  | М   | unavailable     | unavailable                             | Heavy drinker                                  | left side                 | Burr hole,<br>irrigation              | none                | 20% NBCA                      | unavailable           |

| 12 | 87 | M | unavailable | unavailable    | none        | left side | Burr hole, | none | 16% NBCA   | unavailable |
|----|----|---|-------------|----------------|-------------|-----------|------------|------|------------|-------------|
|    |    |   |             |                |             |           | irrigation |      |            |             |
| 13 | 73 | F | unavailable | unavailable    | none        | bilateral | Burr hole, | none | 18% NBCA   | unavailable |
|    |    |   |             |                |             |           | irrigation |      | 200-μm PVA |             |
|    |    |   |             |                |             |           |            |      | particles  |             |
| 14 | 68 | F | no          | Disorientation | warfarin    | left side | Burr hole, | none | 20% NBCA   | 4           |
|    |    |   |             |                | (Cerebral   |           | irrigation |      |            |             |
|    |    |   |             |                | infarction, |           |            |      |            |             |
|    |    |   |             |                | pulmonary   |           |            |      |            |             |
|    |    |   |             |                | Embolism)   |           |            |      |            |             |

CSDH, chronic subdural hematoma; DRPLA, dentatorubral-pallidoluysian atrophy; GDC, Guglielmi detachable coils; MLS, midline shift; NBCA, N-butyl 2-cyanoacrylate; LOC, loss of consciousness; PVA, polyvinyl alcohol

TABLE 2 Factors associated with the recurrence of CSDH\*

| Patient factors                      | Laboratory findings | Imaging findings           | Surgical factors    |
|--------------------------------------|---------------------|----------------------------|---------------------|
| -age                                 | -liver dysfunction  | -brain atrophy             | -duration of        |
| -sex                                 | -renal failure      | -high and mixed-density    | drainage            |
| -low GCS                             | -thrombocytepenia   | CSDH                       | -residual air after |
| -low GOS at discharge                | -coagulation-factor | -bilateral CSDH            | surgery             |
| -hemodialysis                        | deficiency          | -midline shift more than 5 |                     |
| -dementia                            |                     | mm                         |                     |
| -diabetes                            |                     | -width of the hematoma     |                     |
| -epilepsy                            |                     | -cranial base hematoma     |                     |
| -APs or ACs                          |                     | -septum formation          |                     |
| -chemotherapeutic agents             |                     |                            |                     |
| -intracranial hypotension (CSF shunt |                     |                            |                     |
| surgery)                             |                     |                            |                     |
| -hypertension                        |                     |                            |                     |
| -alcoholism                          |                     |                            |                     |
| -postoperative patient posture       |                     |                            |                     |

\*These factors are still controversial

AC, anticoagulant; AP, antiplatelet; CSF,crebrospinal fluid; GCS, Glasgow coma scale, GOS, Glasgow outcome scale

The first treatment that these patients received were: 1) burr hole plus drainage and irrigation, 2) burr hole plus drainage, or 3) burr hole plus irrigation. Several techniques were also used to treat hematoma recurrence, namely an additional surgery, irrigation, an Ommaya reservoir, and steroids. The materials used to embolize the MMA were: polyvinyl alcohol particles, 16–20% N-butyl 2-cyanoacrylate, gelatin sponges, and Guglielmi detachable coils.

No complications were reported after the MMA embolization procedure. Follow-up times ranged from 3 to 18 months, although follow-up times for patients 3, and 11–13 were not available. Importantly, all 14 patients experienced no recurrences or enlargements of CSDH following embolization of the MMA.

### Discussion

CSDH is one of the most common types of intracranial hemorrhage and it is associated with high levels of morbidity and mortality (1.5-8%) (4, 16, 17). Patients with advanced age are more likely to have CSDH because of atrophy of the brain parenchyma. This atrophy enlarges subarachnoid spaces, thereby stretching veins that bridge these spaces (18). Stretched veins are more fragile and tend to rupture, releasing venous blood into the subdural space (19). Men are twice as likely as women to have CSDH because they experience more head injuries (20). Single or double burr holes with closed drainage systems (with or without irrigation) are currently treatments of choice for this condition, but many others techniques have been described (21). The recurrence of CSDH following

surgery is not uncommon, and can result from poor re-expansion of the brain, or inflammation and hemorrhage of the outer membrane (22).

Several factors are associated with CSDH recurrence, including advanced age, a tendency to bleed, brain atrophy, alcohol abuse, kidney and liver diseases, diabetes, epilepsy, anticoagulants, chemotherapy intracranial agents, hypotension, hemodialysis, the density of the hematoma, the width of the hematoma, insufficient intraoperative drainage, postoperative accumulation of air in the subdural space, bilateral CSDH, postoperative position, the Glasgow coma scale, and the Glasgow outcome scale (7, 11, 16, 17, 23-26) (Table 2). Analyses concerning these associations, however, have yielded inconsistent results.

The treatment of recurrent CSDH remains controversial and many techniques have been used to manage this condition, including burr holes plus drainage, repeated irrigations, the installation of reservoirs and aspiration (12), subdural peritoneal shunts (13), capsule removal with large craniotomy, endoscopic perforation of the septum (14), and MMA embolization (15, 27–32).

Embolization of the MMA to treat CSDH was first reported by Mandai et al. (15) for patients with coagulopathy (cirrhosis of the liver). Application of this technique resulted in very positive patient outcomes. In one Japanese article Takahashi et al. (32) reported results concerning three patients with recurrent CSDH that had been subjected to several unsuccessful drainage procedures. After MMA embolization no hematoma

enlargements were seen and all hematomas gradually and completely resolved (32). Tanaka et al. (33) performed superselective angiography of the MMA in a series of 35 patients. They found diffuse dilation of the MMA and scattered networks of abnormal vasculature. which seemed macrocapillaries within the outer membrane (29, 33). To understand the relationship between the MMA and the outer membrane in CSDH, Tanaka et al. (34) studied histological features of the outer membrane and observed many communicating arteries, i.e., arteries that originated from branches of the MMA and then entered the outer membrane. They concluded that the MMA provides nutrients to the abnormal membrane that surrounds the hematoma via these communicating arteries. As direct arterial pressure of the MMA may rupture these communicating arteries, MMA embolization may prevent or delay CSDH recurrence.

The most common materials used for the embolization procedure were: N-butyl 2cyanoacrylate, Guglielmi detachable coils, polyvinyl alcohol particles, and gelatin sponges. Using a low concentration of liquid 2-cyanoacrylate increases therapeutic effect by embolizing more peripheral vessels associated with the MMA with a low risk of catheter adhesion (28, 31). To prevent the aberrant flow of blood into the ophthalmic artery, it is necessary to perform a pre-embolization study of the MMA pathway and its anastomosis with the internal carotid artery through the inferolateral trunk, or the feeding vessel to the facial nerve. If there is anastomosis between the MMA and the

ophthalmic artery, embolization of the proximal MMA using coils or gelatin sponges should be considered (31).

Here we have review 14 cases in which MMA embolization was used to treat CSDH. None of these patients had post-operative complications, CSDH recurrence, or CSDH enlargement. It should be acknowledged, however, that this review does not contain Class I or Class II data, and that this analysis should be supplemented with more definitive data (e.g., randomized trials) when they become available. Results to date indicate that embolization of the MMA represents an effective option for treating recurrent CSDH. It is is good option for patients who cannot tolerate invasive intervention such as repeated surgery (burr hole), capsule removal with large craniotomy or perforation of the septum with an endoscope. It can use as adjuvant therapy in treating recurrence of chronic subdural hematoma.

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