



Review Article

Definitions of intracranial aneurysm size and morphology: A call for standardization

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Received : 09 June 2021

Accepted : 13 August 2021

Published : 06 October 2021

DOI

10.25259/SNI_576_2021

Quick Response Code:



ABSTRACT

Background: Intracranial aneurysms (IAs) are classified based on size (maximal dome diameter) as well as additional parameters such as neck diameter and dome-to-neck ratio (DNR). The neurosurgical literature includes a wide variety of definitions for both IA size and neck classifications. Standardizing the definitions of IA size and wide-neck classifications would help eliminate inconsistencies and potential misunderstandings of aneurysm morphology and rupture risk.

Methods: We queried the MEDLINE (EBSCO) database using the terms “unruptured IA” and (“small” or “medium” or “large”) and filtered based on publication date, language, and scholarly journals. The resulting articles and their references were further screened for eligibility. This identified 286 records, of which 104 were excluded, leaving 182 articles for analysis. The review found several different IA size classifications and neck classifications.

Results: A review of the existing literature describing size and neck classifications revealed 13 size classifications for small aneurysms, four classifications for medium aneurysms, 15 classifications for large aneurysms, and one classification for giant aneurysms. There were also seven different wide-neck classifications found.

Conclusion: It is imperative that a standardization in classification be implemented to help interventionalists make the most informed decisions regarding emerging treatment options as new endovascular technologies and devices are emerging with indications based around these classifications. Based on the database findings, this article recommends standardized quantitative measurement ranges for IA size and neck classifications.

Keywords: Intracranial aneurysm, Neurovascular anatomy, Aneurysm size, Aneurysm morphology

INTRODUCTION

Recent advances in endovascular treatment options for intracranial aneurysms (IAs) have brought increased attention to the various morphological characteristics of IAs and how these characteristics may affect endovascular device selection and the safety and efficacy of treatment. IA classification has typically been based on maximal dome diameter (size) in combination with other parameters such as maximal dome diameter to neck diameter ratio (DNR) and neck diameter [Figure 1].

A wide variety of definitions for both IA size and neck type have been reported, but the optimal definitions for IA size remain unclear. Existing definitions for IA size have been based around

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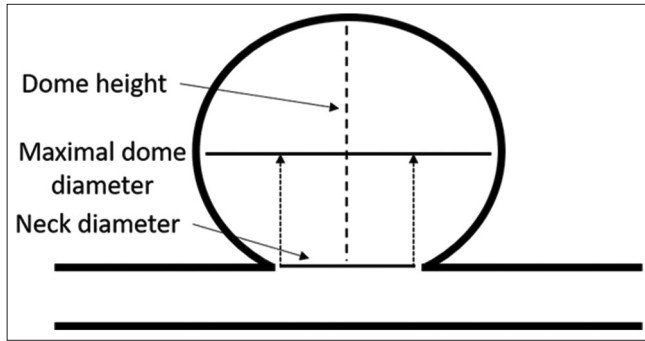


Figure 1: Depiction of relevant intracranial aneurysms dimensions for the calculation of size, neck width, and dome to neck ratio.

the large natural history of IA studies performed, including in the International Study of Unruptured IAs (ISUIA) and The Natural Course of Unruptured Cerebral Aneurysms in a Japanese Cohort (UCAS) study. These studies analyzed varying factors that contribute to IA growth and rupture, to better understand the progression of the disease. A major finding was that the size of the IA correlated to rupture risk and attempted to stratify rupture risk based on IA diameter. The first ISUIA was published in 1998, which was a retrospective study, initially classified aneurysms by the following size definitions based on risk of rupture: small (<10 mm), large (10 mm–25 mm), and giant (>25 mm). The second ISUIA was published in 2003, which was a prospective study, further stratified the aneurysm sizes by including a 4th classification based on risk of rupture: small (<7 mm), medium (7 mm–12 mm), large (12 mm–25 mm), and giant (>25 mm). However, clinicians and researchers in Japan and Finland noticed that they were observing ruptures in small aneurysms much more frequently than what was reported by the ISUIA. To better understand this disparity, the UCAS study was performed and published in 2012. UCAS stratified aneurysms by size based upon rates of rupture: small (<5 mm), medium (5 mm–10 mm), large (10 mm–25 mm), and giant (>25 mm).

As additional research articles and case studies have been published, there has not been a uniform adoption of any single size classification definition system. These inconsistencies in definitions can be a major limitation, as current FDA indications for use include specific IA size and neck size classifications, such as in the expanded small/medium aneurysm indication for Medtronic's Pipeline Embolization Device.^[6] The purpose of this literature review is to highlight the inconsistencies in existing definitions for IA size to highlight the need for standardization of classification schemes for IA neck sizes.

METHODS

This literature review was performed by identifying applicable articles and screening them for eligibility [Figure 2].

A Boolean search of the MEDLINE (EBSCO) database using the terms “unruptured IA” and (“small” or “medium” or “large”) retrieved a total of 100 articles. These articles were then filtered using a publication date between January 1, 1998 and December 31, 2020, an English-language filter, and a scholarly (peer reviewed) journal filter, yielding 81 articles. Of the 81 articles, full-text was found for 65 articles. The total screening group of 65 articles underwent full-text review by searching the PDF file by keyword search for “small”, “medium”, “large”, and “giant”. In addition, the reference lists of the 65 full-text articles were reviewed, and 205 additional articles, including FDA meeting executive summaries and informational webpages, were identified. Screening these references resulted in an additional 117 studies that qualified for full-text evaluation. During the full-text analysis, we also noted any mention of “wide-neck” aneurysm classification and documented if the article defined the term or not. A total of 182 articles underwent full-text evaluation. This literature review uses simple tallying methodology to numerate the number of articles that define/quantify an IA size classification and is not intended to serve as a meta-analysis nor be subjected to statistical analysis.

RESULTS

The reviewed articles consistently classified IAs as “small”, “medium”, “large”, or “giant”. However, the authors did not consistently describe how the measurements were performed (i.e., maximal sac diameter, and midline dome diameter) and measurement ranges for each classification were inconsistent, affecting the ability to compare classifications between articles.

The review identified several different size classifications for IAs. There were 13 quantitative size classification definitions (i.e., 0mm – 5 mm diameter) for “small” aneurysms, 4 classification definitions for “medium” aneurysms, 15 classification definitions for “large” aneurysms, and one classification for giant aneurysms. There were seven wide-neck classification definitions and one very wide-neck classification definition. Articles that mentioned an IA size classification but failed to define/quantify this classification were also noted. The differing classification definitions are listed in [Table 1a-c]. [Table 1a] lists the classification definitions for small – medium IAs, [Table 1b] lists the lists the classification definitions for large – giant IAs, and [Table 1c] lists the classification definitions for wide-neck – very wide-neck IAs.

In total, small IAs were mentioned in 135 articles (74.2%), medium in 20 articles (10.9%), large in 113 articles (62.1%), and giant in 77 articles (42.3%). Wide-neck IAs were mentioned in 64 articles (35.2%), and very wide-neck IAs in two articles (1.1%). Giant IAs were defined the most consistently with all definitions stating IAs >25 mm in maximal diameter.

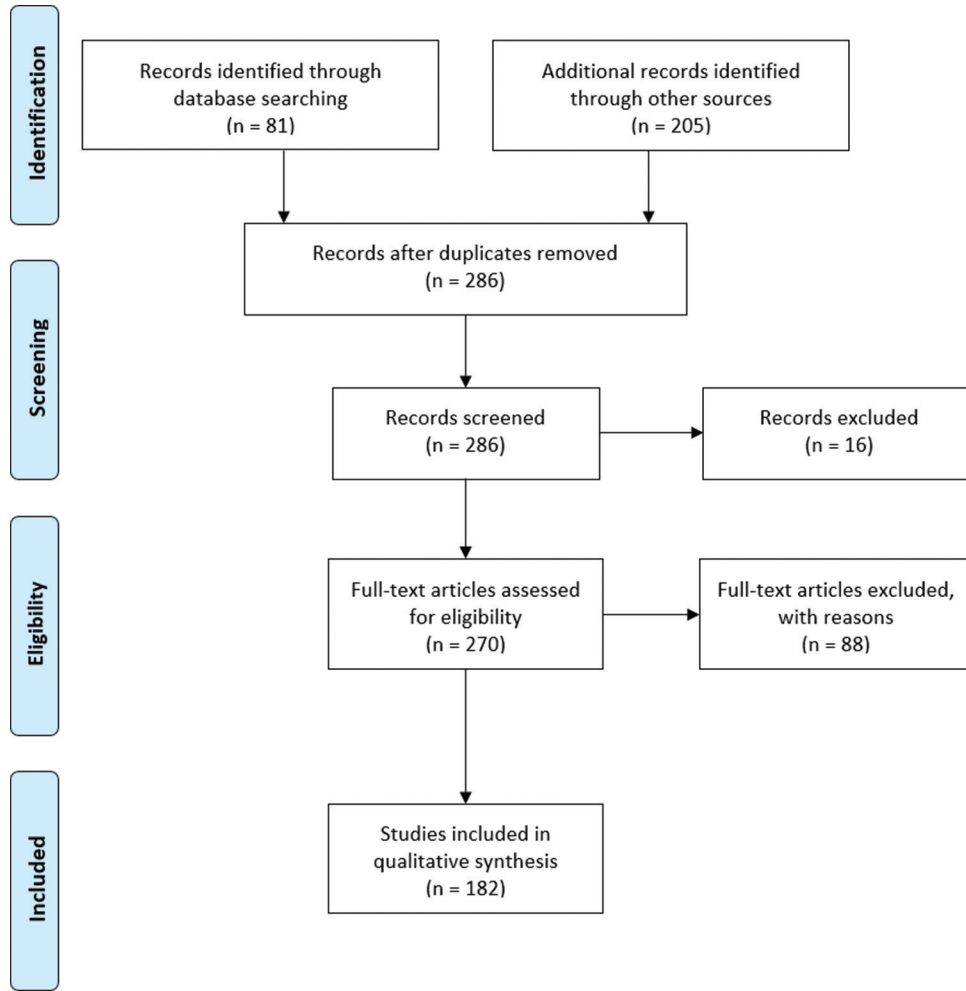


Figure 2: Preferred reporting items for systematic reviews and meta-analysis flow diagram depicting the process of the literature review.

Small IA dimensions, ranging from 0–5 mm to 0–10 mm, were mentioned in 56% of the references. In 31% of the references, no dimension was associated with the small classification [Table 1a]. For medium IAs, 60% of the referenced dimensions either overlapped or were not differentiated from the small classification [Table 1a].

Large IA dimensions, ranging from > 10 mm to 10–25 mm, were mentioned in 21% of the references. In 51% of the references, no dimension was associated with the large classification [Table 1b]. For giant IA dimensions, >25 mm was mentioned in 48% of the references. In 42% of the references, no dimension was associated with the giant classification [Table 1b].

Wide-neck IA dimensions, defining a neck width >4 mm, was mentioned in 44% of the references. DNR values were highly variable, but a wide-neck associated with a DNR <2 was most prevalent (30% of the references). In 47% of the references, no dimension was associated with the wide-neck classification [Table 1c].

DISCUSSION

Existing classifications for IA size have been derived from large natural history studies of unruptured IAs and are derived from risk of rupture. The results from the first ISUIA study (retrospective group) in 1998, the second ISUIA trial (prospective group) in 2003, and The Natural Course of UCAS study in 2012 largely influenced current definitions for IA size classification.^[8,13,14] To take this into consideration, data were also analyzed with respect to publication date: 1998–2002, 2003–2012, and 2012–present [Table 1], to highlight the evolution of IA size classifications as new definitions were introduced by the ISUIA and UCAS.

As new morphology definitions were adopted from these studies, this has led to more variability as many articles continue to use definitions from prior studies. This lack of consistency in size definition is an issue considering that many articles mention IA size classifications without actually defining them (in total, 91.2% of size classifications were not defined) [Figure 3].

Table 1a: IA small - medium size classification definitions, as reported in 155 articles with respect to date published.

Size classification	Dimensions (mm)	After ISUIA 1	After ISUIA 2	After UCAS	Total
		(1998–2002)	(2003–2011)	(2012–2020)	
Small	0–3	1	0	0	1
	0–4	0	0	3	3
	0–5	1	5	14	20
	0–7	0	10	18	28
	0–10	4	5	18	27
	0–11	0	0	1	1
	0–12	0	2	0	2
	0–15	0	0	2	2
	10	0	0	0	0
	2–7	0	0	1	1
	3–5	1	0	0	1
	5–7	0	0	1	1
	5–10	0	0	1	1
	4–10	0	1	0	1
	Combined with medium	0	0	4	4
	Mentioned, not defined	3	13	26	42
Overall Total: 135					
Medium	7–12	0	3	2	5
	5–10	0	1	2	3
	7–10	0	1	1	2
	7–15	0	0	1	1
	Combined with small	0	0	4	4
	Combined with large	0	0	1	1
	Mentioned, not defined	0	1	3	4
	Totals	0	6	14	
Overall Total: 20					

IA: Intracranial aneurysms, ISUIA: International Study of Unruptured intracranial aneurysms, UCAS: Unruptured Cerebral Aneurysms in a Japanese Cohort

It is also important to note the differences between maximal dome diameter and height. Maximum dome diameter is properly defined as the widest IA measurement perpendicular to the aneurysm height.^[10] Aneurysm height has been shown to potentially have independent influence on the treatment outcomes and should be thus differentiated from dome diameter.^[2]

Small IAs

Small IAs are mentioned the most in the reviewed literature. This is likely due to the extensive research performed on small IAs and the implication of their size on risk of rupture.^[8,13,14] Both Japanese and Finnish studies have shown that aneurysms <5 mm in size still pose a significant risk for rupture.^[3]

Medium IAs

Medium IAs are mentioned the least in the reviewed literature. They are often included in either small or large IA

classifications (*i.e.*, “small and medium aneurysms [<10 mm in diameter]”), or are not defined at all. When medium aneurysms were included in small or large IA classifications, it was tallied as “combined.” There were several articles that had a data range for what could have been classified as a medium IA, but the medium classification was not mentioned (surrounding size data ranges were classified as small and large, but no mention of medium IA was made). The medium IA classification was not used in the first ISUIA study but has been adopted in later studies.^[8,14] A distinct medium IA size classification is critical because this has now been used in endovascular device FDA indications for use.^[6]

Large IAs

Large IAs were mentioned but left undefined 51% of the time, more than any other classification in the reviewed literature. In many references that focused on small IAs, the large classification was used as a catch-all for anything that fell outside the small IA classification.

Table 1b: IA large - giant size classification definitions, as reported in 190 articles with respect to date published.

Size classification	Dimensions (mm)	After ISUIA 1	After ISUIA 2	After UCAS	Total
		(1998–2002)	(2003–2011)	(2012–2020)	
Large	>4	0	0	2	2
	>5	0	1	0	1
	>7	0	0	1	1
	>10	1	0	8	9
	>12	0	0	1	1
	6–13	1	0	0	1
	7–25	0	0	1	1
	10–25	1	4	10	15
	10–24	0	1	0	1
	11–25 mm	0	1	2	3
	12–24 mm	0	1	0	1
	12–25 mm	0	4	2	6
	13–24 mm	0	0	1	1
	15–24 mm	0	0	1	1
	15–25 mm	0	1	2	3
	Combined with medium	0	0	1	1
	Combined with giant	0	0	7	7
	Mentioned, not defined	7	16	35	58
	Overall Total: 113				
Giant	>25 mm	4	11	22	37
	Combined with large	0	1	7	8
	Mentioned, not defined	2	9	21	32
	Overall Total: 77				

IA: Intracranial aneurysms, ISUIA: International Study of Unruptured intracranial aneurysms, UCAS: Unruptured Cerebral Aneurysms in a Japanese Cohort

Table 1c: IA wide-neck – very wide-neck size classification definitions, as reported in 66 articles with respect to date published.

Size classification	Dimensions (mm)	After ISUIA 1	After ISUIA 2	After UCAS	Total
		(1998–2002)	(2003–2011)	(2012–2020)	
Wide-Neck	>4 mm	0	5	7	12
	DNR <2	1	0	5	6
	DNR <1.6	0	1	0	1
	DNR <1.5	0	1	0	1
	>4 mm or DNR >0.7	0	1	0	1
	>4 mm and DNR <2	0	2	7	9
	>4 mm and DNR <1.5	0	3	1	4
	>4 mm and DNR <1.2	0	0	0	0
	>4 mm and DNR <1	0	0	0	0
	Mentioned, not defined	0	8	22	30
	Overall Total: 64				
	Very Wide-Neck	DNR <1.2	0	1	0
Mentioned, not defined		0	1	0	1
Overall Total: 2					

IA: Intracranial aneurysms, ISUIA: International Study of Unruptured intracranial aneurysms, UCAS: Unruptured Cerebral Aneurysms in a Japanese Cohort

Giant IAs

Giant IAs were classified most consistently in the reviewed literature. This could be because the giant IA classification is one of the oldest, dating back to 1969.^[1,9]

Wide-neck IAs

Wide-neck IAs were first defined in 1994 as an IA with a neck ≥ 4 mm.^[15] As endovascular treatment became more prevalent, the definition expanded to include DNR as a predictor

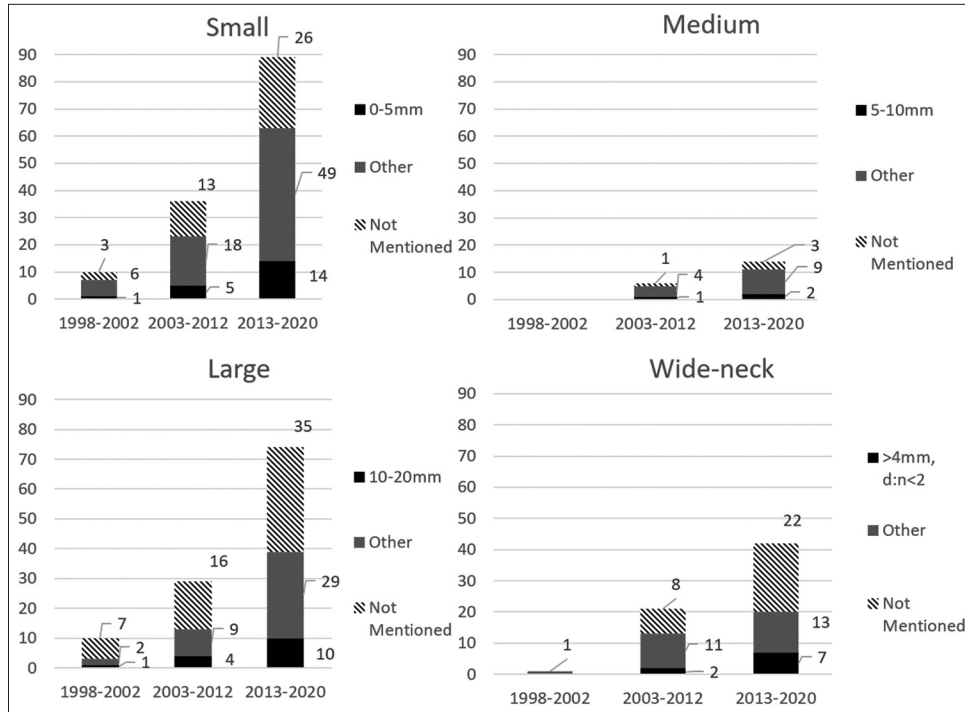


Figure 3: Graphical representation of trends in intracranial aneurysms classification definitions.

of coiling success.^[4] Neck size and DNR remain critical characteristics in determining the success of endovascular IA treatment, particularly with coil embolization. IAs with “wide”, “broad”, or “large” necks often require advanced endovascular techniques, and have implications for IA recurrence, rupture, or other para- and post-operational complications. As technology and endovascular techniques have evolved, some of the neurosurgical literature has pushed for the DNR definition of wide-neck IAs to be decreased from <2 for the definition to be more representative of IAs that are more likely to require adjunctive treatment measures.^[2]

Recently, Hendricks *et al.* (2019) performed a focused systematic review of the neurosurgical literature directed at wide-neck IA definition and clinical implication. This review also found a lack of consistency and identified seven unique definitions for wide-neck aneurysms. The most commonly found definition for wide-neck IA was a neck diameter >4 mm or DNR < 2.^[7]

Taking into account the evolution of the IA size and IA neck classification trends in the literature, and the fact that unclassified aneurysms are still prevalent in the literature, we recommend the following standardized classification for aneurysm geometry: maximal dome diameter classifications, and wide neck classification [Table 2]. This classification system generally agrees with the Japanese UCAS study findings that correlated aneurysm size to risk of rupture that included higher-risk demographics for aneurysm rupture.^[8]

Table 2: Proposed classification definitions for IA size (dome width) and neck type.

Small	Medium	Large	Giant	Wide-neck
<5 mm	>5 mm–10 mm	>10 mm–25 mm	>25 mm	Neck >4 mm and/or DNR <2

IA: Intracranial aneurysms

A major limitation of the ISUIA studies and their resulting size classification systems was that the cohorts evaluated were over 90% Caucasian and did not sufficiently include demographics that demonstrate higher rates of aneurysm rupture such as Japanese and Finnish patients.^[8]

Aspect ratio, defined as the ratio between IA height and neck diameter, has also been shown to have clinical implications on both treatment outcome and risk of rupture. This ratio has not been studied as extensively but may represent an additional important morphological parameter for predicting treatment outcomes of IAs.^[2,5,11,12]

Limitations

This literature review was not intended to be a comprehensive analysis of every paper that has classified an aneurysm by size, but rather a structured sampling of the literature to highlight the inconsistencies and trends in the definitions for IA size classification.

CONCLUSION

There is tremendous variability in the existing definitions for IA size and neck classification. Flexibility in size definition was more acceptable when it mainly related to statistical stratification of IA rupture risk in studies. With new endovascular technologies emerging with indications based around these size classifications, it is imperative that standardization in classification definition is implemented. Standardizing the definitions of IA size and neck classifications would help eliminate inconsistencies and potential misunderstandings. We hope that this review spurs further discussion that results in a more consistent, standardized intracranial classification system that allows surgeons, regulatory bodies, and medical device developers to quickly identify suitable treatment devices for patient-specific IA dimensions and morphologies.

The general trend can be seen in IA size classification moving towards the following definitions: small <5 mm, medium >5 mm-10 mm, large >10 mm-25 mm, and giant >25 mm, which follows the Japanese UCAS study size classifications.^[3] With the current natural history of IA data available, this size classification system represents the most inclusive of higher-risk demographics and thus is recommended as the most appropriate size classification to adopt. The data also suggest that definition of a wide-neck aneurysm be an IA with a neck diameter >4 mm or DNR <2 be adopted as a standard until data can support reducing the DNR to a lower value. Whether, these definitions comprise the most appropriate system for IA size classification is yet to be determined and is a subject for future discussion and research.

Declaration of patient consent

Patient's consent not required as patients identity is not disclosed or compromised.

Financial support and sponsorship

Nil.

Conflicts of interest

There are no conflicts of interest.

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How to cite this article: Merritt WC, Berns HF, Ducruet AF, Becker TA. Definitions of intracranial aneurysm size and morphology: A call for standardization. *Surg Neurol Int* 2021;12:506.