Similar Safety in Centers with Low and High Volumes of Endovascular Treatments for Unruptured Intracranial Aneurysms: Evaluation of the Analysis of Treatment by Endovascular Approach of Nonruptured Aneurysms Study

BACKGROUND AND PURPOSE: The outcome of treatment for unruptured intracranial aneurysm by surgery or endovascular therapy appeared to be related to the volume of patients treated by the hospital. We performed an analysis in the ATENA series to determine whether the outcome of endovascular treatment was different in hospitals with low and high volumes of cases.

MATERIALS AND METHODS: The ATENA series included 649 patients with 739 unruptured intracranial aneurysms. Patients were classified into 2 groups. Group A had 171 patients with 197 aneurysms in 13 centers that treated ≥20 patients; group B, 478 patients with 542 aneurysms in 14 centers that treated ≥21 patients.

RESULTS: Groups A and B had similar patient populations and aneurysm characteristics. Stent placement was used in group B more frequently than in group A (9.6% versus 2.5%, \(P = .0016\)). The global rate of adverse events was not significantly different in groups A and B (16.0% and 14.4%, respectively). Thromboembolic events and intraoperative rupture were not significantly more frequent in group A than in group B. One-month mortality and morbidity rates were not significantly different in groups A (2.3% and 1.8%, respectively) and B (1.0% and 1.7%, respectively). Anatomic outcomes for groups A and B were not significantly different.

CONCLUSIONS: Clinical and anatomic outcomes of endovascular treatments for unruptured intracranial aneurysms were similar in hospitals with low and high volumes of cases.

ABBREVIATIONS: ACA = anterior cerebral artery; AcomA = anterior communicating artery; ATENA = Analysis of Treatment by Endovascular Approach of Nonruptured Aneurysms; CH = centre hospitalier; CHU = Centre Hospitalier Universitaire; CI = confidence interval; GP = group; HTA = hypertension artérielle; ICA = internal carotid artery; MCA = middle cerebral artery; mRS = modified Rankin Scale; NA = not applicable; TE = thromboembolic complications; UIA = unruptured intracranial aneurysm; VB = vertebrobasilar system
Permanent morbidity and mortality rates of treated patients were evaluated at 1 month. Morbidity was defined as an mRS score from 2 to 5. When a preoperative mRS was >1, morbidity was defined as any increase in the mRS score.

The neuroradiologist who performed the treatment evaluated the degree of aneurysmal occlusion on the basis of the modified 3-point classification scale of Raymond et al: complete occlusion, neck remnant, and aneurysm remnant.

**Group Definitions**
The ATENA study centers were classified according to the number of patients treated during the study period. To divide the centers into approximately equal groups, we used an arbitrary limit of 20 patients treated during the period of inclusion.

Group A included 13 centers that treated ≤20 patients: In 1 center, ≤5 patients were treated; in 2 centers, 6–10 patients were treated; in 5 centers, 11–15 patients were treated; and in 5 centers, 16–20 patients were treated.

Group B included 14 centers that treated >20 patients: In 8 centers, 21–30 patients were treated; in 4 centers, 31–40 patients were treated; and in 2 centers, >40 patients were treated.

Separate analyses were also conducted by using different cutpoints, including 1 patient treated per month (≤17 patients or >17 patients during the study period) and 20 patients treated per year (≤28 patients or >28 patients during the study period).

**Statistical Analysis**
Data management and statistical analyses were independently conducted by the Clinical Research Unit of Reims University Hospital (F.V.) to calculate patient demographics, aneurysm characteristics, treatment, adverse events, and outcome. Mean and frequency comparisons were performed with the Student t test and χ² test or the Fisher exact test, respectively. Differences were considered significant at P = .05. Statistical analysis was performed with SAS Version 8.0 software (SAS Institute, Cary, North Carolina).

**Results**
Analysis was performed by using different cut-points: 17, 20, and 28 patients during the study period (see “Materials and Methods”). Results were similar regardless of the cut-point used, and only results obtained with the cut-point of 20 patients are presented in this section.

**Patient Population**
The distributions of sex, age, and risk factors (smoking and hypertension) of groups A and B were not significantly different (Table 1).

### Table 1: Patient characteristics in centers with low (group A) and high (group B) numbers of cases/hospital

|                  | Group A (n = 171) | Group B (n = 478) | P Value |
|------------------|-------------------|-------------------|---------|
| **Sex ratio**    |                   |                   |         |
| Women            | 122               | 346               | .80     |
| Men              | 49                | 132               |         |
| **Age**          |                   |                   |         |
| Mean (yr)        | 52.1 ± 12.2       | 50.8 ± 11.0       | .21     |
| Range (yr)       | 22–80             | 23–83             |         |
| **HTA**          |                   |                   |         |
| No. %            | 51/168* (30.4%)   | 141/468* (30.1%)  | .95     |
| **Smokers**      |                   |                   |         |
| No. %            | 62/167* (37.1%)   | 198/470* (42.1%)  | .25     |

* The information was not available in some cases.

### Table 2: Aneurysm characteristics in centers with low (group A) and high (group B) numbers of cases/hospital

|                  | Group A (n = 197) | Group B (n = 542) | P Value |
|------------------|-------------------|-------------------|---------|
| **Location**     |                   |                   |         |
| ICA              | 89                | 235               | .54     |
| ACA, AcomA       | 30                | 107               |         |
| MCA              | 60                | 158               |         |
| VB               | 18                | 42                |         |
| **Aneurysm size (mm)** |   |                   |         |
| 1–3              | 31                | 100               | .65     |
| 3–6              | 86                | 218               |         |
| 7–10             | 54                | 161               |         |
| 11–15            | 26                | 63                |         |
| **Dome-to-neck ratio** |   |                   |         |
| ≤1.5             | 109               | 278               | .30     |
| >1.5             | 88                | 264               |         |

**Aneurysm Characteristics**
The aneurysm characteristics (location, aneurysm size, dome-to-neck ratio) of the groups were not significantly different (Table 2).

**Modalities of Endovascular Treatment of UIAs**
The numbers of nonselective and selective treatments were not significantly different between groups A and B (Table 3). However, the selective treatments were performed significantly differently in group A compared with group B: The standard coiling technique was more frequently used in group A versus group B (62.4% and 50.4%, respectively; P = .001). Conversely, stent placement was more frequently used in group B versus group A (9.6% and 2.5%, respectively; P = .0016). The remodeling technique was used with a higher fre-
Adverse Events Related to Endovascular Treatment of UIAs

The global rate of adverse events was not significantly different in groups A and B (16.0% and 14.4%, respectively) (Fig 1 and Table 4). Similarly, the rate of specific adverse events (thromboembolism and intraoperative rupture and device-related complication) was not significantly different in groups A and B (12.8% and 12.5%, respectively). Thromboembolic events and intraoperative rupture were not significantly different in group A than in group B (Table 4).

One-Month Clinical Results

One-month mortality and morbidity rates were not significantly different in groups A and B (Table 5).

Duration of the Hospital Stay

The median duration of a hospital stay was the same in each group: median \( \text{Group A} = 5 \) days and median \( \text{Group B} = 5 \) days \((P_{\text{Mann-Whitney}} = 0.09)\). In group A, the duration of the hospital stay was 1–5 days for 147 procedures (78.6%), 6–10 days for 29 procedures (15.5%), and >10 days for 11 procedures (5.9%). In group B, the duration of the hospital stay was 1–5 days for 440 procedures (85.8%), 6–10 days for 44 procedures (8.6%), and >10 days for 29 procedures (5.7%). The difference between the groups was significant \((P = .03)\).

Anatomic Results

The postoperative aneurysmal occlusion was not significantly different in groups A and B (Table 6).

Discussion

The ATENA study was conducted in 27 neurointerventional centers in France and Canada from June 2005 to October 2006. It showed that the endovascular treatment of unruptured intracranial aneurysms was feasible in a high percentage of cases with low morbidity and mortality rates. However, an important unanswered question was whether the results were similar among centers that treated high and low volumes of patients—that is, it was unknown whether the treatment in a center with a high volume of cases was associated with a better outcome.

Previously, patients with ruptured or unruptured aneurysms with surgery or endovascular treatments have been studied to evaluate the relationships between physician experience, volumes of patients treated, and clinical and anatomic outcomes. Most series were based on a retrospective analysis of a database of hospital discharges and were fraught with the limitations of this type of evaluation.

Previous analyses were conducted in the patient population that had been treated surgically. Solomon et al reviewed all discharges in the state of New York from 1987 to 1993 that had been recorded in the Statewide Planning and Research Cooperative System of the New York Department of Health. They analyzed the outcomes of all patients who underwent craniotomy for cerebral aneurysms \((n = 4034)\). They found that hospitals that performed >30 craniotomies per year had a 43% lower mortality rate than hospitals that performed fewer craniotomies (8.8% versus 15.5% mortality rates, respectively). They found similar results for patients who underwent craniotomy for unruptured cerebral aneurysms. Hospitals that performed >30 craniotomies per year had a 43% lower mortality rate than hospitals that performed fewer craniotomies (4.6% versus 8.1% mortality rates, respectively). Similar results were reported for older patients with ruptured aneurysms treated by the surgical approach. The 30-day mortality rates for patients with subarachnoid hemorrhage who were treated surgically were 20.5%, 18.4%, and 14.3% in hospitals averaging <1, between 1 and 5, and ≥5 interventions per year, respectively. Similar results were observed in a more recent series that studied mortality and morbidity after surgical treatment of unruptured intracranial aneurysms in the United States (1996-2000). The hospitals that treated ≥20 patients/year had fewer adverse outcomes (15.6%) and lower mortality rates (1.6%) compared with hospitals with <4 cases/year (23.8% adverse outcomes and a 2.2% mortality rate).

One study analyzed the effects of endovascular services and volume of cases/hospital on cerebral aneurysm treatment outcomes. They showed that patients were less likely to die in the hospital when treated at institutions that more frequently used coil embolization and angioplasty for vasospasm. However, the volume of cases/hospital was not independently associated with in-hospital deaths. Johnston et al analyzed the data base from the Office of Statewide Health Planning and Development of California for the period of 1990-1998. They showed that compared with surgery, endovascular therapy of unruptured aneurysms was associated with a lower risk of adverse outcomes and in-hospital deaths. The volume of cases/hospital was also associated with outcome.

A subsequent analysis of patients treated for ruptured or unruptured aneurysms was conducted in New York State. Again, that study showed that the volume of cases/hospital and the propensity of a hospital to use endovascular therapy were both independently associated with better outcomes. In cases that involved clipping of either ruptured and unruptured aneurysms, a higher volume of cases/hospital was associated with a better outcome. In cases that involved embolization, the volume of cases/hospital was either not significantly associated with outcome or demonstrated a statistically weaker association than clipping. In addition, Hoh et al conducted a retrospective cohort study with the Nationwide Inpatient Sample of 1996-2000. They demonstrated that hospitals with >23 admissions per year had fewer adverse outcomes (5.2%) than hospitals with <4 admissions per year (17.6%). The mortality rate was lower at hospitals with high volumes of cases (1.0% versus 3.7%), but the difference was insignificant. Fi-

### Table 3: Modalities of treatment in centers with low (group A) and high (group B) numbers of cases/hospital

| Modality             | Group A | Group B | P Value |
|----------------------|---------|---------|---------|
| Nonselective treatment | 5       | 7       | .31     |
| Selective treatment   | 123     | 273     | .001    |
| Coiling               | 62      | 209     | .07     |
| Remodeling            | 5       | 52      | .0016   |
| Stenting              | 2       | 1       | NA      |

*Boston Scientific, Natick, Massachusetts.*
nally, the length of stay was shorter at hospitals with high volumes of cases.

We used a completely different approach in our analysis of the relationship between the volume of cases/hospital and the outcome of endovascular treatment for UIA. We did not use a discharge data base but the results from a prospective consecutive multicenter controlled study. This methodology can avoid much of the bias that might be encountered in a retrospective analysis of a discharge data base, but it is also associated with some limitations (see below). The population of the ATENA study was divided into 2 groups according to the volume of cases/hospital. The groups were defined to ensure approximately the same number of centers in each group. Group A included 13 centers that treated ≤20 patients, and group B included 14 centers that treated >20 patients during the inclusion period. With other cut-points (see “Materials and Methods”), similar results were obtained (not shown). Although the patient and aneurysm characteristics were similar in both groups, the endovascular treatment was performed differently in groups A and B. The standard coiling technique was more frequently used in group A (62.4% versus 50.4% in group B, \( P = .001 \)), and stent placement was more frequently used in group B (9.6% versus 2.5% in group A, \( P = .0016 \)). These results suggested that centers with high volumes of cases may facilitate the use of complex techniques like stent placement.

The most important factor in evaluating any treatment is certainly clinical outcome, as assessed by morbidity and mortality. Morbidity was not significantly different in groups A and B (1.8% and 1.7%, respectively). The mortality rate was higher in group A (2.3% versus 1%), but the difference was not significant. Hoh et al\(^9\) reported somewhat different outcomes. They found that discharge to a facility other than home occurred at a rate of 5.2% from hospitals with high volumes of cases compared with 17.6% from hospitals with low volumes of cases. However, with findings similar to ours, they reported a lower mortality rate at hospitals with high volumes of cases (1.0% versus 3.7%), but the difference was not significant.

Several factors may explain the discrepancies between studies. Our series was conducted more recently (2005–2006) and during a shorter period than the analysis of Hoh et al\(^9\) (1996–2000).\(^9\) Endovascular treatment started in France in 1993–1994; thus, most teams participating in the ATENA study had long experience with the endovascular treatment of unruptured aneurysms. This was probably not the case in the series of Hoh et al. Moreover, the development of more appropriate devices (coils, microcatheters, microguidewires, etc) has facilitated the performance of endovascular treatments. Also, the series of Hoh et al included a very large (81) number of centers compared with the number included in our series (27); thus, the levels of clinical experience were presumably more heterogeneous in the series of Hoh et al. Finally, we defined high- and low-volume hospitals differently than Hoh et al did. They defined high-volume hospitals as those with >23 admissions per year and low-volume hospitals as those with <4 admissions per year; in contrast, we defined high- and low-volume hospitals as those with greater or less than 20 admissions during 17 months, respectively.

We found that the anatomic outcomes were similar in both groups. Despite the difference in treatment techniques (see above), the relative percentages of complete occlusion, neck remnant, and aneurysm remnants were not different in groups A and B. However, this result was based on the occlusion rates determined by the practitioners that performed the procedures. A core lab analysis of anatomic results is necessary to confirm that the results were similar in both groups.

Our study had several limitations. First, it was not possible to evaluate the results according to the physician volume. Because 1, 2, or more physicians can be involved in the treatment of a single patient, it was not possible to calculate the physician volume by simply dividing the number of patients treated by the number of physicians in each center. Second, it was not possible to evaluate the effect of physician experience because we had no information regarding the years of experience or the number of unruptured aneurysms treated before the study for each physician. It was previously shown that the risk of complications with coil embolization of unruptured aneurysms was dramatically reduced with physician experience.\(^11\) Third,
we were unable to take into account the global experience within each center for the treatment of intracranial aneurysms, including ruptured aneurysms, and of neurovascular procedures in general. Finally, our definitions of high- and low-volume centers for groups A and B were arbitrary; in fact, there is a continuum from group A to group B, and this may partially explain our results. However, results were similar whatever the cut-point used to define the low and high-volume centers.

Summary
Our analysis demonstrated that at least in France (and Canada), clinical and anatomic outcomes of endovascular treatments for unruptured intracranial aneurysms are similar in centers with low and high volumes of cases/hospital. This is probably related to the fact that most neurointerventional teams participating in ATENA have had long intensive experience with the endovascular treatment of intracranial aneurysms (unruptured and ruptured).

Appendix

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