

# Vascular Ultrasound Statement from the Department of Cardiovascular Imaging of the Brazilian Society of Cardiology – 2019

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**Note:** These Guidelines are for information purposes and are not to replace the clinical judgment of a physician, who must ultimately determine the appropriate treatment for each patient.

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#### 1. Introduction and equipment

#### 1.1. Introduction

The use of ultrasound (US) in Medicine started in the decade of 1940 and, since then, it has played a significant role in the diagnosis of cardiovascular diseases (CVD). It is a diverse technology that follows the physical principles of propagation of US waves in different media. Due to its broad applicability, relatively low-cost, and reproducibility, US has a secure place in diagnostic aid. Members of the Department of Cardiovascular Imaging (DCI) - experts in vascular ultrasound (VUS) - prepared this guideline to indicate the best use of this technique, according to recommendations from the current medical literature. We included the main aspects related to equipment components, software, transducers, and their evolution since the last guideline, and addressed the most relevant topics in the field of diagnosis of vascular diseases. The foundation behind using VUS to diagnose important pathologies, such as carotid atherosclerotic disease, diseases in vertebral arteries, abdominal aorta and its branches, was

based on recommendations from the DCI panel of experts in 2015 and 2016.  $^{\!\!\!1,2}$ 

This guideline does not aim to compare VUS with other imaging examination methods or expound on the use of VUS in the follow-up of vascular diseases after the initial diagnosis. For this content, the reader should consult more extensive and specific publications on the subject.

Our goal is to disseminate the best practices in VUS to various services in the country, standardize the interpretation of examinations, and contribute to the proper use of this noninvasive, widely available, and low-cost tool.

#### 1.2. Equipment

In a country with continental dimensions and different economic realities like Brazil, it is difficult to determine what ideal equipment is. We cannot demand from a small laboratory in the interior of the country to work with equipment that has the same technological resources needed for a laboratory that assists a large number of patients. This standardization intends to suggest the appropriate minimum resources of equipment and the ideal way to perform, with safety and accuracy, the examinations whose protocols will be described below, always remembering that this area is in constant evolution.

#### 1.2.1. Machine

The equipment must be capable of delivering the following types of image and Doppler: (a) two-dimensional image; (b) color flow imaging (CFI); (c) pulsed wave Doppler; (d) continuous wave Doppler (for some types of transducers, but not required for vascular examinations); (e) power Doppler, also known as power angio and Doppler energy – way of mapping the flow without indicating the direction, based on the amplitude of the signal (ideal resource, but not essential to the examination).

More advanced tools, such as second-harmonic imaging, B-Flow, inversion recovery pulse sequences – for the use of microbubble contrast –, and even transducers capable of producing three-dimensional images, are useful in complex examinations, but not yet part of our daily practice. They will also be covered for information purposes as a way to encourage the technological progress that brings additional benefits to patients.

#### 1.2.2. Applications (Software)

Among the application options, the equipment must have specific presets to each type of study to expedite and facilitate the task of the examiner.

#### 1.2.3. Multi-frequency or broadband transducers

• Linear transducer: ideal frequency between 5 and 10 MHz (in some cases, frequencies of 4 or 12 MHz can be useful); for studies of superficial structures, since transducers of higher frequency have better axial image resolution, but their use is limited due to the large sound damping when traveling through tissues.

• Convex transducer: ideal frequency between 2 and 5 MHz; used in studies of deeper structures, such as abdominal

ones, with the advantage of covering a larger area compared to sector transducers of similar frequencies.

• Low-frequency sector transducer: 2 to 4 MHz; useful when the examiner needs continuous Doppler in studies of abdominal arteries.

• **High-frequency sector transducer**: 4 to 10 MHz; useful when the acoustic window is limited by bone structures.

• Micro-convex transducer: frequency between 4 and 8 MHz; adaptable to sites with limited window, such as bone structures, dressings, wounds, or other situations in which the contact surface available for the probe is reduced, without loss of lateral resolution in distal fields, as presented by sector transducers.

#### 1.2.3.1. Image orientation

In longitudinal images, most vascular imaging guides recommend displaying cranial structures on the left side of the screen, and caudal structures on the right. In transverse planes, structures on the left side of the screen must correspond to the marking on the upper left corner of the monitor. That way, transverse planes will display right lateral structures, as well as left medial structures on the left side of the monitor screen.<sup>3,4</sup>

#### 2. Carotid and vertebral arteries

According to the World Health Organization (WHO), CVDs are the main causes of morbidity and mortality worldwide. In 2012, 17.5 million people died from CVDs, the equivalent to 31% of all deaths occurred in the period, with estimates that 7.4 million resulted from coronary artery disease (CAD) and 6.7 million from cerebrovascular accident (CVA).<sup>5</sup>

Ultrasound of carotid arteries is valuable and widely used in cardiovascular risk assessment, as it measures the intima-media thickness (IMT), detects atherosclerotic plaques, and can evaluate the morphology of plaques and degree of stenosis, characteristics associated with cerebrovascular events.

# 2.1. Intima-media thickness and detection of carotid artery plaques for cardiovascular risk assessment

With the publication of the Brazilian Guidelines for Dyslipidemia and Atherosclerosis Prevention in 2007 and 2013,<sup>1,6-8</sup> the Mannheim consensus documents of 2004-2011,<sup>9</sup> and the American Society of Echocardiography consensus,<sup>10</sup> Brazilian experts in the VUS field joined forces to disseminate the correct way to measure IMT and detect atherosclerotic plaques in carotid arteries. In the latest update of the Brazilian guideline in 2017,8 IMT measurement was not included separately in the stratification of cardiovascular risk, but in the characterization of atherosclerotic plaque as IMT > 1.5mm. Another aspect that shows the importance of correctly measuring IMT is its use in several research protocols. Since the American and European expert consensuses use IMT as an aggravating factor for cardiovascular risk, we decided to include the measurement technique in this guideline. This section aims to standardize the technique to measure IMT and detect carotid plaques.

# 2.2. Ultrasound definition of intima-media thickness and carotid plaque

in two-dimensional images, IMT is characterized by a double line with defined intima-lumen and media-adventitia interfaces. IMT is the distance between the two acoustic interfaces. The atheromatous carotid plaque (CP) can be defined as a focal structure that spreads at least 0.5 mm into the arterial lumen, and/or measures more than 50% of the surrounding IMT value, and/or has an IMT > 1.5 mm.<sup>9</sup> Figure 1 schematically illustrates the IMT measurement and the three ways to define CP.

#### 2.2.1. Indications

The European and American consensuses<sup>9,10</sup> recommend measuring IMT in specific groups (Table 1). In these groups of individuals, IMT is considered increased when above the 75 percentile for their age, gender, and ethnicity, according to one of the normative tables, assisting in the discussion of clinical treatment and change in lifestyle. If a CP is found, regardless of the obstruction degree, IMT measurement does not need to be reported, except for exams explicitly requested for this purpose. In these cases, if the CP is located in the IMT measurement, it should be included in the value.

#### 2.2.2. Measurement protocol

The recommended protocol is similar to the one described by the ELSA-Brasil study<sup>13</sup> (Table 2).

After collecting numerical IMT data, the average values will be compared with existing reference numbers, according to normative tables of the studies ELSA Brasil,<sup>13</sup> CAPS,<sup>14</sup> or

MESA.<sup>15</sup> The decision about which table to use will depend on the gender, age, and ethnicity of the individual.

Although the manual point-to-point measurement is less reproducible – considering the differences in ultrasound equipment used in our country –, the consensus was that it could be used if the equipment does not automatically measure IMT, strictly respecting the technical recommendations. The examiner should pay special attention when placing the cursor in the intima-lumen and media-adventitia interfaces and be very cautious not to overestimate the values. In this case, measure each side at least five times to obtain the mean value (mean IMT).<sup>16</sup>

#### 2.2.3. Interpretation of results

Mean IMT [mean of measurements in the right or left common carotid artery (RCCA and LCCA, respectively)]: most types of equipment provide this measurement automatically or semi-automatically (in the latter case, the examiner can make small adjustments based on the automatic one). After calculating the mean IMT values from each side, compare them to the table to find the equivalent percentile. The individual under study is classified in the highest percentile.

To determine the percentile of IMT measurement, consult tables 3 to 5, according to the age group of the individual under study.

#### 2.2.4. How to prepare a report

Describe the mean IMT on each side, in mm, in the body of the report. Also, include in conclusion if the value is



Figure 1 – Schematic illustration showing examples of IMT and plaque measurements. IMT measurement (A). Different measurements of 3 carotid plaques: encroaching ≥ 0.5 mm on the arterial lumen (B); measurement > 50% of the surrounding IMT value (C); large plaque (D).

#### Table 1 – When to measure the intima-media thickness.

1. Intermediate cardiovascular risk: use the IMT measurement as an aggravating factor for high risk reclassification<sup>10</sup>

2. Patients known to have a higher cardiovascular risk and hard clinical classification:

· Patients with familial hypercholesterolemia11

Patients with autoimmune diseases or who use immunosuppressants, corticosteroids, antiretroviral drugs, or other medicines that induce elevation of cholesterol<sup>12</sup>

· History of early cardiovascular disease in first-degree relatives<sup>10</sup>

Individuals < 60 years with a severe abnormality in a risk factor<sup>10</sup>

Women < 60 years with at least two risk factors<sup>10</sup>

IMT: intima-media thickness.

#### Table 2 – Protocol to measure intima-media thickness.

· Two-dimensional fundamental imaging

- Do not zoom
- Transducer with frequency > 7 MHz
- · Proper gain adjustment; depth between 3.0 and 4.0 cm
- · Longitudinal plane of the common carotid and carotid bifurcation

• Capture images in the anterior and posterior accesses or the sternocleidomastoid muscle, with the most rectilinear image possible and with a well-defined double-line pattern, and choose the best one

· Measure it in the posterior wall of the common carotids on the right and left sides, 1 cm from the bifurcation, in automatic/semi-automatic mode

above or below the 75 percentile and the table used, with its bibliographic reference.

Inform the presence of carotid plaques, with their specific characteristics and quantification, according to the criteria recommended by the recent Brazilian consensus.<sup>1</sup>

# 2.3. Morphological evaluation of carotid atherosclerotic plaques

CP morphology plays an essential role in the incidence of cerebrovascular events and can also be an important predictor of events.<sup>17,18</sup>

Recognizing the ultrasound characteristics of the plaque can help to identify unstable ones. Describe the following properties: location, extension, echogenicity, texture, surface, presence of movable components, and anechoic areas next to the fibrotic capsule. Report these characteristics for the most important plaques, particularly those with more than 50% stenosis.

• Location: we recommend subdividing the carotids into distal and proximal common carotid, bifurcation, external branch, and proximal and medial internal branch (Figure 2).

• **Extension**: must be measured, as it can be correlated with events and affects the choice of surgical and endovascular treatment.<sup>19</sup>

• **Echogenicity**: defined by comparing the plaque echogenicity to that of adjacent structures (blood, muscle, adventitia of the vessel, and bone), and classified into:<sup>20</sup>

- Hypoechoic or echolucent: darker, that is, echogenicity similar to that of blood and less echogenic than the sternocleidomastoid muscle.

- Isoechoic: echogenicity close to that of muscle.
- Hyperechoic: lighter than the adjacent muscle.

- Calcified: very echogenic, creating acoustic shadowing due to calcium deposition. Echogenicity is comparable to that of the bone.

• Echotexture: Reilly et al.<sup>21</sup> classified the texture of the plaque as homogeneous or heterogeneous.

- Homogeneous: uniform in both low and high echo levels.

- Heterogeneous: a mixture of high, medium, and low echo levels.

 $\bullet$  Surface: lumen surface is categorized into three classes:  $^{\rm 20,22}$ 

- Smooth: irregularities of less than 0.4 mm depth.
- Irregular: from 0.4 to 2 mm depth.
- Ulcerated: crater greater than 2.0 mm depth.

#### 2.4. Quantification of carotid artery stenosis

Several institutions published evaluation criteria for carotid stenosis, with some differences in interpretation.<sup>3,23-25</sup> However, in 2003, a consensus document was published in the USA to make recommendations on the performance of VUS of carotid arteries. The United Kingdom followed them in 2009, and the DCI of the Brazilian Society of Cardiology (BSC) in 2015.<sup>1,3,4</sup>

Меа	n IMT LCCA	(mm)							Меа	n IMT RCCA	(mm)						
	Ethnicity	Percentile	40y	45y	50y	55y	60y	65y		Ethnicity	Percentile	40y	45y	50y	55y	60y	65y
		P 25	0.47	0.49	0.52	0.54	0.57	0.60			P 25	0.45	0.48	0.51	0.53	0.56	0.59
	\A/bite	P 50	0.53	0.57	0.60	0.64	0.67	0.71		\A/bite	IMT RCCA (mm)           Ethnicity         Percentile         40y         45y         50y         55y         60y         6           P25         0.45         0.48         0.51         0.53         0.56         0.           White         P50         0.51         0.54         0.58         0.61         0.65         0.           P90         0.66         0.71         0.76         0.81         0.85         0.           P90         0.66         0.71         0.76         0.81         0.85         0.           P90         0.66         0.71         0.76         0.81         0.85         0.           Multiracial         P50         0.50         0.54         0.58         0.62         0.66         0.           P90         0.69         0.74         0.79         0.84         0.89         0.           P90         0.69         0.74         0.79         0.84         0.89         0.           Black         P50         0.54         0.58         0.62         0.66         0.70         0.           P90         0.70         0.77         0.83         0.89         0.95         1.	0.69					
	vvnite	P 75	0.60	0.65	0.69	0.73	0.77	0.81		vvnite	P 75	0.59	0.63	0.67	0.71	0.75	0.79
		P 90	0.70	0.75	0.80	0.85	0.90	0.95			P 90	0.66	0.71	0.76	0.81	0.85	0.90
		P 25	0.48	0.50	0.53	Mean M1 RCCA (mm)           Main M1 RCCA (mm)           Part M1 RCCA (mm)           Part M1 RCCA (mm)           Main M1 RCCA (mm)           Main M1 RCCA (mm)           Part M1 RCCA (mm)           Main M1 RCCA (mm)           Part M1 RCCA (mm)           Main RCCA (mm)     <	0.56	0.60									
Man	Madding at all	P 50	0.53	0.57	0.61	0.65	0.69	0.73	Man	Marking sight	CCA (mm)           ity         Percentile         40y         45y         50y         55           P 25         0.45         0.48         0.51         0.5           P 50         0.51         0.54         0.58         0.6           P 75         0.59         0.63         0.67         0.7           P 90         0.66         0.71         0.76         0.8           P 25         0.44         0.47         0.50         0.5           cial         P 50         0.50         0.54         0.58         0.6           P 75         0.58         0.63         0.68         0.7           P 90         0.69         0.74         0.79         0.8           P 90         0.69         0.74         0.79         0.8           P 90         0.61         0.67         0.73         0.7           P 90         0.70         0.77         0.83         0.6           P 75         0.61         0.67         0.73         0.7           P 90         0.70         0.77         0.83         0.6           P 90         0.70         0.77         0.83         0.6           P 50	0.62	0.66	0.69			
	wuttracial	P 75	0.60	0.65	0.70	0.75	0.80	0.85		wuttracial	P 75	0.58	0.63	0.68	0.73	0.77	0.82
		P 90	0.69	0.75	0.80	0.86	0.92	0.97			P 90	0.69	0.74	0.79	0.84	0.89	0.94
		P 25	0.49	0.52	0.55	0.58	0.62	0.65			P 25	0.46	0.50	0.53	0.57	0.60	0.64
	Diask	P 50	0.56	0.59	0.63	0.67	0.71	0.75		Diask	P 50	0.54	0.58	0.62	/         55y         60y         65y           1         0.53         0.56         0.59           8         0.61         0.65         0.69           7         0.71         0.75         0.79           6         0.81         0.85         0.90           0         0.53         0.56         0.60           8         0.62         0.66         0.69           9         0.84         0.89         0.94           3         0.57         0.60         0.64           2         0.66         0.70         0.74           3         0.78         0.84         0.90           3         0.78         0.84         0.90           3         0.78         0.84         0.90           3         0.78         0.84         0.90           3         0.78         0.84         0.90           0         0.53         0.55         0.58           6         0.59         0.63         0.66           2         0.66         0.70         0.75           9         0.74         0.79         0.84           0         0.53         0.56	0.74	
	BIACK	P 75	0.64	0.68	0.72	0.77	0.81	1.62     0.65     P 25     0.46     0.50     0.53     0.57     0.57       1.71     0.75     P 50     0.54     0.58     0.62     0.66     0       1.81     0.86     P 75     0.61     0.67     0.73     0.78     0       0.97     1.03     P 90     0.70     0.77     0.83     0.89     0       60y     65y     Etnia     Percentile     40y     45y     50y     55y	0.84	0.90							
		P 90	0.71	0.78	0.84	0.91	0.97	1.03			P 90	0.70	0.77	0.83	0.89	0.95	1.02
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		P 25	0.44	0.47	0.50	0.53	0.56	0.59			P 25	0.44	0.47	0.50	0.53	0.55	0.58
	\A/bite	P 50	0.49	0.52	0.56	0.59	0.63	0.66		\A/bite	P 50	0.48	0.52	0.56	0.59	0.63	0.66
	vvnite	P 75	0.54	0.58	0.63	0.67	0.71	0.75		vvnite	P 75	0.53	0.58	0.62	0.66	0.70	0.75
		P 90	0.61	0.66	0.71	0.76	0.81	0.86			P 90	0.59	0.64	0.69	0.74	0.79	0.84
Ę		P 25	0.45	0.48	0.51	0.54	0.57	0.60	5		P 25	0.44	0.47	0.50	0.53	0.56	0.59
/oma	Multingeigt	P 50	0.50	0.53	0.57	0.60	0.64	0.67	loma	Multinesiel	P 50	0.49	0.52	0.56	0.60	0.64	0.68
S	wuttracial	P 75	0.56	0.60	0.64	0.68	$\begin{array}{c ccccccccccccccccccccccccccccccccccc$	0.72	0.76								
		P 90	0.63	0.68	0.73	0.78	0.83	0.88			P 90	0.62	0.67	0.72	0.77	0.82	0.87
		P 25	0.46	0.49	0.52	0.55	0.58	0.61			P 25	0.46	0.49	0.53	0.56	0.59	0.63
	Diask	P 50	0.51	0.55	0.59	0.63	0.67	0.70		<b>D</b> 1 1	P 50	0.51	0.55	0.59	0.63	0.67	0.71
	BIACK	P 75	0.57	0.62	0.66	0.70	64 $0.67$ $0.71$ White         P 50 $0.51$ $0.54$ $0.71$ 73 $0.77$ $0.81$ White         P 75 $0.59$ $0.63$ $0.65$ 85 $0.90$ $0.95$ P 90 $0.66$ $0.71$ $0.55$ 56 $0.69$ $0.73$ $P 50$ $0.50$ $0.54$ $0.71$ $65$ $0.69$ $0.73$ $P 50$ $0.50$ $0.54$ $0.74$ $0.75$ $75$ $0.80$ $0.85$ $0.62$ $0.65$ $0.54$ $0.50$ $0.54$ $0.57$ $67$ $0.71$ $0.75$ $0.69$ $0.74$ $0.57$ $0.60$ $0.69$ $0.74$ $0.57$ $67$ $0.71$ $0.75$ $0.56$ $0.57$ $0.61$ $0.67$ $0.77$ $0.77$ $0.77$ $0.77$ $0.77$ $0.77$ $0.77$ $0.77$ $0.77$ $0.77$ $0.77$ $0.77$ $0.77$ $0.77$ $0.77$ $0.77$	0.67	0.71	0.76	0.80						
		P 90	0.64	0 70	0.76	0.82	0.88	0 94			P 90	0.64	0.71	0 77	0.83	0.90	0.96

Table 3 – ELSA Brasil:<sup>13</sup> individuals of both genders, aged 40 to 65 years, of white, multiracial, or black ethnicity.

RCCA: right common carotid artery; LCCA: left common carotid artery; IMT: intima-media thickness.

Table 4 – CAPS lable. Individuals of both genuers, aged 25 to 45 years. There is no ethnic classifica	able 4 -	– CAPS table: <sup>14</sup>	individuals of bot	n genders.	aged 25 to 45	years. There is r	to ethnic classificat
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	Densentile		Age							Deveentile	Age						
	Percentile	25	35	45	55	65	75	85	E	Percentile	25	35	45	55	65	75	85
Man	%25	0.515	0.585	0.634	0.68	0.745	0.814	0.83	Voma	%25	0.524	0.575	0.619	0.665	0.718	0.771	0.807
	%50	0.567	0.633	0.686	0.746	0.83	0.914	0.937	>	%50	0.567	0.615	0.665	0.719	0.778	0.837	0.880
	%75	0.633	0.682	0.756	0.837	0.921	1.028	1.208		%75	0.612	0.66	0.713	0.776	0.852	0.921	0.935

#### 2.4.1. Measurement techniques to quantify stenosis

The assessment of carotid stenosis with VUS is based on measurements of flow velocity and their relationships using spectral Doppler, associated with the evaluation of twodimensional and color Doppler imaging. With the patient in the supine position, use transverse and longitudinal ultrasound planes that allow the visualization of the right and left carotid system. Figure 3 shows the recommendation from DCI-BSC for the sequence of evaluation of carotid stenosis.

# 2.4.1.1. Quantification of carotid stenosis with hemodynamic parameters

Arterial flow velocity measured by Doppler is presented as flow velocity waveforms (spectrum) for each site examined.

#### Table 5 – MESA table:<sup>15</sup> individuals of both genders, aged 65 to 84 years, of white, black, Chinese, or Hispanic ethnicity.

							Mean	IMT RCC	A								
		Whit	e man			White	woman			Black	k man			Black	woman		
Percentile		A	ge		Age			Age				A	ge				
	45-54	55-64	65-74	75-84	45-54	55-64	65-74	75-84	45-54	55-64	65-74	75-84	45-54	55-64	65-74	75-84	
%25	0.52	0.57	0.65	0.72	0.51	0.55	0.65	0.72	0.58	0.61	0.71	0.74	0.55	0.60	0.65	0.71	
%50	0.62	0.68	0.77	0.83	0.58	0.65	0.75	0.83	0.67	0.74	0.85	0.85	0.64	0.71	0.76	0.83	
%75	0.71	0.81	0.92	0.97	0.67	0.76	0.87	0.93	0.80	0.92	0.99	1,02	0.74	0.81	0.92	0.96	
		Chine	se man			Chinese	woman			Hispar	nic man			Hispanic woman			
Percentile	rcentile Age				Α	lge Age			Age								
	45-54	55-64	65-74	75-84	45-54	55-64	65-74	75-84	45-54	55-64	65-74	75-84	45-54	55-64	65-74	75-84	
%25	0.54	0.56	0.62	0.66	0.55	0.54	0.59	0.67	0.53	0.60	0.65	0.71	0.51	0.57	0.65	0.63	
%50	0.64	0.70	0.73	0.79	0.60	0.63	0.71	0.77	0.62	0.67	0.78	0.81	0.58	0.69	0.76	0.78	
%75	0.73	0.83	0.92	0.98	0.70	0.77	0.84	0.96	0.73	0.82	0.90	0.92	0.67	0.77	0.87	0.92	
							Mean	IMT LCC	A								
		Whit	e man			White	woman			Black	k man			Black	woman		
Percentile	tile Age					Α	ge			Α	ge			A	ge		
	45-54	55-64	65-74	75-84	45-54	55-64	65-74	75-84	45-54	55-64	65-74	75-84	45-54	55-64	65-74	75-84	
%25	0.54	0.57	0.67	0.71	0.50	0.55	0.63	0.70	0.56	0.63	0.69	0.72	0.54	0.59	0.63	0.68	
%50	0.63	0.69	0.81	0.85	0.58	0.64	0.73	0.80	0.69	0.75	0.82	0.85	0.63	0.67	0.76	0.78	

%75	0.78	0.82	0.95	1,00	0.67	0.75	0.85	0.94	0.81	0.92	0.99	1,02	0.73	0.80	0.90	0.91		
Chinese man					Chinese	e woman			Hispar	nic man			Hispanic	woman	man			
Percentil		A	ge			Α	ge			Α	ge			Ag	ge			
	45-54	55-64	65-74	75-84	45-54	55-64	65-74	75-84	45-54	55-64	65-74	75-84	45-54	55-64	65-74	75-84		
%25	0.55	0.57	0.62	0.69	0.49	0.52	0.58	0.64	0.55	0.61	0.68	0.72	0.51	0.58	0.62	0.68		
%50	0.63	0.70	0.72	0.84	0.58	0.63	0.71	0.76	0.64	0.72	0.80	0.86	0.58	0.68	0.72	0.77		
%75	0.73	0.84	0.86	0.97	0.67	0.72	0.87	0.94	0.75	0.85	0.98	0.97	0.68	0.79	0.86	0.91		

RCCA: right common carotid artery; LCCA: left common carotid artery; IMT: intima-media thickness.

Wave characteristics depend on the type of local flow, as well as changes in blood velocity resulting from conditions proximal and distal to the measurement site. In the internal carotid artery (ICA), the resistance of flow velocity waveforms is usually of low. In the external carotid artery (ECA), the resistance of velocity waveform is high, while in the common carotid artery (CCA), it is intermediate between ICA and ECA<sup>13</sup> (Figure 4).

Measure the velocities in the proximal and distal segments of CCA. To evaluate the velocity ratio in the quantification of ICA stenosis, assess CCA at approximately 2 cm from the bifurcation, preferably in a rectilinear segment, without plaques.<sup>4</sup> Analyze ICA in the proximal and middle segments, since atherosclerotic lesions usually affect the proximal 2 cm. The cursor must be parallel to the vessel wall, with a volume sample smaller than the lumen, and positioned in its center and toward the flow, so the insonation angle is  $\leq 60^{\circ}$  (Figure 5).<sup>1,26</sup> After identifying the stenosis with B-scan and/or color Doppler, document the spectral tracing in this point, and pre-stenotic and post-stenotic regions. The velocity evaluation given by the spectral tracing must include the peak systolic velocity (PSV) and end-diastolic velocity (EDV) in CCA and ICA. Systolic and diastolic velocities provide the following relationships (ratios): PSV ICA/PSV CCA, PSV ICA/EDV CCA, and EDV ICA/EDV CCA.

Subocclusions are diagnosed based on the narrowing of the vessel lumen using color/power Doppler, with thin flow (string sign or trickle flow); however, it might be associated with high, low, or undetectable velocities, which occasionally hinders the diagnosis.<sup>1</sup>

In turn, carotid occlusions can be diagnosed with ultrasound as the absence of patent lumen in grayscale and undetected flow with color/power Doppler and/or spectral Doppler, in addition to the presence of high-resistance flow



Figure 2 – Right carotid and its anatomical subdivisions recommended by the group (adapted from the Mannheim study).9 CC: common carotid; IB: internal branch; BCA: brachiocephalic artery.

in CCA and staccato flow – flow with minimal velocity and very high-resistance at the occlusion or pre-occlusion site.<sup>1</sup>

The study group from DCI-BSC suggests the use of table 6 to quantify ICA stenosis.<sup>1</sup>

# 2.4.1.2. Quantification of carotid stenosis with anatomic parameters

The anatomical criterion (Figure 6) is based on the assessment of lumen reduction and should be used to characterize, in particular, stenosis below 50% (without hemodynamic repercussion); however, it is also a great contributor in stenosis greater than 50%, in which the hemodynamic criterion can fail to quantify stenosis accurately (e.g., severe aortic stenosis, significant contralateral carotid stenosis, among others). Lumen reduction is preferentially measured by diameter, and the result of the carotid stenosis range should be reported in intervals of 10%. It is recommended not to measure plaques smaller than 20% to avoid possible differences in the measurement of diameter reduction when the thickened intima is included or stopped being included.<sup>1</sup>

# **2.4.2.** Circumstances that can change the measurement of flow velocities and anatomic evaluation

Velocity evaluation can be compromised in some situations that affect measurements of spectral analysis. They can be located in the carotid bifurcation – distal or proximal – or even, in the contralateral carotid. Among the conditions proximal to the bifurcation, we underline aortic valve diseases (stenosis or insufficiency), atherosclerotic stenosis, or arteritis with involvement of the aortic arch, branches, and common carotid<sup>1</sup> (Table 7). Anatomic evaluation can be affected in circumstances such as arterial calcification with acoustic shadowing, improper adjustment of equipment, among others.

#### 2.4.3. Report description

Relevant information for the report:

• Specify the type of transducer used.

• Inform the technical quality of the examination (report situations that can lower its quality – e.g., presence of catheters).

• Describe the presence of atherosclerotic plaques, their location, extension, morphological characteristics, and degree of stenosis – quantified in deciles according to the DCI-BSC recommendation.<sup>1</sup>

• Report other findings of or related to carotid arteries (e.g., tortuosities, dissections, tumors, arteritis).

#### 2.5. Ultrasound assessment after carotid intervention

Treatment of symptomatic and asymptomatic carotid atherosclerotic disease has been the subject of multidisciplinary debate. Interventional treatment can be done by carotid endarterectomy or carotid stenting.

Ultrasound is the examination of choice for the followup after carotid intervention, and its protocol has the same sequence of the examination of carotid arteries without intervention, with some peculiarities in the intervention site. For more information about what to report on the vascular intervention site, the basic protocol of ultrasound follow-up, and velocity parameter tables, we suggest consulting the DCI recommendations recently published.<sup>1</sup>



Figure 3 – Recommendation from the Department of Cardiovascular Imaging of the Brazilian Society of Cardiology for the sequence of evaluation of carotid stenosis. PSV: peak systolic velocity.



Figure 4 – Normal flow patterns of carotid arteries. (A) Common carotid artery. (B) Internal carotid artery. (C) External carotid artery.



Figure 5 – (1) Diagram illustrating the placement of the cursor and the insonation angle. (A) Parallel to the jet in case of stenosis. (B) Parallel to the vessel. (2) Cursor and insonation angle toward the flow jet in case of stenosis (arrow).

% Anat Dist St (Nascet)	PSV cm/s	EDV cm/s	PSV IC/ PSV CC	PSV IC / EDV CC	EDV IC / EDV CC
< 50%	< 140	< 40	< 2.0	< 8	< 2.6
50 to 59%	140 to 230	40 to 69	2.0 to 3.1	8 to 10	2.6 to 5.5
60 to 69%	-	70 to 100	3.2 to 4.0	11 to 13	-
70 to 79%	> 230	> 100	> 4.0	14 to 21	-
80 to 89%	-	> 140	-	22 to 29	> 5.5
> 90%	> 400	-	> 5.0	> 30	
Subocclusion	Variable – thin flow				
Occlusion	Lack of flow	Lack of flow	Not applicable	Not applicable	Not applicable

#### Table 6 – Quantification of internal carotid artery stenosis (Department of Cardiovascular Imaging of the Brazilian Society of Cardiology).

The colors represent, from left to right, the most relevant criteria according to the literature. CC: common carotid; IC: internal carotid; EDV: end-diastolic velocity; PSV: peak systolic velocity.

#### 2.6. Ultrasound evaluation of vertebral arteries

The VUS evaluation of extracranial vertebral arteries contributes to the carotid study. It is divided into four segments: three extracranial and one intracranial (Figure 7).<sup>27,28</sup>

#### 2.6.1. Methodology to perform the examination

The patient's position is the same as that adopted for the carotid study.

To evaluate the extracranial part of vertebral arteries, start the examination by the V2 segment. At this point, with the aid of color Doppler and small angulation movements, try to identify the artery (as well as its vein) and record the spectral curves, adapting the scale and insonation angle of the vessel. Past this point, the artery is displayed toward its origin. The V3 segment of the vertebral artery lies below the mastoid process of the temporal bone (anatomic mark for the study). This region presents the vessel end of the transverse foramen and its course around the mastoid process (also called "atlas loop" due to its anatomical relationship with this vertebral body).<sup>3</sup> The typical parameters expected are:

Diameter: ranges from 2 to 4 mm. Caliber asymmetry among vertebral arteries is common (73% of cases), and the left one has a more dilated caliber in 50% of cases.<sup>28,29</sup> The normal standard with pulsed wave Doppler is waveforms with laminar antegrade flow of low-resistance, PSV between 20 and 60 cm/s in the V2 segment, and that can range from 30 to 100 cm/s in the origin of the vertebral artery. Due to the frequent diameter asymmetry, there is a considerable difference in PSV and resistance index (RI) among the normal vertebral arteries of an individual.<sup>27</sup>

Vertebral artery hypoplasia is defined as a vessel diameter equal to or lower than 2.0 mm (or caliber asymmetry among vertebral arteries with ratio > 1:1.7).<sup>28</sup>



Figure 6 – Measurement of lumen reduction. (A) Smooth atheromatous plaque in the lumen. (B) Irregular atheromatous plaque in the lumen.

#### Table 7 – Circumstances that can change the measurement of flow velocities.

Pathology	Abnormalities in VUS	Assessment alternatives				
Stenosis proximal to the common carotid artery or brachiocephalic artery	Reduced absolute flow velocities (PSV and EDV)	Using the velocity ratio and evaluation by the anatomical criterion				
Significant stenosis or contralateral carotid occlusion	Compensatory increase in flow velocities	Using the velocity ratio and evaluation by the anatomical criterion				
Arrhythmias (atrial fibrillation)	Variable velocity peaks	Waiting for the most regular period, or using an average of five beats and anatomical criterion				
Aortic valve stenosis	Reduced absolute flow velocities (PSV and EDV)	Using the velocity ratio and evaluation by the anatomical criterion				
Aortic valve insufficiency	Increase in PSV flow, with the possibility of retrograde diastolic flow	Using anatomical criterion or velocity ratio that does not involve EDV				

VUS: Vascular Ultrasound; EDV: end-diastolic velocity; PSV: peak systolic velocity.

#### 2.6.2. Quantification of stenosis

# Proximal stenosis (V0-V1) diagnosis results from the increase in flow velocities at the lesion site. The DCI-BSC standardization<sup>3</sup> suggests the values presented in table 8, adapted from the study by Hua et al.<sup>30</sup> Evaluate stenosis in the remaining segments with VUS based on multi-parameter analysis, such as turbulent flow with color Doppler, local increase in flow velocities, increase in velocity rates, and distal flow damping, since there are no tables of quantification of stenosis for these segments.

#### **3.** Abdominal aorta and branches

#### 3.1. Abdominal aortic aneurysm

#### 3.1.1. General considerations

Aneurysms are defined as a local dilation equal to or greater than 50% of the proximal or normal arterial diameter, necessarily involving all vessel layers. Even though the diameter of the abdominal aorta changes with age, gender, and biotype,



Figure 7 – Extracranial segments of the vertebral artery (V0-V3).

# Table 8 – Cut-off velocity values for proximal vertebral artery stenosis.

Stenosis	< 50%	50 to 69%	70 to 99%
V <sub>max</sub>	≥ 85 cm/s	≥ 140 cm/s	≥ 210 cm/s
VVR	≥ 1.3	≥ 2.1	≥ 4
EDV	≥ 27 cm/s	≥ 35 cm/s	≥ 55 cm/s

VVR: maximum velocity rate at the stenosis site and the V2 segment; EDV: end-diastolic velocity.

the mean diameter of the infrarenal aorta is approximately 2.0 cm, with upper normal limit < 3.0 cm. Thus, abdominal aortic aneurysm (AAA) is defined as an aorta that measures > 3.0 cm. AAAs are located between the diaphragm and the aortic bifurcation and can be classified as suprarenal, juxtarenal, and infrarenal. Approximately 85% of AAAs are infrarenal, and 5% involve the suprarenal aorta.<sup>31</sup> About 25% of patients with AAA have associated iliac artery aneurysm.<sup>2</sup>

Aneurysms can be fusiform, saccular, or with eccentric shapes. The type of asymmetry can significantly influence the risk of rupture and, as aneurysms grow, they can form laminated thrombi that preserve the arterial lumen.<sup>2</sup>

VUS is the most used examination to screen and diagnose asymptomatic patients in emergency units without a prior diagnosis and symptomatic ones. Computed tomography angiography (CTA) is the examination of choice for preand postoperative assessment; however, VUS does not lose its value for being more accessible, costing less, and not using nephrotoxic contrast. The current availability of microbubble contrast makes VUS quite attractive, particularly in postoperative assessments.<sup>32</sup>

#### 3.1.2. Clinical indications<sup>2</sup>

• Screening (Table 9).

 $\,$  Follow-up: monitor the growth and determine the appropriate time for surgery.  $^{\rm 33}$ 

• Evaluation of pulsating abdominal mass, signs of rupture, or growth.

• Preoperative AAA examination: report data on the access route, abnormalities in iliac arteries, fixation site of the endoprosthesis, aspect and measurements of the aneurysmal sac, and presence of parietal thrombi.

• Postoperative AAA examination.

#### 3.1.3. Examination instructions and protocols (Table 10)

# 3.1.4. Examination protocol for abdominal aortic aneurysm according to recommendations from the department of cardiovascular imaging<sup>2</sup>

The examination must be conducted from the subxiphoid region to the aortic bifurcation, followed by the study of

 
 Table 9 – Recommendation from the Department of Cardiovascular Imaging for screening and follow-up of abdominal aortic aneurysm.

#### Screening

· Men aged 65 to 75 years

- Men aged 55 to 75 years with a family history of AAA and/or who smokes
- Women aged 55 to 75 years with a family history of AAA and/or who smokes

#### Follow-up interval

- 2.6 to 2.9 cm reevaluate in 5 years (sub-aneurysmal dilation)
- 3.0 to 3.9 cm 24 months
- 4.0 to 4.5 cm 12 months
- 4.6 to 5.0 cm 6 months
- > 5.0 cm 3 months

#### Indication for intervention

- ≥ 5.5 cm
- · AAA-related symptoms
- Growth rate > 1.0 cm per year

AAA: abdominal aortic aneurysm.

right and left common iliac arteries and their external and internal branches.

The evaluation uses B-scan, with transverse, coronal, and longitudinal planes to detect atheroma plaques and measure the diameters, especially if dilations are found. Take the anteroposterior (AP) measurement of the aneurysm during the peak systolic expansion, reporting if it was made from outer wall to outer wall (OTO) or inner wall to inner wall (ITI).

• Screening: use the B-scan during the subxiphoid part of the echocardiogram or in routine abdominal ultrasound.

• **Diagnosis and follow-up**: screen the aorta from the subxiphoid region to its bifurcation, followed by the study of iliac arteries and branches.

• **Preoperative assessment**: descriptions, necessary measurements, and relevant data are described in figure 8.

• **Postoperative assessment**: inform the surgical techniques used. They are described in detail in the DCI recommendations.<sup>2</sup>

Essential information to include in the medical report (Figure 8):

#### Table 10 – Examination instructions and protocols for the study of abdominal aorta and branches.

Examination instructions	Abdominal aorta	Aortoiliac segment	Mesenteric arteries and celiac trunk	Renal arteries
Low-frequency convex or sector transducers (2 to 5 MHz)	Х	Х	х	Х
Preferentially in the morning with 6- to 8-h fasting	x	х	х	x
The patient should not smoke, chew gums, or consume carbonated beverages	х	Х	x	x
Optional antiflatulent	x	х	Х	х
Supine position with head raised at $30^{\circ}$	х	х	Х	х
Lateral position	x	х	-	х
Transverse, coronal, and longitudinal planes	х	х	Х	х
What to evaluate:				
B-scan:				
Dimensions	x	х	x	х
Anatomic changes	x	х	x	х
Morphology of walls and plaques	х	х	Х	х
Presence of thrombi	х	х	-	-
Color Doppler:				
Aliasing	х	х	Х	х
Lack of flow (occlusion)	х	х	Х	х
Spectral Doppler:				
PSV	-	х	Х	х
PSV ratio (V2/V1)	-	х	-	-
EDV	-	-	х	х
Renal aortic ratio	-	_	-	X

EDV: end-diastolic velocity; PSV: peak systolic velocity.



Figure 8 – Preoperative assessment of abdominal aortic aneurysm (AAA).

#### · Diagnostic examination:

- Report technical difficulties.
- Measure the largest diameter of the aorta.

- Inform the presence of tortuosities or enlargement of the aorta.

- Define the location of the aneurysm: supra-, juxta-, or infrarenal.

- Describe the anatomical shape of the aneurysm: saccular, fusiform, or other.

- Inform the presence or absence of wall thrombi, the intraluminal diameter, and signs of rupture.

• Additional information for the preoperative examination:

- Diameters:

- Neck in the lowest renal artery plane.
- o Neck 15 mm below the origin of the lowest renal artery.
- Maximum AAA AP transverse plane (ITI or OTO).
- o Bifurcation plane (when present).
- Both common iliac arteries.
- Iliac artery bifurcation.
- Longitudinal extension of the proximal neck.
- Neck angle with aneurysm axis.
- Longitudinal extension of AAA.

#### 3.2. Aortoiliac atherosclerotic disease

VUS allows the identification, localization, and anatomic extension of atherosclerotic lesions, and evaluates the aortic wall to register not only the presence of atherosclerotic lesion but of ulceration, calcification, thrombus, dissection, and dilation. When performed by trained and experienced professionals, this technique has good diagnostic accuracy for aortoiliac atherosclerotic disease, with 86% sensitivity and 97% specificity for lesions > 50% stenosis.<sup>34</sup>

# 3.2.1. Clinical indications for venous ultrasound in cases of aortoiliac atherosclerotic disease

• Symptoms of acute ischemia (distal embolism) and intermittent claudication with decreased or absent femoral pulse, gluteal claudication, erectile dysfunction, and pain at rest.

• Clinical signs such as abdominal bruit and reduced anklebrachial index.

• Prior VUS showing abnormalities in the velocity curve pattern of femoral arteries.

- $\bullet$  Follow-up of grafts and endoprostheses for the treatment of aortoiliac obstruction.\*
  - Suspected diagnosis of aortic dissection.
  - Suspected diagnosis of arteritis.

<sup>\*</sup>This document will not cover the VUS assessment in the postoperative follow-up of aortoiliac obstructions, in case of suspected diagnosis of aortic dissection and arteritis. We suggest the recent publication of DCI recommendations as additional reading.<sup>2</sup>

3.2.2. Examination protocol for aortoiliac atherosclerotic disease according to dci recommendations (Table 10)

#### 3.2.2.1. Diagnostic criteria

• **Stenosis**: measure PSV at the lesion site (V2) and 1 to 2 cm proximal to the lesion (V1) and calculate the velocity ratio (V2/V1). Determine the spectral curve with an angle  $\leq 60^{\circ}$  parallel to the turbulent flow axis (Figure 9). The degree of stenosis should be classified according to table 11.

• **Occlusion**: lack of flow in any aortoiliac segment, even with scan parameters that detect low-velocity flows. Presence of typical preocclusive waveform (high peripheral resistance, low peak systolic velocity, and lack of diastolic flow). Collateral vessels can be found in occluded pre- and post-segment (reentry point). The post-occlusive spectrum is characterized by monophasic waveform, with reduced PSV and prolonged acceleration time - *parvus/tardus* (Figure 9). Hypoechoic image with a concave interface in colored flow and spectrum in preocclusive *staccato* pattern suggests thromboembolic occlusion.

- Essential information to include in the medical report:
- Diagnostic examination:
  - $\circ$  Report if there were technical difficulties during the examination.
  - $\circ$  In case of dilations, inform the largest diameter of the aorta and/or iliac arteries.

#### - Additional information for the preoperative examination:

• Inform the presence, aspect, and location of atherosclerotic plaques, as well as the degree of stenosis of lesions. Table 12 lists the general limitations of VUS examination.

#### 3.3. Mesenteric arteries

#### 3.3.1. General considerations

Mesenteric vessels are represented by the celiac trunk (CT) and superior and inferior mesenteric arteries (SMA and IMA, respectively). Anatomically, CT starts just below the aortic hiatus of the diaphragm and originates the splenic and hepatic arteries.

SMA and IMA begin approximately 0.5 to 2 cm below CT and 4 to 5 cm above the aortic bifurcation, respectively (Figure 10). $^{35}$ 

# Table 11 – Classification of the degree of stenosis with pulsed wave Doppler.

Classification	Systolic velocity ratio
Stenosis < 50%	V2/V1 < 2.0
Stenosis ≥ 50%	V2/V1 ≥ 2.0
Stenosis ≥ 70%	V2/V1 ≥ 4.0
Occlusion	Lack of flow



Figure 9 – Color flow imaging showing the flow proximal to the lesion in red and the turbulent flow at the lesion site (arrow). The diagrams A and C demonstrate the velocity spectrum with Doppler. (A) Cursor proximal to the lesion to measure V1. (B) Cursor at the lesion site to measure V2. (C) Cursor distal to the lesion with damped waveform.

Aorta and iliacs	Mesenteric arteries/Celiac trunk	Renal arteries	
Hostile abdomen	Hostile abdomen	Hostile abdomen	
Obesity	Obesity	Obesity	
<ul> <li>Intestinal meteorism</li> </ul>	<ul> <li>Intestinal meteorism</li> </ul>	<ul> <li>Intestinal meteorism</li> </ul>	
• Examiner-dependent	<ul> <li>Patient with severe abdominal pain – in acute ischemia</li> </ul>	Anatomic changes	
<ul> <li>Low-quality equipment</li> </ul>	<ul> <li>Examiner-dependent</li> </ul>	Examiner-dependent	
	Low-quality equipment	<ul> <li>Low-quality equipment</li> </ul>	

Table 12 – General limitations of vascular ultrasound in the evaluation of abdominal aorta and branches



Figure 10 – Abdominal aorta and branches. SA: splenic artery; HA: hepatic artery; RCIA: right common iliac artery; LCIA: left common iliac artery; IMA: inferior mesenteric artery; RRA: right renal artery; LRA: left renal artery; SMA: superior mesenteric artery; CT: celiac trunk.

Mesenteric artery obstructive disease progresses chronically and asymptomatically. The clinical manifestation represented by postprandial abdominal pain (mesenteric angina) and/or progressive weight loss occurs when two or more mesenteric vessels are involved. Older men are more frequently affected. Atherosclerosis is responsible for more than 90% of diseases that strike mesenteric arteries and is usually dissemination of the atheromatous process that involves the entire aorta.<sup>36</sup>

Arteriography is the standard diagnostic method; however, VUS is the first examination indicated for the study of symptomatic chronic intestinal ischemia for being non-invasive and risk-free.<sup>37</sup>

The objective of VUS is to determine the presence, location, extension, and severity of the stenotic lesion. The study must include the SMA, CT, and IMA, with the evaluation of SMA and CT being more important.<sup>38</sup>

#### 3.3.2. Clinical indications

• Recurrent postprandial abdominal pain (mesenteric angina).

- Weight loss without a known cause.
- Abdominal bruit.

The DCI recommendations describe in detail the protocols of follow-up after surgical or endovascular treatment and assessment of compression syndromes.<sup>2</sup>

# 3.3.3. Examination preparation and general protocol (Table 10)

#### • Specific examination protocol:<sup>39,40</sup>

- Place the transducer in the epigastric region; using the transverse plane, identify the SMA anteriorly, the aorta posteriorly, and the right renal vein between these two vessels (Figure 11A). In this same location, using the longitudinal plane of the aorta, find the CT and SMA (Figure 11B).

- B-scan: evaluates the presence of atherosclerotic or aneurysmal disease.

- Color flow imaging: assesses vessel patency and indicates flow turbulence, suggesting the probable stenosis site.

- Spectral analysis: used to analyze PSV and EDV. Measure these velocities in the origin of the vessel and/or in places with suspicion of stenosis. Position the insonation angle toward the blood flow.

- Velocities of mesenteric vessels are influenced by respiration; therefore, patients should hold their breath during the measurement.



Figure 11 – Mesenteric vessels and abdominal aorta. (A) Transverse plane with B-scan showing the superior mesenteric artery (SMA) anteriorly and the abdominal aorta (AA) posteriorly. (B) Longitudinal plane of the abdominal aorta and emergence of the celiac trunk and SMA.

 $\bullet$  Table 13 presents the ultrasonographic criteria to assess flow in CT and SMA.  $^{2,41\cdot43}$ 

#### 3.3.4. Essential information to include in the medical report

 ${\boldsymbol \cdot}$  Report if there were technical difficulties during the examination.

- Presence or absence of atherosclerotic disease.
- Lesion site.
- Measurement of stenosis.
- Measurement of PSV and EDV.

#### 3.3.5. Examination limitations

Table 12 lists the examination limitations.

#### 3.4. Renal arteries

#### 3.4.1. General considerations

Prevalence of renal artery stenosis (RAS) changes according to the population studied. RAS is the most common cause of secondary hypertension among the general population of hypertensive patients, representing approximately 1 to 6% of cases.<sup>44,45</sup>

The most frequent cause of RAS is atherosclerosis (85% to 90% of cases), which often strikes the origin of and/ or the segment proximal to the renal artery and can be unilateral or bilateral. Its prevalence increases with age, diabetes, and atherosclerosis in other arterial sites. It is considered an independent predictor of adverse events such as acute myocardial infarction, CVA, and death due to cardiovascular causes.<sup>45,46</sup>

Fibromuscular dysplasia – a non-inflammatory disease – is responsible for 10% of RAS cases. Its frequency among the general population is unknown, but it is more usually reported in young women. Renal artery involvement occurs in its middistal segment and is often bilateral.<sup>47</sup>

With the quality improvement of imaging methods, RAS diagnosis became feasible and of great interest, aiming to identify patients who would benefit not only from drug therapy but also from renal revascularization procedures. Among the imaging examinations, VUS is the initial method

|--|

Artery	Normal Doppler	Stenosis ≥ 50%	Stenosis ≥ 70%	Occlusion
СТ	Low-resistance	PSV > 240 cm/s EDV > 40 cm/s	PSV > 320 cm/s EDV > 100 cm/s	Lack of flow Retrograde CHA flow
SMA Fasting	High-resistance	PSV > 295 cm/s EDV > 45 cm/s	PSV > 400 cm/s EDV > 70 cm/s	Lack of flow
SMA Postprandial	Low-resistance	PSV > 295 cm/s EDV > 45 cm/s	PSV > 400 cm/s EDV > 70 cm/s	Lack of flow

CHA: common hepatic artery; SMA: superior mesenteric artery; CT: celiac trunk; EDV: end-diastolic velocity; PSV: peak systolic velocity.

of choice to investigate RAS due to its advantages, such as being non-invasive, low-cost, having no risk of radiation, and, mainly, the lack of contraindications related to the use of nephrotoxic contrast. Ultrasound evaluation of renal arteries has high specificity in competent laboratories, demonstrating that, when the vessels are correctly assessed, its results rarely differ from those obtained by arteriography.<sup>45,48-50</sup>

Table 14 shows the main indications to investigate RAS, according to the principal American guidelines.<sup>51,52</sup>

The principal objectives of the study of renal arteries are:

• Identify the main renal arteries and, if possible, the accessory ones.

• Locate and grade stenotic lesions resulting from atherosclerotic disease or not.

- Monitor the progression of RAS.
- · Follow-up after renal artery revascularization.

#### 3.4.2. Examination protocol (Table 10)

• Patient's position: Supine, using transverse and longitudinal ultrasound planes to analyze the abdominal aorta and origin of renal arteries.

• Lateral, using the coronal plane to assess all middle and distal segments of renal arteries. Use this plane to measure the longitudinal diameter of the kidney and analyze the intrarenal flow in segmental or interlobular arteries.<sup>2</sup>

• B-scan: used to identify atheroma plaques in the renal artery and evaluate the echogenicity and size of the kidney.

• Color flow imaging and/or power Doppler: evaluate artery patency. Power Doppler indicates possible stenosis sites through flow turbulence or reduction in the vessel lumen.

• Spectral analysis: Essential to measure systolic and diastolic velocities (PSV and EDV). Use the transverse or coronal plane, keeping the cursor directed at the flow jet, with an insonation angle  $< 60^\circ$  in the renal artery and 0° in intraparenchymatous arteries.  $^{53}$ 

• Measure PSV and EDV at the origin of the renal artery or in any segment with suspicion of stenosis.

• Use the longitudinal plane of the aorta, near the origin of the SMA, to measure its PSV and calculate the renal aortic ratio (RAR).

#### 3.4.3. Diagnostic criteria for renal artery stenosis

Diagnostic criteria for RAS are classified as direct and indirect. The first consists of evaluating the renal artery from its origin in the aorta and, if possible, in all its extension. The second analyzes the hemodynamic repercussion of the proximal lesion of the renal artery on intraparenchymatous arteries.

The direct criterion comprises renal artery PSV and EDV and aortic PSV to calculate RAR (renal artery PSV/aortic PSV). The low-resistance waveform is considered normal for the renal artery. PSV is the most accurate parameter to grade RAS, with values that range between 180 and 250 m/s from study to study. EDV and RAR are used to aid the RAS evaluation. According to several studies, RAR ranges from 3.2 to 3.7 to estimate hemodynamically significant stenosis. Situations such as aortic coarctation, severe left ventricular dysfunction, aortic dissection or aneurysm, and systolic aortic velocities > 100 cm/s or < 40 cm/s can change the velocities in renal arteries and their relationships.<sup>2</sup>

Indirect criteria, represented by the measurement of the kidney size and analysis of intraparenchymatous artery flow, should be combined with direct criteria to optimize the

results. A difference of 1.5 cm in kidney size can result from hemodynamically significant stenosis or even renal artery occlusion. Regarding the analysis of intraparenchymatous artery flow, initially, there is a reduction in the first systolic peak (FSP), prolonged acceleration time (AT) with decreased acceleration rate (AR), and flattening of the systolic wave until the *parvus/tardus* pattern is found. AT > 70 ms is associated with 60% RAS, while the *parvus/tardus* flow is present in more severe stenosis (80%).<sup>2</sup>

#### Table 14 – Clinical indications to investigate renal artery stenosis.

- Onset of hypertension in patients aged  $\leq$  30 years
- Onset of severe hypertension in patients aged ≥ 55 years
- · Patients with accelerated hypertension (sudden or persistent worsening of previously controlled hypertension)
- Patients with resistant hypertension (treatment failure with full doses of three antihypertensive drug classes, including diuretics)

• Patients with malignant hypertension (with target-organ damage: acute renal failure, acute congestive heart failure, new visual or neurological disorder, and/or advanced retinopathy)

- Patients with worsening of renal function after administration of angiotensin-converting enzyme inhibitor or angiotensin-receptor blocker
- Patients with unexplained renal atrophy or discrepancy > 1.5 cm from kidney sizes
- Patients with sudden and unexplained pulmonary edema (flash pulmonary edema)
- · Patients with renal failure or unexplained congestive heart failure
- Patients with refractory angina
- Patients with multiple vessel coronary artery disease
- Patients with abdominal aortic aneurysm

Table 15 summarizes the criteria recommended by DCI-BSC for hemodynamically significant RAS (> 60%). All references to these criteria are detailed in the guideline.<sup>2</sup>

Renal artery occlusion might be suggested if the vessel flow is not seen during color flow imaging and/or power Doppler, and not detected with pulsed wave Doppler, associated with a longitudinal diameter of the ipsilateral kidney < 8.5 cm.

Follow-up after renal revascularization is not part of the scope of this publication and can be found in the DCI guidelines.<sup>2</sup>

#### 3.4.4. Limitations of the renal artery study

Listed on table 12.

# 3.4.5. Essential information to include in the medical report

• Report if there were technical difficulties during the examination.

• Inform the presence or absence of atherosclerotic disease or signs of fibromuscular dysplasia.

• Lesion site.

- Measurement of stenosis.
- Measurement of renal artery PSV and EDV.
- Measurement of aortic PSV.

• Measurement of pulsatility index (PI) and RI in intraparenchymatous arteries (preferably the segmental artery).

• Measurement of kidney size.

#### 4. Lower-limb arteries

VUS can evaluate peripheral arterial diseases (PADs) with high accuracy, enabling the anatomical and functional assessment of arterial lesions, in addition to identifying the location, extension, and hemodynamic repercussion of stenosis or occlusion.<sup>54,55</sup>

#### 4.1. Clinical indications

 $\bullet$  Anatomic diagnosis of stenosis or occlusion in the stenotic PAD in symptomatic patients considered for revascularization.  $^{56-59}$ 

• Follow-up of the progression of stenotic disease previously diagnosed.

 $\,$  Surgical the rapeutic planning for patients diagnosed with PAD.  $^{\rm 60,61}$ 

- Diagnosis and follow-up of peripheral arterial aneurysms.<sup>62</sup>
- Diagnosis, follow-up, and treatment of pseudoaneurysms.<sup>63,64</sup>

 $\bullet$  Evaluation of autogenous or synthetic vascular grafts, with follow-up and diagnosis of complications.  $^{65\cdot 67}$ 

• Monitoring of arterial sites submitted to percutaneous intervention, such as angioplasty, thrombolysis, thrombectomy, atherectomy, and stenting.<sup>68-71</sup>

• Confirmation of significant arterial abnormalities detected by another imaging method.

• Evaluation of vascular and perivascular abnormalities, such as masses, aneurysms, pseudoaneurysms, dissections, thrombosis, embolism, vascular malformation, and arteriovenous fistula (AVF).

• Evaluation of arterial integrity in trauma.

• Evaluation of artery compression syndromes, such as popliteal artery entrapment.

#### 4.2. Examination protocol (Table 16)

#### 4.3. Diagnostic criteria

Stenosis: measure PSV at the lesion site (V2) and 1 to 4 cm proximal to the lesion (V1) and calculate the velocity ratio (V2/V1). Obtain the spectral curve with an angle  $\leq 60^{\circ}$  parallel to the turbulent flow axis (Figure 9). The degree of stenosis should be classified according to table 17.<sup>60,61</sup> Other criteria that can assist in grading stenosis are: prolonged AT in distal arteries, which could indicate hemodynamically significant lesions in proximal segments.

Occlusion: lack of flow in any lower-limb arterial segment, even with scan parameters that detect low-velocity flows. Presence of typical preocclusive waveform (high peripheral resistance, low peak systolic velocity, and lack of diastolic flow). Collateral vessels can be found in occluded pre- and post-segment (refilling point). The post-occlusive spectrum is characterized by monophasic waveform, with reduced PSV and prolonged AT (*parvus/tardus* pattern). Hypoechoic image with concave interface in colored flow and spectrum in standard pre-occlusive staccato suggests thromboembolism (Table 18).

Table 15 – Velocity criteria to quantify renal artery stenosis both native and after stenting.

Degree of stenosis	Renal artery PSV	Renal aortic ratio	Renal artery EDV	Intrarenal flow	Renal artery PSV after stenting	Renal aortic ratio after stenting
Normal	< 200 cm/s	< 3.5	< 150 cm/s	AT < 70 ms	< 390 cm/s	< 5
< 60%	≥ 200 cm/s	< 3.5	< 150 cm/s	AT < 70 ms	< 390 cm/s	< 5
≥ 60%	≥ 200 cm/s	≥ 3.5	< 150 cm/s	$AT < or \ge 70ms$	> 390 cm/s	≥ 5
≥ 80%	≥ 200 cm/s	≥ 3.5	≥ 150 cm/s	AT ≥ 70 ms <i>tardus/</i> <i>parvus</i> flow	≥ 390 cm/s	≥ 5
Occlusion	-	-	-	Might have <i>tardus/</i> <i>parvus</i> flow	-	-

AT: acceleration time; EDV: end-diastolic velocity; PSV: peak systolic velocity.

	4.0	<b>D</b> ( )	e					
Iahla	16.	- Protocol	tor di	adnoetic	examination	and	nronorativa	manning
TUDIC	10	11010001	101 01	agnostic	CAUTINIALION	unu	proportative	, mapping.

Artony	B-s	can	Puls	ed Wave Doppler	Color Imaging
Antery	Normal	Abnormal	Normal	Abnormal	Color Imaging
CFA				Longitudinal: evaluate the flow in	
Proximal SFA				the stenosis, and proximal and distal to the stenosis	Assess vessel patency with intraluminal color
Mid SFA	Transverse: evaluate the			Use the peak systolic velocity in	filling
Distal SFA	diameter and aspect of	Transverse: measure the dilations and, if possible,	Longitudinal: use an angle $\leq 60$ degrees	the site of highest velocity (V2), with angle $\leq$ 60 degrees, and 1 to	Occlusion: lack of color filling
PA	Les situdis de sustante	the intraluminal stenosis	laminar flow, and	4 cm proximal to the lesion (V1) to calculate the velocity ratio (V2/V1)	
PTA	the aspect of the wall and intraluminal diameter	Longitudinal: measure the lesion extent	multiphase curve (triphase)	Flow in the post-stenotic segment:	Shows flow turbulence – mosaic aspect (aliasing)
ATA				velocity turbulence or decrease	Guide the volume scan of
FA				Post-stenotic velocity curve of the parvus/tardus type indicates	the sample to detect the point of highest velocity
TFT				nemodynamic repercussion	

FA: fibular artery; CFA: common femoral artery; DFA: deep femoral artery; SFA: superficial femoral artery; PA: popliteal artery; ATA: anterior tibial artery; PTA: posterior tibial artery; TFT: tibial-fibular trunk.

#### Table 17 – Classification of the degree of stenosis in native arteries with pulsed wave Doppler.

Classification	Systolic velocity ratio
Stenosis < 50%	V2/V1 < 2.0
Stenosis ≥ 50%	V2/V1 ≥ 2.0
Stenosis ≥ 70%	V2/V1 ≥ 4.0
Occlusion	Lack of flow

Aneurysm: report the identification and location of the aneurysm. Measure the largest diameter including the adventitial layer (out-out) with two-dimensional image in transverse plane. Investigate and document the presence of intraluminal thrombus with color flow imaging.

Pseudoaneurysm: characterized by dilation that does not compromise all arterial layers and having a connecting channel with the arterial lumen. Evaluate all of these structures with two-dimensional image and color imaging, measuring the calibers. Use spectral Doppler in the connecting channel, where the typical "to-and-fro" flow can be found (Table 18). Therapeutic interventions should use color imaging and pulsed wave Doppler as a guide, evaluating blood flow in the native artery, aiding in the puncture – in cases of thrombin treatment –, and verifying the thrombosis of the pseudoaneurysm in all treatment modalities.<sup>63</sup>

Table 18 presents the different patterns of arterial flow, with their names, clinical meanings, and main occurrence situations.

#### 4.4. Essential information to include in the medical report

• Report if there were technical difficulties during the examination.

• Wall aspect and diameter of all arteries studied, when necessary.

• Presence, aspect, location, and degree of stenosis of atherosclerotic plaques.

• Stenosis velocity and characteristics of velocity curves in the post-stenotic segment.

• Signs of segmental or complete arterial occlusion.

# 4.5. General limitations of the peripheral artery ultrasound

- Examiner-dependent.
- · Equipment of low technical quality.
- · Lower-limb edema.
- Unhealed ulcer.

#### 5. Arteriovenous fistulas for hemodialysis

#### 5.1. Introduction

AVFs can be congenital, traumatic, or specifically created for hemodialysis. This guideline will cover only the technical aspects of VUS evaluation of AVF for hemodialysis (AVFH), which can be of two types:

1. Autogenous – radiocephalic or brachiocephalic (Brescia-Cimino) fistula. Figure 12 shows the latero-lateral connection between artery and vein in A; the terminal connection between the artery and lateral of the vein in B; the terminal connection between the vein and lateral of the artery in C; and the termino-terminal connection between artery and vein in D.<sup>72</sup>

2. Polytetrafluorethylene (PTFE) grafts. Figure 13 shows an example of a straight PTFE graft between the basilic vein and

#### Table 18 – Arterial flow patterns in various situations.

	Name	Clinical meaning	Occurrence situation
AAAAA	Triphasic	Found in young and normal individuals	Normal
	Triphasic without the elastic component	Found in old adults with reduced vessel elasticity	<ul><li>Normal old adults</li><li>Bad US beam alignment</li></ul>
Man han han ha	Biphasic hyperemic	Without retrograde component, but reaches the baseline. Increased velocities	<ul> <li>Inflammatory process</li> <li>Normal reactive vasodilation</li> </ul>
July Mark	Biphasic post-obstructive	Without retrograde component, but reaches the baseline. Low velocities	After moderate obstructions
1	Biphasic with retrograde pandiastolic component	Retrograde pandiastolic component. Usually with normal velocities	Vessels that provide collateral to other stenotic arteries. In most cases, vessels without proximal lesions
A mark	Biphasic "to-and-fro"	Pronounced retrograde pandiastolic component	<ul><li>Pseudoaneurysm</li><li>Type II endoleak</li></ul>
- 80 - 60 - 40 - 20 - 0	Acute monophasic	Low velocity, with little increase in acceleration time	Moderate and moderate to severe post- stenosis, WITH vasodilation reserve
	Damped monophasic	Low velocity, with increased acceleration time	Severe post-stenosis or post-occlusion, WITH vasodilation reserve
- 30 15 0	Extremely damped monophasic	Very low velocity, with high increase in acceleration time	Post-occlusion, WITH vasodilation reserve
<b>A</b>	Monophasic without diastole	Extremely low velocity, with increased acceleration time, without diastole flow	Post-occlusion, WITHOUT vasodilation reserve. Severe multisegmental lesions
ndelang anto bean pagenainan dituri	Continuous	Extremely low velocity, with acceleration time so increased that it is not possible to differentiate PSV from EDV	Post-occlusion, WITH vasodilation reserve. Severe multisegmental lesions
· · · · · · · · · · · · · · · · · · · ·	Retrograde	Low velocity, can be damped or not, and can even have a retrograde component	Retrograde filling of a vessel with proximal occlusion

EDV: end-diastolic velocity; PSV: peak systolic velocity.

radial artery in A; a loop PTFE graft between the basilic vein and radial artery in B; a curved PTFE graft between the artery and brachial vein in C; and a loop PTFE graft between the great saphenous vein and femoral artery in the lower limb in D.<sup>73</sup>

The most common AVFH types are the radiocephalic and brachiocephalic;<sup>73</sup> however, as an alternative, they can be created between other vessels, such as the ulnar artery and basilic vein, or the brachial artery and basilic vein, but they must be superficialized.

#### 5.2. Examination indications

The main indications for VUS in cases of AVFH are:

• Preoperative vascular mapping, with planning for the procedure, including evaluation of central vessels.<sup>74,75</sup>

• Maturation assessment, especially in obese patients. Maturation occurs when AVFH can withstand repeated venous punctures with large-caliber needles. This process can fail in up to 60% of cases.<sup>74,76,77</sup>

• Functional follow-up of AVFH to detect complications early.

• Recommendation: perform an examination before the procedure for planning and two after to reduce its failure rate.<sup>74,78</sup>

• Contraindications for the procedure: Paget-Schroetter syndrome and deep venous thrombosis (DVT).

#### 5.3. Technique<sup>79</sup>

• High-tech equipment – with high-frequency and/or variable frequency linear probe.

• Patient at rest in a room at ambient temperature, with the upper limb extended parallel to the body.

• Measure the anteroposterior diameters of vessels in transverse planes.

• Possible sites to measure flow volume: afferent – in the artery, 1 to 2 cm before the anastomosis; and efferent – in the vein, 1 to 2 cm after the anastomosis.

• Observe the abnormalities in two-dimensional planes and analyze turbulent flows with color flow imaging

 $\bullet$  The Doppler angulation should be parallel to the blood flow (closer to  $60^\circ).^{79}$ 

• Spectral Doppler characteristics: artery – low-resistance flow; vein – arterialized flow pattern.

#### 5.4. Diagnostic criteria

# 5.4.1. Protocol to create arteriovenous fistulas for hemodialysis

• Choose the non-dominant member, if possible.

 $\bullet$  Follow the order:  $1^{st}$  option – wrist;  $2^{nd}$  option – elbow;  $3^{rd}$  option – prosthesis.

 $\bullet$  Measure the arterial and venous diameter (a tourniquet can be used in the member to evaluate venous distensibility).  $^{80,81}$ 

- Ideal venous or arterial diameter: elbow – 4 mm; wrist – 3 mm.

- Minimum functional diameter: elbow – 2 mm; wrist – 1.8 mm.  $^{\rm 72}$ 

 $\cdot$  Measure the depth between vessels and skin surface; the ideal for puncture is < 5 mm.  $^{79}$ 

• Measure the distance between artery and vein to be used (Figure 14).



Figure 12 – Types of brachiocephalic fistulas (Brescia-Cimino). (A) Latero-lateral between artery and vein. (B) Terminal artery – lateral vein. (C) Terminal vein – lateral artery. (D) Terminal between artery and vein.

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## **Special Article**



Figure 13 – Types of polytetrafluoroethylene (PTFE) grafts. (A) Straight PTFE graft between the basilic vein and radial artery. (B) Loop PTFE graft between the basilic vein and radial artery. (C) Curved PTFE graft between the brachial artery and vein. (D) Loop PTFE graft between the great saphenous vein and femoral artery.

#### 5.4.2. Protocol after procedure<sup>77,79,81</sup>

• Measurements of artery, vein, and anastomosis;

- Ideal venous or arterial diameter: elbow – 4 mm; wrist – 3 mm

- Minimum functional diameter: 2 mm

• Flow volume: to obtain the effective flow volume through an AVFH, measure the diameter of the drainage vein by its inner edges (in cm), preferably with a transverse plane, to calculate its radius (R) and, consequently, its area (in cm<sup>2</sup>). Take this measurement farther from the anastomosis site of the fistula, where the color flow imaging shows no flow turbulence, usually 2 to 5 cm from the anastomosis. Find the flow in the same place of the vein where the diameter was assessed with pulsed wave Doppler. Locate the sample volume in the center of the vessel and correct the Doppler flow angle to up to 60°. Determine the mean flow velocity (V<sub>MEAN</sub>) in cm/s with pulsed wave Doppler. Calculate the mean of 3 to 5 cardiac cycles. Apply the following formula:

Mean flow (ml/min) =  $V_{\text{MEAN}}$  (cm/s) × R<sup>2</sup> (cm<sup>2</sup>) ×  $\pi$  × 60 (s)

The values below are used for normal flow volume of homologous and heterologous AVFH:

- Brescia-Cimino fistula (radiocephalic) =  $614 \pm 242$  ml/min.
- $PTFE = 464 \pm 199 \text{ ml/min.}$
- Mean normal value = 514 ml/min.

Fistulas with flow volume < 450 ml/min have a high risk of thrombosis in 2 to 6 weeks.

The minimum functional flow volume for AVFH in the elbow is  $\geq 200$  ml/min, and in the wrist is  $\geq 150$  ml/min.

Hyperflow is considered when the flow volume is > 3,000 ml/min.

• Maturation protocol:<sup>73,74,77</sup> AVFH maturation consists of an increase in caliber and flow of the vessels used.

Assess vessel diameter and structure ( $\geq$  6 mm), flow volume ( $\geq$  600 ml/min), and the distance between the vein and skin surface  $\leq$  6 mm.

Wait at least 14 days; the ideal scenario is individualization with follow-up for each patient and interval superior to 30 days. If possible, wait 60 or even 90 days before the first canalization.

Table 19 summarizes the main objectives of VUS findings.



Figure 14 – Measurement of the distance between the brachial artery and basilic vein before the creation of alternative arteriovenous fistulas.

# Table 19 – Objectives of arteriovenous fistulas for proper hemodialysis (95%)<sup>77.</sup>

- Vascular diameter: > 4 mm
- Flow volume: > 500 ml/min
- Maturation time: > 30 days
- Diameters < 3mm and flow volume < 400 ml/min = high probability of failure
- · Vessel diameter should increase with time

# 5.4.3. Causes and types of failures (autogenous and prosthetic) $^{74,76,79,82}\,$

- Arterial: diabetes mellitus and significant atheromatosis.
- Venous: fibrosis.
- Anastomosis site: turbulence and intimal hyperplasia.
- Significant escape through tributary veins.

• Steal phenomenon, in which retrograde flow is found in the radial artery distal to AVFH (Figure 15).

- Pronounced tortuosities.
- Intimal hyperplasia (valves).
- Thrombosis (dissection by puncture).

· Idiopathic: puncture and surgical technique.

• Significant stenosis with V2/V1  $\ge$  4 (if  $\ge$  2 indicates stenosis > 50%; angioplasty is recommended in case of clinical and/or hemodynamic abnormalities associated).

Occlusion.

• Aneurysm, dilation due to prosthesis degeneration and pseudoaneurysm.

- Infection.
- Hematoma, seroma, and lymphocele.

#### 5.4.4. Examination limitations

Inexistent, with rare exceptions in highly significant edemas, fibrotic scars, and presence of orthopedic devices.

#### 5.4.5. Suggestions to elaborate the report and conclusion

1. Examination before the procedure (Table 20):

• Report deep and superficial venous thrombosis; abnormalities in the arterial system; and anomalous anatomic changes, such as the number of vessels, paths, and forearm vessels that continue in the arm

• Inform the measurements of calibers and proper flows.



Figure 15 – Steal phenomenon with retrograde flow in the radial artery (in blue), in the segment distal to the anastomosis of arteriovenous fistulas for hemodialysis.

#### Table 20 – Mapping of arteriovenous fistulas for hemodialysis before the procedure.

	Arterial diameter – mm				Venous diameter – mm			
Axillary	Brachial	Radial	Ulnar	Brachial	Radial	Ulnar	Cephalic	Basilic
Proximal arm/forearm								
Mid arm/forearm								
Distal arm/forearm								
Elbow/wrist								
Skin-vessel depth								
Distance A-V								

2. Examination after the procedure (Table 21):

• Describe the type of AVFH and its location; inform if it is functional, dysfunctional, or non-functional, mentioning the cause, location, and extension of the involvement

• Report the afferent and efferent volumes (ml/min).

#### 6. Deep venous thrombosis

#### 6.1. Introduction

DVT and pulmonary embolism (PE) are part of the same disease spectrum: venous thromboembolism (VTE). DVT

Table 21 – I	Mapping of	arteriovenous	fistulas fo	r hemodial	sis after the	procedure

Anastomosis:	Arte	erial diameter – r	nm	Venous diameter – mm				
Axillary:	Brachial	Radial	Ulnar	Brachial	Radial	Ulnar	Cephalic	Basilic
Proximal arm/forearm								
Mid arm/forearm								
Distal arm/forearm								
Elbow/wrist								
Skin-vessel depth								

represents approximately two-thirds of the cases and PE, one-third. Between 85% and 90% of DVT cases occur in the lower limbs.  $^{\rm 83}$ 

VTE is a severe, preventable, and high-incidence disease – the third most common CVD, after acute myocardial infarction and CVA. Therefore, VTE is a serious and potentially lethal condition that can affect both inpatients and outpatients. After the first VTE episode, the chance of recurrence is high.<sup>84,85</sup>

The venous thrombus often begins at the venous cuspid level (Figure 16) – either superficial or deep veins – and extends proximally in 13% of cases, retrogradely in 4%, and in both directions in 10%.<sup>86-88</sup> It can be partial – when occupying part of the lumen of the vein involved – or total. Thrombus located in the superficial system indicates superficial venous thrombosis, while DVT involves the deep venous system, and can strike one or more veins.<sup>89</sup> DVT in lower limbs is considered proximal if it affects the popliteal vein and/or proximal veins, with or without the involvement of other leg veins, and distal if it affects deep infrapatellar veins.<sup>87,90</sup>

The objective examination is crucial as the clinical diagnosis alone is not reliable. The consequences of diagnostic error are severe. At an early stage, it can result in death and, at a later stage, depending on the pathophysiology (obstruction, reflux, or both), it can cause chronic venous hypertension, leading to debilitating conditions, such as post-thrombotic syndrome (PTS) and, in case of pulmonary involvement, pulmonary hypertension.<sup>83,85,91</sup> Furthermore, although effective, the unnecessary use of anticoagulant therapy results in higher costs and risk of hemorrhage.<sup>92</sup>

#### 6.2. Post-thrombotic syndrome

PTS is defined as a combination of symptoms and objective findings in patients with DVT in lower or upper limbs. PTS is a debilitating disease and the most common and less known consequence of DVT.<sup>93,94</sup> One to five years after the DVT episode, approximately 30% to 50% of patients develop PTS, with 5% to 10% of them being severe cases, even when treated correctly.<sup>94,95</sup>

PTS is a combination of venous hypertension secondary to flow obstruction or valvular incompetence and microcirculation and lymphatic abnormalities.

#### 6.2.1. Deep venous thrombosis diagnosis

As the accuracy of the clinical diagnosis of DVT is low (< 50%), few patients with suspicion of DVT effectively have the disease (12% to 31%). Therefore, the recommendation is to perform an accurate and objective examination that can confirm or rule out DVT.<sup>87,92,96</sup> The current gold standard examination to diagnose DVT is compression VUS.<sup>92,97</sup>

Three categories of examinations are used to determine the probability of  $\mathsf{DVT}:^{92,98}$ 

1. Clinical probability based on anamnesis and clinical examination

2. D-dimer levels

3. Imaging studies – the most commonly used is venous VUS, and the less used are venography, CTA, and venous magnetic resonance angiography. This guideline will cover only VUS.



Figure 16 – Two-dimensional image of a recent partial thrombus (acute) located in the valve sinus of the common femoral vein, shown through longitudinal (A) and transverse (B) planes. There is no complete collapse of the vein during compression (C).

#### 6.2.2. Vascular ultrasound

VUS is considered the current gold standard examination to diagnose acute DVT.<sup>99</sup> The PIOPED II study showed 95.5% agreement between CTA and ultrasound in diagnosing or ruling out DVT. In addition, we must remember that VUS is useful not only in diagnosing DVT in symptomatic or asymptomatic (with a high risk of DVT) patients but also in identifying other conditions that cause signs and symptoms indistinguishable from DVT.

Knowledge about vascular anatomy and its variations is crucial since the frequency of variability in the number of veins is high, with the possibility of only one of them being involved, in addition to the different levels of confluences and paths.

To diagnose thrombosis, we adopt several criteria, listed below.

1. Venous compression: the normal vein has thin, smooth, and regular walls, and is completely collapsible with transducer compression (Figure 17).

Incompressibility is the main criterion for DVT diagnosis (Figure 16).

A few conditions might hinder this compressibility, and sometimes special maneuvers are necessary. They are:

• Presence of a strong muscle group in the path to be compressed; for instance, the adductor canal requires compression in the external side of the thigh.

• Neurogenic bladder needs draining through urinary catheterization.

• Proximity to bone structures; for example, compression is limited in infrapatellar veins due to the size of the transducer.

2. Vein caliber: usually the vein has twice the caliber of the adjacent artery. Acute venous thrombosis presents great distension of the vessel wall, with an increase in vein caliber and loss of this relationship. Over time, with the process of recanalization, the vein caliber decreases disorderly in some segments, losing the uniformity usually observed along the vessels. In the most chronic stage, a complete vein retraction might occur, with calibers becoming smaller than those of arteries, at times, hindering their recognition during the examination.

3. Characteristics of the venous wall: parietal irregularities or diffuse parietal thickening, intraluminal trabeculations or synechia, structural valve abnormalities, loss of anatomical relationship with adjacent structures, even if tenuous, which can mean a previous DVT already recanalized.

Intraluminal echoes indicate presence of thrombus and its echogenicity can characterize or not its age.

4. Color imaging: color flow imaging is an important tool for venous thrombosis.

While color flow filling all lumen in longitudinal and transverse planes indicates normality, the lack of color or flaws in filling the vessel can point to partial or complete thrombosis, particularly in recent partial thrombi with low echogenicity.

Flow around the vessel strongly suggests acute DVT. Flow permeating the thrombus indicates recanalization and chronicity of the process.

Take precautions regarding the scale and gain of the equipment, and the proper performance of maneuvers that increase venous flow, generating a color signal.

5. Spectral analysis: the normal venous flow with spectral Doppler is spontaneous and phasic with respiration, increases with distal compression maneuvers, and ceases with Valsalva maneuvers (proximal veins) or proximal compression.<sup>85</sup> The spontaneity of the flow might not be observed in distal veins of patients in the supine position, and distal compression maneuvers and/or mobilization of the limb to displace the blood column become necessary. Loss of phasicity, with continuous flow pattern, is an indirect sign of occlusion or proximal compression. Damped response to distal



Figure 17 – Venous compression maneuver. (A) Artery and vein without compression. (B) Normal vein with total compression. (C) Dilated and incompressible vein, with recent thrombus. (D) Incompressible vein, with old thrombus (chronic). (E) Rethrombosis.

compression indicates obstruction between the compression site and the location of the sample flow.

Retrograde flow occurs when there is valve damage. Thus, valve regurgitation is a pathological sign, which can warn to the presence of previous thrombosis, and valve agenesia or venous reflux secondary to hyperflow should be ruled out.

Table 22 describes VUS characteristics that assist in the identification of stages of venous thrombosis.

Valsalva and inspiration and expiration maneuvers help and add information (Table 23).

#### 6.3. Rethrombosis

High risk of recurrent venous thromboembolic disease persists after DVT treatment, with a cumulative incidence of these complications close to 30% in eight years of follow-up; among them, mortality reaches 30%, mainly due to malignant diseases.<sup>100,101</sup> Relevant predictive factors for risk of recurrence are male gender, thrombus location, and D-dimer.<sup>102</sup> Other less known risk factors are residual thrombus occupying 50% of the vessel lumen diameter after treatment and failure in normalization of compression with VUS.<sup>101</sup>

As the non-invasive diagnosis of recurrence is difficult, some ultrasound criteria can be used, such as:

- Measurement of the residual thrombus mass.
- · Abnormalities in the thrombus extension.

• Ultrasound characteristics of the thrombus – low echogenicity, slight compressibility, presence of tail, adhesion to the wall, and increase in venous diameter ( $\geq 2$  times the contralateral vein, or compared to the diameter of the adjacent artery).<sup>100,101,103</sup>

#### 6.4. Examination technique

Some protocols assess only the proximal segment (femoropopliteal) or the compression VUS of two points (common femoral vein and popliteal vein). These protocols, known as point of care, facilitate the examination by emergency physicians and are proving to be an alternative in the emergency room.<sup>106</sup> However, evaluating the entire venous system is important for a proper DVT diagnosis, better assessment in case of recurrence, and to assist in the differential diagnosis with other pathologies. This guideline recommends always performing a full examination.<sup>107,108</sup>

In lower limbs, the patient should be in a comfortable supine position with the torso and head elevated up to 30°, close to the edge of the bed, on the same side of the examiner, with a slight lateral rotation of the hip and slight knee flexion.

In upper limbs, the patient should be in the supine position, with the limb stretched alongside and slightly away from the body.

Examine the deep venous system starting with the inguinal fold, gently compressing the veins with the transducer, using transverse planes.<sup>85,104,105</sup> The goal is to confirm the absence

## Table 23 – Venous flow variation according to the phase of the respiratory cycle.

Flow	Inspiration	Expiration
Lower limbs	$\downarrow$	↑
Upper limbs	↑	$\downarrow$
Subclavians	↑	$\downarrow$

Stage	Acute	Intermediate (subacute)	Chronic	
Event time	Up to 14 days	14 to 28 days	> 28 days	
Vessel caliber	Dilated	Still dilated, normal, or slightly reduced	Usually reduced	
Incompressibility	Total or partial	Total or partial	Partial or absent	
			Hyperechoic	
Thrombus aspect Hy			Parietal irregularities	
	Hypoechoic	Isoechoic	Fibrotic residues	
			Residual thrombus: caliber > 2 mm after maximum compression in 6 months or > 3 mm in two consecutive examinations	
			Multiple flow channels permeating the thrombus	
Flow 4			Complete lumen filling	
	Absent or partial	Flow channels permeating the thrombotic mass	Presence of collaterals	
			Continuous flow or reduced amplitude spectrum	
			Can have orthostatic reflux	

Table 22 – Characteristics of different stages of deep venous thrombosis observed in vascular ultrasound.

or presence of recent or residual thrombi, every 5 cm, throughout the venous trunk in the femoropopliteal segment by the medial thigh, the infrapatellar segment for the popliteal and trifurcation, the posteromedial leg for calf muscle veins (gastrocnemius, soleus, intergemellar), posterior tibial and fibular veins up to the ankle, as well as superficial veins (great and small saphenous veins along its entire extension).

The fibular and soleus veins can be seen in the lateral leg, using the fibula as an anatomical reference. In this case, fibular veins are more superficial in relation to the transducer and posterior tibial veins, deeper. To that end, the patient should change positions, flexing the knee and placing the foot on the bed. Do not neglect this maneuver even in hospitalized patients due to the high prevalence of DVT in segments proximal to fibular veins in this group of individuals.

Anterior tibial and foot veins are rarely studied unless there is a sign of local involvement.

Investigate iliac veins when the patient has DVT in common femoral veins, and it is not possible to identify the end of the thrombus, or when the flow detected in these veins is continuous or has low amplitude.

Alternate between transverse and longitudinal images using color imaging to evaluate flow and, when necessary, spectral Doppler.

Save all stages of the examination (videos and still images) digitally. Currently, clinical research protocols of international multicenter studies recommend recording the maneuver of compressing the vein – with and without compression – in the same picture (dual image). In case of thrombus, measure the venous diameter, in determined sites, to evaluate the residual thrombotic mass.<sup>107</sup>

Follow the same evaluation protocol for deep and superficial venous systems in upper limbs, middle and distal segments of subclavian veins, and jugular veins.

During the follow-up of patients with DVT and recanalization, they preferably should stay in the orthostatic position or, when not possible, sitting for the study of venous reflux.

Do not forget that the temperature in the room can significantly influence the performance of these examinations. Cold induces vasoconstriction and should be avoided. The ideal temperature is 22 to  $25^{\circ}$ C.

#### 6.4.1. Transducers

In general, high-frequency (5 to 12 MHz) linear transducers are used for normal-weight and thin patients. In overweight/ obese patients, the study of the adductor canal and even of leg veins in individuals with moderate/severe edema can use transducers of greater depth range, such as convex abdominal ones, whose frequency varies from 3 to 5 MHz.<sup>108</sup>

#### 6.4.2. Information for the report

• Presence or lack of signs of deep and/or superficial venous thrombosis.

• Information about the characteristics of the thrombus.

• Extension of thrombosis: essential and, if possible, with approximate measurements of anatomical reference points,

such as the anterior-superior iliac spine, inguinal fold, knee, malleolar or plantar region, axillary fold, elbow fold, and wrist fold.

• In case of chronic thrombosis:

- Measurements of residual thrombotic masses can be included in the report and/or images, with their locations so that they can be compared later.

- Presence of orthostatic reflux.

#### 7. Transcranial doppler

The main purpose of the study called transcranial Doppler (TCD) is to gather hemodynamic information from intracranial trunk arteries non-invasively, using flow insonation with pulsed wave Doppler.<sup>109,110</sup> The skull has always represented a barrier to reach the vessels since US does not penetrate the calcium in bone tissue and there are limited areas (transorbital and transtemporal windows, and foramen magnum) for examination. Also, arteries are located in deeper parts of the brain, making it difficult to obtain proper images and safely collect a sample capable of providing the spectral curves needed to interpret the hemodynamic status registered at a given moment. These unfavorable characteristics determine that a transducer capable of insonating flow in all trunk arteries of anterior and posterior circulations needs to be small and have a low frequency (2.0 MHz or less). CFI safely identifies vessels and analyzes flows.

#### 7.1. Types of transcranial doppler

• "*Blind" TCD*: only uses transducer with pulsed wave Doppler, without B-scan image (Table 24).

• *TCD with color Doppler*: transducer with image from B-scan, pulsed wave Doppler, and CFI associated.

• *TCD with microbubble contrast*: color TCD associated with intravenous infusion of microbubble contrast.

• *TCD with macrobubbles*: color TCD associated with peripheral intravenous infusion of saline or glucose solution mixed with ambient air and shaken (macrobubbles).

#### 7.2. Examination technique and protocol

The main focus of any TCD examination must be identifying all trunk arteries safely and recording the spectral flow curve of each vessel. Therefore, the usefulness of CFI is indisputable. Currently, using devices exclusively dedicated to "blind" Doppler is warranted in the monitoring of continuous flow during surgeries and in neurointensive units.

The basic TCD protocol should include the use of all possible windows to transmit US to intracranial arteries:<sup>109</sup> a) *transorbital (right and left)*, to insonate carotid siphons and ophthalmic arteries; b) *transtemporal (right and left)*, to visualize distal internal carotids, anterior cerebral (A1 and A2 segments), middle cerebral (throughout M1 segments and at the beginning of M2), top basilar, and posterior cerebral arteries (throughout P1 and P2 segments); c) *transforaminal*, to study V4 segments of vertebral and basilar arteries (proximal and middle segments), with the possibility of insonating posterior-inferior cerebellar branches of vertebral arteries.

The patient can be in a supine position or sitting. The head should be turned according to the window used at the time. The examiner can follow any sequence and must record images of spectral flow curves of each vessel examined – with identification –, according to the international standard. Including pictures or videos done with CFI ensures safety in confirming identification and possible anatomical variants (very common) or technical difficulties during the examination.

Visualizing brain structures with B-scan determines the ability of the US in penetrating the bone wall in the window chosen and its usability for the examination. CFI will show (or not) flow in lumens of regional arteries, guiding the sample volume of pulsed wave Doppler with precision up to the lumen segment to be insonated to obtain the spectral flow curve. In addition to flow waveform morphology of each vessel, the routine of integral measurement of velocities has to be followed to ensure the collection of essential data to the hemodynamic analysis required in various pathologies with suspicion of DTC: PSV, EDV, mean velocity, RI, and PI (make sure that the equipment preset includes these calculations).

Table 25 lists the clinical indications for TCD.

\*According to the **Brazilian Guidelines for the Application** of Transcranial Ultrasound as a Diagnostic Test for the **Confirmation of Brain Death** (defined by a group of experts from the Department of Neurosonology of the Brazilian Academy of Neurology in 2012), the criteria<sup>122</sup> are:

• A single TCD is enough to confirm brain death.

• TCD should be performed only in patients with previously established clinical diagnosis, in accordance with the current Brazilian regulations; moreover, the patient needs to have stable hemodynamic conditions (with or without the use of drugs), and minimum systemic systolic blood pressure of 90 mmHg (below this value, the examination will not have diagnostic validity).

• A full standard TCD study is mandatory, with images in B-scan and color flow imaging (if available), and spectral flow curves of all intracranial trunk arteries.

• Characteristic TCD findings in *cerebral circulatory arrest*: spectral flow curves with low-amplitude systolic wave (velocity < 50 cm/s) or curves with an *alternating flow* pattern (waves with antegrade systolic component followed by retrograde diastolic component).

• Lack of flow in intracranial trunk arteries is not a criterion for brain death, except in cases in which a prior TCD was performed during the same hospitalization, with a record of flow in the arteries analyzed.

• Anterior circulation: in case of inadequate transtemporal windows, reporting criteria for "vascular collapse" in both carotid siphons becomes mandatory to diagnose brain death.

• Posterior circulation: if it is not possible to detect flow in the basilar artery, findings of "vascular collapse" in both intracranial vertebral arteries are crucial to diagnosing brain death; on the other hand, findings related to basilar "vascular collapse" in the presence of residual blood flow in at least one vertebral artery will invalidate the conclusive diagnosis of the examination as an indication of brain death in this region.

• Residual blood flow can be detected in almost 20% of patients, especially in intracranial carotid arteries and in patients who underwent craniotomy (but this finding tends to disappear in a few hours).

• The examination report must have a detailed account of the findings in each anterior and posterior circulation artery and be conclusive regarding the presence or lack of criteria for *cerebral circulatory arrest* that corroborate the clinical diagnosis of brain death.

#### \*\*\*Spencer scale

The number of embolic spikes shown in the device screen is also useful information since the greater the number of macrobubbles, more significant the size of the shunt(s) through the foramen ovale (Spencer scale): grade 0 - lack of HITS; grade 1 - 1 to 10 HITS; grade 2 - 11 to 30 HITS; grade 3 - 31to 100 HITS; grade 4 - 101 to 300; grade 5 - > 300 HITS ("curtain effect"). Above grade 2, the right-left cardiac shunt is significant. In case of countless spikes ("curtain effect"), consider the possibility of pulmonary AVF.

Examination protocol: peripheral intravenous infusion of solution with "macrobubbles" (8 ml of saline or glucose solution mixed with 2 ml of ambient air and shaken until it becomes homogeneous), followed immediately by vigorous Valsalva maneuver performed by the patient for 5 seconds and simultaneous insonation of spectral flow curves (pulsed wave Doppler) in cerebral and basilar arteries. The examination must be conducted in basal conditions (flow record with Valsalva maneuver in the right and left middle

Table 24 – Identification of intracranial trunk arteries with "blind" transcranial Doppler<sup>114</sup>.

Artery	Depth	Flow Vm	Flow direction in relation to the transducer
Carotid siphon	55 to 70 mm	40 to 50 cm/s	Positive or negative
Ophthalmic	40 to 60 mm	20 cm/s	Positive
Distal internal carotid	55 to 70 mm	45 cm/s	Positive
Anterior cerebral	60 to 70 mm	60 cm/s	Negative
Middle cerebral	35 to 60 mm	70 cm/s	Positive
Posterior cerebral	55 to 70 mm	40 cm/s	Positive (P1), negative (P2)
Vertebral	55 to 70 mm	40 cm/s	Negative
Basilar	70 to 120 mm	45 cm/s	Negative

#### Table 25 – Clinical indications for transcranial Doppler<sup>115,116</sup>.

Pathology	Objective	Observation	Findings
Brain death*	Additional examination judicially valid to confirm cerebral circulatory arrest	Standard study of anterior and posterior circulation arteries is mandatory. In case of residual blood flow, repeat after 12 hours	Spectral curve with short peak systolic (< 50 cm/s) pattern and lack of diastolic flow; or "alternating" pattern (antegrade systolic and retrograde diastolic flow)
Intracranial hypertension <sup>111</sup>	Adjuvant indirect monitoring, including after decompressive craniectomy	Qualitative analysis of the curve pattern, which can vary dynamically (inversion of the diastolic component delimits the irreversible stage)	Progressive reduction of the diastolic component of the spectral flow curve according to the severity of hypertension
Ischemic CVA (acute phase) <sup>112,113</sup>	Monitoring of vessel reperfusion in case of thrombolysis (up to 4.5 hours after the start of the event), which lasts approximately 40 minutes (but can take more than 1 hour)	Monitoring can be intermittent (conventional equipment) or continuous (transducer with "blind" Doppler fixed to a helmet adjustable to the patient's head)	Reappearance of gradual flow according to the degree of reperfusion (TIBI scale of spectral curve pattern)
Subarachnoid hemorrhage** 114 (Table 26)	Diagnosis, assessment of severity, and monitoring of vasospasm, recommending early intervention	Perform the examination at hospital admission and repeat it daily in case of vasospasm (critical period: 4 to 14 days after the event). Insonate all arteries at each examination	Increase in mean flow velocity, according to the severity. Lindegaard ratio (velocity ratio between the middle cerebral artery and ipsilateral internal carotid) differentiates true spasm from hyperemia
Patent foramen ovale*** 115	Shunt study in patients with ischemic CVA (transient or permanent)	Intravenous infusion of shaken saline solution ("macrobubbles") associated with the Valsalva maneuver	HITS (gaseous emboli) recorded in spectral curves are counted and classified according to the Spencer scale
Sickle cell disease**** <sup>116-118</sup> (Table 27)	Diagnosis and grading of intraluminal stenosis to stratify the risk of ischemic CVA and define the therapeutic approach. Monitoring of therapeutic response	Mandatory in patients aged 2 to 16 years	Mean flow velocity defines the periodicity of follow-up and approach (blood exchange)
Migraine <sup>119</sup>	Support for clinical diagnosis and differentiation from other headaches	It can be performed in the intercritical period or during a painful crisis (different results)	Measurement of pulsatility index and mean flow velocity in all vessels
Transoperative monitoring <sup>120,121</sup>	Preoperative assessment of CVA risk (monitoring of spontaneous microemboli; study of cerebral flow reserve) and peroperative monitoring of emboli and cerebral flow reduction during neurological and cardiovascular surgeries	Continuous flow monitoring of middle cerebral arteries using two transducers with "blind" Doppler fixed to a helmet adjustable to the patient's head. Monitoring should continue in the postoperative (due to microemboli during this period)	Report of emboli rate (solid and/or gaseous) and reduction in mean flow velocity compared to the baseline value (> 15%) determine the risk of ischemic CVA in the immediate postoperative period*****

CVA: cerebrovascular accident; HITS: high-intensity transient signals; TIBI: thrombolysis in brain ischemia (scale). Information followed by asterisks (\*) have additional data, presented in the text below.

cerebral arteries and basilar arteries) and after the infusion of macrobubbles (that is, six times in total). In the event of a record with "curtain" pattern, stop the study (which will be considered positive).

\*\*\*\*\*Microembolic signals are detected in up to 70% of cases during the first hour after endarterectomy. A rate of 50 "microembolic signals"/hour occurs in up to 10% of cases and is predictive of ipsilateral focal ischemia.<sup>121</sup>

#### 7.3. Limitations of transcranial color doppler

Limitations of TCD basically result from the barrier the cranial bone represents to US. The use of contrast agents ("microbubbles") greatly reduced the cases of inconclusive examinations due to a "lack of adequate windows." The inexperience of the examining physician is also a crucial limiting factor; the learning curve is relatively long and requires dedication.

# 7.3.1. Essential information to include in transcranial doppler reports

The basic structure of any additional examination must have:

• Patient's identification (full name and age).

• Clinical indication (the purpose of the examination will determine the type of TCD needed).

• Technical quality of the examination (reporting possible issues that interfere in obtaining the necessary images for the study).

• Record of all ultrasound windows used, and vessels examined (justifying the cases that were impossible to study).

• Description of specific characteristics found in each technical resource used:

- Color flow imaging – lumen patency or occlusion, laminar or turbulent ("mosaic") pattern, direction (antegrade or retrograde).

- Pulsed wave Doppler - laminar or turbulent pattern,

#### \*Table 26 - Vasospasm classification.

Artery: MCA	MFV (cm/s)	Lindegaard ratio
Mild	120 to 130	3 to 6
Moderate	130 to 200	3 to 6
Severe	> 200	> 6
Artery: ACA	> 50% of MFV in 24h	-
Artery: PCA	> 110	-
Artery: VA	> 80	-
Artery: basilar		Soustiel ratio
Mild	80 to 95	2 to 2.49
Moderate	> 85	2.5 to 2.99
Severe	> 115	> 3

MCA: middle cerebral artery; ACA: anterior cerebral artery; PCA: posterior cerebral artery; VA: vertebral artery. Note: Lindegaard ratio is the ratio between the highest mean flow velocity (MFV) in the middle cerebral artery (M1) and MFV in the ipsilateral extracranial internal carotid. Soustiel ratio is the ratio between vertebral MFV and basilar MFV.

direction (antegrade or retrograde), velocities of spectral flow curves (peak systolic, end-diastolic, and mean), and PI and resistance according to the clinical indication of the examination (e.g., sickle cell disease requires mean velocity).

- Contrast solution with "macrobubbles" – record and count of HITS in the study of patent foramen ovale and paradoxical embolism.

- Use of contrast agent with "microbubbles" (SonoVue®).

- In the intraoperative monitoring, report the occurrence and count of HITS/hour and variations in flow velocities with potential risk for ischemic CVA in the postoperative period.

**Conclusion**: avoid etiological diagnosis (TCD is a study of cerebral hemodynamics and findings should be correlated to the clinical indication of examination: for instance,

Low flow

Normal

"Low conditional"

"High conditional"

Abnormal

Abnormal

the presence or lack of criteria for vasospasm, with grade classification and time reference to the onset of symptoms of subarachnoid hemorrhage).

Note: in cases of sickle cell disease, it is mandatory to specify, after the conclusion, the recommended date to repeat the reassessment examination (following the Brazilian Guidelines for Transcranial Doppler in Children and Adolescents with Sickle Cell Disease, 2010).<sup>123</sup> In other pathologies, this suggestion is prohibited due to professional ethics.

#### 8. Contrast in vascular ultrasound

The introduction of contrast agents in ultrasound examinations considerably expanded the clinical value of this method. B-scan ultrasound is an excellent method to demonstrate the anatomical and morphological characteristics of parenchymal tissue but does not provide any information about the viability and integrity of microcirculation. Color Doppler ultrasound (CDUS) added data about blood flow velocity but is not able to quantify the volume and show with enough sensitivity the blood flow in microcirculation,124-126 where velocity is too low to be detected without artifacts created by tissue and transducer maneuvers. Also, the spatial resolution of CDUS is limited, the representation of velocity is angle-dependent, and numerous artifacts can influence the diagnostic interpretation of images. Contrast-enhanced Doppler ultrasound substantially increased the sensitivity of CDUS and reduced some limitations.

The technical innovation was the introduction of specific image components for contrast in US equipment, allowing direct visualization of signals emitted by microbubble contrast agents, regardless of their velocities. Due to typical characteristics of microbubble signals (which are fundamentally different from those originating in tissues), "specific microbubble" images (contrast only) are created, which can show volume and perfusion of the parenchymal tissue with extremely high sensitivity and spatial resolution.<sup>124</sup> The creation of each "specific microbubble" signal requires

Repeat the examination in 30 days

Repeat the examination in 12 months

Repeat the examination in 3 months: if < 170 cm/s, repeat in 12 months Repeat the examination in 30 days: if < 170 cm/s, repeat TCD every 3 months; if two

consecutive examinations are abnormal, consider long-term blood exchange Repeat the examination in 30 days: if > 200 cm/s, blood exchange; if "high conditional",

repeat TCD in 3 months; if "low conditional", repeat it in 6 months

Imminent risk of CVA, consider long-term blood exchange

Mean flow velocity	CVA risk group	Approach
Inadequate ultrasonic windows	Inconclusive	Use another imaging method to assess cerebrovascular events
Difficult execution: uncooperative patient	Inconclusive	Repeat in 3 months, with a different examiner if possible

\*\*\*\*Table 27 – Recommendations of the Brazilian Guidelines for Transcranial Doppler in Children and adolescents with Sickle Cell Disease (2010)<sup>123</sup>.

CVA: cerebrovascular accident; TCD: transcranial Doppler.

< 70 cm/s

< 170 cm/s

170 to 184 cm/s

184 to 199 cm/s

200 to 220 cm/s

> 220 cm/s

proper interaction between signals of the microbubble contrast agent and the insonated US beam.

#### 8.1. Properties of ultrasound contrast agents

Unlike contrasts for magnetic resonance imaging (MRI) and computed tomography, which use physical and chemical characteristics of cells to generate their effect, microbubble contrast uses the physical attributes of US, that is, the higher the density difference between media, the greater the reflection of energy emitted and larger the amplitude of US signal. Unquestionably, the gaseous medium provides the greatest difference, corresponding to a signal increase of approximately 30 decibels.

US contrast agents are microbubbles of gas inside capsules with flexible and stable phospholipid membranes and a defined size. The SonoVue  $\mathbb{R}^{127}$  agent (produced by Bracco Imaging S.p.A., Milan, Italy), sold in Europe, the United States, parts of Asia, and South America, is the only product currently authorized in Brazil by the Brazilian Health Regulatory Agency (Agência Nacional de Vigilância Sanitária – Anvisa) and members of the National Regulatory Agency for Private Health Insurance and Plans (Agência Nacional de Saúde Suplementar – ANS). SonoVue® consists of encapsulated microspheres of sulfur hexafluoride gas. Microbubbles have an average diameter of 2.3  $\mu$ m (a size that prevents them from crossing blood vessel walls and reaching the interstitial space). As a lipophilic gas, its solubility in blood is low, and it does not spread outside the capsule. This protein shell composed of a single layer of phospholipids acts as a surfactant, giving it stability and flexibility along its path in the blood macroand microcirculation. Therefore, SonoVue® is considered an integral agent of the blood pool and a marker of blood circulation (property that distinguishes it from contrasts used in MRI and computed tomography, which can cross into the extracellular space).

After the microbubble bursts, the gas is almost entirely exhaled through the lungs during respiration, without undergoing liver metabolism or renal excretion.<sup>124</sup> Thus, there is no contraindication to its use in patients with renal failure.

# 8.2. Technical aspects that influence the acquisition of contrast images

Currently, most US manufacturers have exclusive software for studies with contrast, which can be included in the original configuration of the machine or purchased separately. However, even equipment without a specific image component for contrast has some parameters the operator can configure. The following concepts and adjustments of the equipment must be known to get the best result during the contrast study:

#### 8.2.1. Mechanical index

A unique characteristic of contrast agents for US (nonexistent in those used for MRI and computed tomography) is that they are modified by the waves used to detect them. When exposed to US, microbubble behavior changes according to the US power emitted, that is, the amplitude of the acoustic wave [called mechanical index (MI) in the equipment]. In non-contrast studies, MI ranges from 1.6 to 1.9; under this acoustic power, the microbubble always oscillates vigorously and bursts, causing two unwanted effects: a sudden increase in signal intensity with an excessive blurring of the image, and a significant reduction in contrast concentration, consequently shortening the examination time. This image mode, called "image by acoustic stimulation," does not require equipment with contrast technology (but, on the other hand, it does not use the full potential of the contrast agent, limiting it to its echo-enhancer function).

After reducing MI to  $\leq 0.1$ , we can not only keep the integrity of microbubbles but also make them oscillate nonlinearly (initial compression followed by expansion) and resonate, emitting different frequencies (known as "harmonic frequencies") from the fundamental one emitted by the transducer. Equipment with this technology can filter signals emitted specifically by microbubbles, leading to a more lasting study that highlights microbubble over tissue signal (virtually nulled in the image, appearing as a dark background).

This type of study, also called "contrast study with low MI", allows the examiner to evaluate the contrast wash-in continuously in the local studied, the enhancement period, and the concentration of microbubbles in the target structure [essential for situations such as the study of vessels of vessels (*vasa vasorum*), carotid plaques, renal capillary (perfusion) distribution, and masses].<sup>124</sup>

An undesired effect of the contrast study with low MI is the depth limitation of the pulse wave, which undergoes greater attenuation as it moves through tissues. Some ways to minimize this effect are: adopting alternative acoustic windows that allow closer proximity with the structure of interest, using broadband transducers with lower frequencies, and, as a last resort, increasing the MI and consequently destroying more bubbles in the proximal field.<sup>128</sup>

#### 8.2.2. Image gain

A noteworthy equipment control in the contrast study is the image gain, which amplifies the signal received during the post-processing in the equipment. High gain produces a bright image and widespread increase in background noise, obscuring the contrast signal (once the saturation level of the equipment has been reached, there will be no margin to amplify the signal caused by contrast). Therefore, during the contrast study, the equipment gain should be reduced until the image is virtually black, except for highly echogenic structures. Some manufacturers have gain adjustment controls for contrast studies that can easily be turned on and off during the examination.

When performing a manual adjustment, make sure to have the least amount of acoustic signals before contrast injection and understand if this signal results from an increase in MI (when specific structures are seen in the image) or gain (which causes a widespread rise in noise across the image).<sup>6</sup>

#### 8.2.3. Contrast level

A last technical aspect worth mentioning is the dose of contrast injected. High doses initially blur (saturation) the

signal and attenuate (acoustic shadowing) structures in the distal field until contrast concentrations drop to a proper level. In addition, distinguishing small enhancing differences between structures is not possible, given that the upper limit of dynamic range (grayscale) of the equipment has been exceeded.<sup>128</sup> One way of highlighting different enhancing levels caused by contrast in a structure is to adjust the dose of contrast to allow adequate opacification, without blurring or attenuation, and increase the dynamic range of the equipment. In turn, low doses will not reach the opacification level desired.

#### 8.3. Indications for contrast in vascular ultrasound

Table 28 lists the main indications for microbubble contrast agents in US studies of different vascular systems (consensus of the European Federation of Societies for Ultrasound in Medicine and Biology, published in 2011).<sup>129</sup> The levels of evidence, based on multicenter and/or single-center studies, were classified as A (good), B (moderate), and C (recommended by expert consensus).

# 8.4. Preparation, Application, and dose of microbubble contrast

SonoVue<sup>®</sup> is a kit that includes: a vial with 25 mg of lyophilized powder in an atmosphere of sulfur hexafluoride; a syringe filled with 5 ml of sodium chloride 9 mg/ml (0.9%) solution; a transfer system. The contrast is easy to prepare at the bedside, following the manufacturer's instructions. After transferring the content of the syringe to the vial with powder, shake it for 20 seconds to create microbubbles and transform the saline solution into a milky suspension (indicating homogeneous microbubble distribution). In this state, the suspension can be stored for up to 6 hours. If microbubbles accumulate on the surface during rest, shake the solution again, so the microbubbles regain homogeneous distribution before use.

The usual route of administration is by intravenous bolus injection in a vessel with caliber suitable for puncture with a needle of 20G in diameter (preferably in the antecubital fossa). Administer a small initial volume, followed by a flush of 5 ml of saline solution at 0.9% to push the contrast agent into the central vein (which happens in seconds).

In most publications, the recommended dose for a single injection in VUS studies is 2.4 ml, ranging from 1 to 4.8 ml, according to the organ studied, the probe used, and the sensitivity of the equipment available (always remembering that probes with higher frequencies need higher doses, in this case, 4.8 ml).<sup>7</sup> The first 10 to 40 seconds after bolus injection correspond to the contrast enhancement curve (wash-in and wash-out) and should be continuously recorded for later analysis. In some specific situations, such as the study of late endoleaks, the period of evaluation can reach 5 minutes; in these cases, shorter videos can be recorded. Bear in mind that the higher the MI, the greater the destruction of bubbles, and shorter the duration of contrast. After the bubbles burst, the lungs quickly (2 minutes) and fully eliminate the sulfur hexafluoride (Anvisa).

SonoVue<sup>®</sup> is a safe agent, with a low complication rate. Reports of anaphylactic reaction correspond to < 0.002% of cases.

#### Table 28 – Indications for contrast agents in vascular ultrasound

System	Application	Level of evidence	Probe	
Carotids	Occlusion or subocclusion In-stent stenosis Dissection Plaque neovascularization	B B C B	Linear	
Aorta	Dissection, extension of the flap to branches, false lumen patency, re-entry points, and perfusion level of branches that originate from false lumen	С	Convex abdominal or sector cardiac	
	Differentiation between an inflammatory aneurysm and contained rupture Endoleak	B		
Intracerebral vessels	Signal increase in unsatisfactory basal study Perfusion in ischemic CVA	A C	Sector cardiac	
Complications in vascular accesses	Arteriovenous fistulas Pseudoaneurysms, hematomas	C C	Linear or convex abdominal or sector cardiac	
Kidney	Signal increase in the renal artery Renal perfusion	C C	Convex abdominal or sector cardiac	
Lower limbs	Obstructive atherosclerotic disease (assessment of collateral circulation and microcirculation)	С	Linear or convex abdominal	
	Deep venous thrombosis (signal improvement and inflammatory reaction in perithrombus)	С		

CVA: cerebrovascular accident.

# 8.5. Basic protocol for vascular ultrasound with microbubble contrast

After defining the indication for microbubble contrast in VUS examination, follow the mandatory basic routine:

• Repeat and record the standard CDUS examination of the organ of interest

• Secure venous access for injection of contrast solution with microbubbles (peripheral vein puncture or deep vein already in use)

• Prepare the contrast solution with microbubbles (SonoVue®) following the manufacturer's instructions

• Activate the specific image component for contrast in US equipment; if there is no specific software, adjust MI (< 0.6 and the closest possible to 0.1), image gain (darken the background), and choose the appropriate windows to reduce the depth of the target-organ under study

• Administer the contrast solution, make adjustments to reduce excessive enhancement, and record digital images (videos) for 10 to 40 seconds after the initial bolus injection; in specific longer examinations, record the necessary videos during the procedure (which can reach 5 to 8 minutes) for later analysis.

**Note**: analysis of examination with microbubble contrast is fundamentally dynamic and the duration of the study is

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short due to the fast destruction of microbubbles by US waves, even when using very low MI. Thus, recording it in digital media is essential for later processing and careful review of images, ensuring a safe diagnosis and permanent storage of results.

# 8.6. limitations of the use of contrast in vascular ultrasound

• Inexperience of the examiner (proper training in ultrasound with contrast is crucial for an effective and safe diagnosis).

• Equipment without a specific software for image with contrast makes it difficult (but not impossible) to conduct a conclusive examination.

• Access to microbubble contrast in units of the Brazilian public health system.

• Complete lack of "window" that allows US transmission to the organ of interest (rare).

• Hypersensitivity to microbubble contrast (rare).

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