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**Case Report** 

### Incidental intracranial saccular aneurysm in a patient with post-Covid-19 headache: What to do with the incidentaloma?

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# Introduction

In the last decades, there was a growing increase in incidentalomas.<sup>1.6</sup> In the radiological investigation of patients with headaches, it is not infrequent to find incidentalomas.<sup>7.8</sup> Incidentaloma is classically defined when, during a radiological investigation, a tumoral lesion is identified, and such lesion is not related to the clinical condition that justified the request for the examination. In other words, it was an incidental finding, *i.e.*, a lesion found by coincidence without any clinical symptoms or suspicion.<sup>9</sup>

The term incidentaloma also serves to designate other lesions found incidentally, such as in the case of an aneurysm. Unruptured intracranial saccular aneurysms develop in cerebral arteries over months or years and are rarely symptomatic.<sup>10-14</sup> Eventually, during an imaging evaluation, these brain aneurysms are discovered. For many of these aneurysms, the chosen approach is conservative, depending on the size, location, associated diseases, risk of bleeding, family history, among other factors that modify the risk of rupture.<sup>13,15</sup> If the approach is conservative, it is essential to follow up with serial reassessment by angiography (e.g., MR angiography).

Approximately 2–4% of the population harbours an unruptured intracranial aneurysm<sup>16-21</sup>, and up to 30% and 42% of these patients may have multiple aneurysms in imaging or autopsy studies, respectively.<sup>22:23</sup> With increased utilization of noninvasive imaging, the detection of these aneurysms has become very common, the so-called incidentaloma.

The discovery of an incidental unruptured intracranial saccular aneurysm implies decision-making strategy about its treatment, either conservative or surgical (open or an endovascular approach), depending on the size, location, angioarchitecture of the aneurismal sac, associated dis

eases, age, risk of bleeding, family history, among other factors that may modify the risk of rupture.<sup>13,15</sup> If the approach

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is conservative, following up the aneurysm with a serial reassessment by magnetic resonance angiography (MRA) is essential. This article will discuss the best management of an elderly patient with an incidentally discovered saccular aneurysm originated in the left middle cerebral artery (MCA).

#### **Case report**

An 83-year-old woman in January 2021 developed persistent headache and decline in cognitive functions that appeared after the acute onset of Covid-19. She is a regular smoker and has hypertension treated with bisoprolol, oral

anticoagulants (rivaroxaban) to control cardiac arrhythmia, and oral rosuvastatin to control dyslipidemia. She denied a family history of a brain aneurysm. She was evaluated with computed tomography (CT), magnetic resonance imaging (MRI), and digital angiography to investigate the causes of symptoms (Figures 1 and 2). CT scan showed an image compatible with aneurysmal dilatation in the left middle cerebral artery bifurcation, confirmed by contrast magnetic resonance angiography (Figure 1). According to digital angiography, the saccular dilatation had the following dimensions: 7.7 x 4.7 x 3.5 mm, with a neck measuring 2.5 x 2.4 mm (Figure 2). In this case, after evaluating the advantages and disadvantages of conservative versus preventive surgical intervention (clipping or coiling), considering the patient's advanced age and expectations, associated morbid conditions, we decided on a non-surgical approach, with conservative measures such as blood pressure control, use of beta-blockers, statin, and follow-up by CE-MRA.

# Comments

Despite advancements in the management of subarachnoid hemorrhage (SAH) due to aneurysmal rupture, overall case fatality is quiet high. (40-60%).<sup>14</sup> Perhaps, for this reason, patients harboring unruptured intracranial aneurysms have performed preventive surgical intervention for many decades to eliminate any conceivable aneurysm rupture.

Accordingly, managing of the patient with unruptured intracranial aneurysms<sup>24</sup> remains a challenge with medical, ethical, and legal implications. Some key questions remain: what is the risk of rupture for a specific aneurysm in a patient in which specific risk factors can be identified? So, we should consider important clinical and morphological points before deciding when and how an accidental aneurysm should be treated.

The prospective arm of the ISUIA<sup>14</sup>, one of the most extensive studies examining rupture risk of unruptured asymptomatic intracerebral aneurysms, examined especially patients with no previous history of subarachnoid hemorrhage that location and size are the major key points to consider in terms of rupture risk. A majority of ruptured aneurysm shows diameter less than 7 mm, coming up the theory that faster the aneurysm grows, soon will be ruptured. Recently, Liu and coworkers<sup>9</sup> monitoring unruptured intracranial aneurysms based on a volumetric analysis found that a group of the aneurysm can present with accelerated growth rate mainly depending on the location, suggesting that linear measurements could be fallible in predict a warning aneurysm behavior.

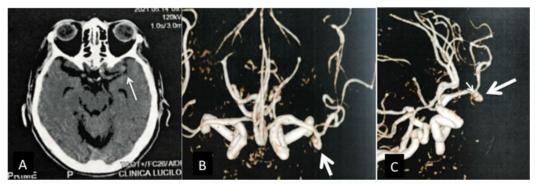


Figure 1. (A) Brain axial CT scan revealed a saccular hyperdensity on the distal left middle cerebral artery (MCA) (arrow); (B) 3D CE-MRA reconstruction on frontal and (C) oblique views showing a 7.1 mm lobulated saccular aneurysm (arrows) on the bifurcation of the MCA. Lobulation on the aneurismal wall is seen.



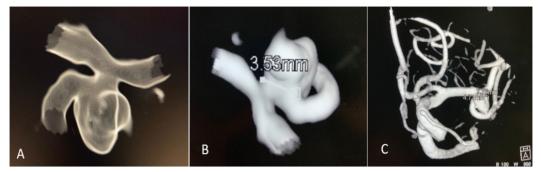


Figure 2. Lobulation is observed in the aneurismal wall.

Age is another crucial point to be taken into consideration for a decision-making process. Although increasing age after 50 does not substantially impact the risk of rupture, preventive surgical intervention brings an additional benefit to morbidity and lethality rates, regardless of whether one chooses to endovascular or open surgery.<sup>15</sup>

In Table 1 are listed meta-analysis studies that evaluated different conditions that may modify the risk of rupture of a still unruptured intracranial aneurysm.

Table 1. Published meta-analyses on risk factors for aneurysm growth

Risk factor for aneurysm growth	Studies
Smoking	Jin et al. <sup>25</sup> ; Backes et al. <sup>26</sup> ; Wermer et al. <sup>27</sup>
Age >70-years-old	Brinjikji <i>et al.</i> <sup>28</sup>
Arterial hypertension	Backes et al. <sup>26</sup> ; Wermer et al. <sup>27</sup>
Woman	Jin et al. <sup>25</sup> ; Backes et al. <sup>24</sup> ; Brinjikji et al. <sup>28</sup> ; Wermer et al. <sup>27</sup>
Aneurysm size >7 mm	Brinjikji <i>et al.</i> <sup>28</sup> ; Wermer <i>et al.</i> <sup>27</sup>
Middle cerebral artery aneurysm	Brinjikji <i>et al.</i> 28; Wermer <i>et al.</i> 27

Using longitudinal analysis of the aneurysm volume, Liu and coworkers<sup>29</sup> demonstrated in a series of 112 aneurysms in 95 patients (the mean follow-up time was 4.0 years.) with UIA that there was evident growth at one year of follow-up in 8% of cases. 36/112 (32%) of the aneurysms exhibited noticeable growth during the study, and 11/36 (31%) aneurysms with noticeable change had an episode of abrupt growth (only 11 of the 112 aneurysms (9.8%) presented an episode of sudden growth).<sup>29</sup> The relative growth rate was dependent on age and the location of the aneurysm, with aneurysms of the anterior cerebral artery showing the fastest growth rate of 4.07% per year. In this study, aneurysms of the middle cerebral artery presented an annual growth rate of 1.41%.<sup>29</sup>

A recent study<sup>11</sup> evaluated the role of statins in unruptured intracranial aneurysm growth and rupture, observing whether atorvastatin (20 mg daily) was associated with aneurysm growth or rupture in patients with less than 7 mm aneurysms. Among the 1.087 enrolled patients, 489 (45%) took atorvastatin (mean follow-up duration of  $33.0 \pm 12.5$ months). Current smoker and uncontrolled hypertension were associated with aneurysm rupture, whereas atorvastatin use was not. The authors concluded that unruptured aneurysms sized 5 to <7 mm and uncontrolled hypertension both were associated with a high growth rate, whereas atorvastatin was associated with a reduced growth rate.<sup>11</sup>

In another study<sup>30</sup>, chosen aneurysm locations (i.e., anterior communicating artery, posterior communicating artery, and middle cerebral artery) and an aneurysm size of 5 to <7 mm were associated with a high risk of aneurysm growth, whereas aspirin and well-controlled blood pressure were associated with a low risk of growth. However, in high-risk patients (>1 risk factor), the cumulative annual growth rates were as high as 40.0 and 53.3 per 100 person-years.<sup>30</sup>

It is worth remembering the headaches associated with aneurysmatic rupture with subarachnoid hemorrhage due to the risk of death due to the lack of an accurate and immediate diagnosis. Headache associate with aneurysm rupture or aneurysm expansion presents as a sudden-onset, high-intensity explosive thunderclap headache, usually occurring during physical exertion. Intracranial aneurysms can simulate a primary headache as well, such as cluster headache. In this context, it is recalled that every patient with cluster headache should be investigated by imaging, including an MRA and cerebral digital subtraction angiography.<sup>31</sup>

Regarding the choice of a conservative management of UIA, results from two studies support this decision. Lawson and colleagues<sup>32</sup> built a methodological model to compare the probability of poor outcomes from intervention (clipping and endovascular) compared to the natural history of the aneurysm, as a function of age and aneurysm size, using data from previous studies. Poor outcomes included discharge to a skilled nursing facility, long-term care facility,



hospice, or in-hospital death. They demonstrated a poor outcome for both clipping or coiling beyond the age of 80 years. Considering the patient's characteristics reported here, an advantage in intervention chose is observed in only one of the four comparisons made, in which endovascular treatment would show benefit up to 86 years old.

Additionally, at the PHASES score study<sup>13</sup>, many of patients with an unruptured intracranial aneurysm derived from six extensive cohort studies, and information was included in a pooled database. They built a model indicating the probability of rupture based on well-known risk factors for aneurysm rupture. The PHASES study incorporated in the evaluation other risk factors, such as age, the maximal diameter and the previous history of subarachnoid hemorrhage. Considering the PHASES score, we calculate a risk of rupture of 2.4% per year (95% Cl; 1.6-3.3), considering the sum total of 7 points from a population, hypertension, age, aneurysm size and site, and absence of a history of SAH. Both studies include the patient's history of hypertension and the first includes the history of smoking. As for limitations, the PHASES score considers populations that do not belong to the reality of the case, making the risk of rupture with the application of the score an estimate.

We conclude by commenting on the significant chance to encounter an asymptomatic intracranial lesion during a patient's headache investigation.<sup>7</sup> Thus, the physician must decide how to treat this incidental finding, which may increase the risk of death.

#### Conflict of interests: No

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