ABSTRACT
The aim of this review is to describe the acquisition and reformattting of state of the art high resolution cone beam CT (HR-CBCT) and demonstrate its role in multiple neurovascular conditions as a tool to improve the understanding of disease and guide therapeutic decisions. First, we will review the basic principle of CBCT acquisition, followed by the injection protocols and the reformattting paradigms. Next, multiple applications in different pathological conditions such as aneurysms, arteriovenous malformations, dural arteriovenous fistulas, and stroke will be described. HR-CBCT angiography, widely available, is uniquely useful in certain clinical scenarios to improve the understanding of disease and guide therapeutic decisions. It rapidly is becoming an essential tool for the contemporary neurointerventionalist.

INTRODUCTION
Clear understanding of the complex angioarchitecture and vasculature in relation to intracranial bony structures is essential in endovascular clinical decision-making. Two-dimensional digital subtraction angiography (2D-DSA) is the standard imaging technique for the evaluation of cerebrovascular conditions, and has been the mainstay of diagnostic cerebral angiography since its inception. Since the introduction of rotational DSA with three-dimensional (3D) reconstruction in the late 1990s, it was quickly recognized as giving paramount additional information for the evaluation of neurovascular lesions, leading to its routine adoption. Multiple subsequent variations were developed, including 3D-3D fusion, 4D DSA, and arterial and venous (AV)-3D-DSA.
All volumetric datasets acquired by modern angiographic machines are based on the cone beam CT (CBCT) technique. While commercial and marketing considerations produce various proprietary names, reflecting the suggested purpose of a particular algorithm (DynaCT, 3D-DSA, VasoCT, Innova CT for GE), to which we refer in this paper as HR-CBCT to distinguish it from the above described 3D-DSA. It is again emphasized that both are based on the same acquisition principles, and differ only in various trade-offs, such as speed of acquisition, resolution, radiation dose, etc.

The 3D-DSA acquisition achieves comparatively low soft tissue resolution due to the relatively low number of projection images (n=133), resulting in more stripe artifacts, and a low detector dose level, contributing to a relatively low signal-to-noise ratio. Advantages of the technique, however, include faster acquisition, allowing for better arteriovenous phase separation and lower X-ray exposure. DynaCT differs from the 3D-DSA acquisition mode, resulting in higher resolution, due to...
the significantly higher number of projection images (n=500) acquired, and the higher detector dose used, both factors contributing to a much better resolution, at the expense of longer acquisition times and higher radiation exposures.

Vendors generally provide two HR-CBCT (DynaCT) protocols, a low and a high kV option. CBCT protocols targeted to visualize iodinated contrast media have their kVp set to lower values, typically 70 kV, closer to the k-edge of iodine (33 keV) to maximize x-ray absorption. For more challenging CT scenarios, such as the detection of soft tissue or hemorrhage adjacent to the skull without the application of contrast media, a higher kVp (109 for Siemens) is advantageous as this reduces the beam hardening artifacts at the skull. The higher kVp HR-CBCT protocol is also best suited for imaging high density structures such as metallic implants or bone. In practice, however, many factors seem to contribute to the final appearance of the image, with 109 kVp acquisitions appearing to often yield superior results in terms of image quality compared with 70 kVp for all HR-CBCT applications in our experience.

Detector readout bandwidth limitations sometimes make it necessary to combine detector pixels (making 4 pixels act as 1) to keep the acquisition times within an acceptable range. This comes at the expense of spatial resolution. Alternatively, to maximize the spatial resolution, a small (22 cm typically) detector area can be used, which activates the acquisition in a non-binned mode (called DynaCT Micro in Siemens), as opposed to the 2×2 pixel binning (combining pixels) typically used for acquisitions with a larger FOV. The caveat of using a higher magnification is that the proceduralist must more carefully set up the acquisition to ascertain the inclusion of the area of interest in the middle of the FOV.

**INJECTION PROTOCOLS**

The typical protocols are to inject full contrast for arterial vessel imaging (such as 3D-DSA) and dilute contrast (20–30% contrast in saline) for post-implant evaluation. However, longer full contrast injections such as during DynaCT can be extremely
useful for dramatically enhancing the resolution of vascular imaging.

It is our experience that an ad hoc approach works best when selecting optimal injection protocols. A few factors need to be taken into consideration when deciding what flow rate/volume/injection delay protocols are optimal. The aim is to have full opacification of the target vessel throughout the entire acquisition without any remarkable inflow of non-opacified blood.

Factors to consider are: (1) the size of the vessel in which the injection catheter is located; (2) the distance between the target anatomy and the catheter location; and (3) the acquisition protocol chosen. These factors will imply what flow rate/volume/delay injection rate should be used. For example, in a typical situation where the catheter is located in a nominally adult-sized distal cervical internal carotid artery (ICA), and the aim is to characterize the intracranial ICA using a 20s acquisition CBCT, a flow rate of 3 mL/s is usually sufficient to obtain full vessel opacification. Given the distal cervical location of the catheter and assuming a normal heart rate and cardiac output, 1–1.5 s delay from the start of the injection to the beginning of image acquisition typically is enough to visualize the ICA territory in its entirety. The delay chosen can be more carefully refined using 2D-DSA imaging previously obtained as part of routine angiography. Since a 20s acquisition protocol is chosen, one needs to make sure that the vessel of interest is opacified throughout the entire acquisition. This duration requirement will drive the calculation of the contrast volume to inject, which for this particular case, equates to a volume of $3 \times 21 = 63$ mL of undiluted contrast. In terms of pressure, our standard is 300 PSI (pounds per square inch) but that can be adjusted based on the catheter used and the catheter location.

A specific situation arises in the case of HR-CBCT angiography obtained through a microcatheter injection, usually during cranial or spinal vascular malformation or during tumor embolization procedures. In these cases the PSI rating of the microcatheter must be accurately confirmed to verify the safety of the injection, and we usually use a flow rate of 0.6 mL/s (with a volume of 12 mL for a 20s CBCT). Alternatively, sometimes injections are done by manual injection with appropriate shielding of the operator.

**REFORMATTING PARADIGM: SMALL VOLUME OF INTEREST, MAXIMUM INTENSITY PROJECTION, AND VOLUME RENDERING**

The final image quality is dependent not only on the acquisition protocol, but on the manner in which the source images are reformatted.

The reconstruction and data processing/display protocols vary and are highly proprietary. Standard options include volume rendering and maximum intensity projections (MIPs)
for volumetric images, and MIP/averaged projections for cross-sectional datasets.

While large FOV acquisition and large volume of interest (VOI) reconstruction can generate beautiful and detailed images, it is by reducing both that the potentials of CBCT are fully realized. Indeed, while a 48 cm FOV with a full VOI generates a volume with a voxel size of 0.5 mm, use of a smaller 22 cm FOV coupled with a small VOI reconstruction is capable of generating images with a voxel size up to 0.02 mm. Such small VOI reconstructions are essential for the accurate detailed study of small branches such as small perforators, retinal branches, spinal arteries and so forth, as will be demonstrated below with practical scenarios. For the reconstruction of the volumetric data as axial images a Hounsfield unit (HU) versus edge enhancement (EE) kernel and a reconstruction image characteristic (normal, sharp or smooth) can be chosen. It is our experience that the best algorithm varies patient by patient rather than being the same for the corresponding structures, possibly in relationship to heterogeneities introduced by additional factors, such as skull thickness, or patient motion artifact—best curbed by use of apnea during general anesthesia—throughout image acquisition. Once the volume is reconstructed, it can be visualized using MIP or multiplanar reformatting (MPR) techniques; please refer to traditional radiology literature for better understanding of the different roles of these techniques. In our experience the MIP slab provides superior rendering of the vasculature with its inherent capability of having the whole maximally opacified vascular structure highlighted in a single image. The thickness of the MIP image must be finely tuned based on the vessel of interest and vessel location in order to include the whole vessel in the image—the thicker the better—trying to avoid other irrelevant structures such as other vessels or bone. Regardless of the reformatting method, MIP or MPR, a volume rendered image may be generated from the selected VOI and rotated as desired according to the interpretation needed. Notably, the volume rendering reformats obtained with HR-CBCT can be overlayed in real-time to the 2D images in the same way that the 3D-D SA reformats are traditionally used, enabling appropriate views to be selected during the procedure.

APPLIC ATION IN DISEASE

HR-CBCT angiography has proven useful in understanding vascular anatomy due to several factors. First and foremost, the inherent 3D capability makes it possible to visualize the curvilinear vascular structures through 3D space along any plane warranted; the visualization of bone, including its canals and foramina, allows a very detailed analysis of a vessel’s course, something difficult with the traditional 2D-D SA. HR-CBCT also has higher resolution compared to the more commonly used 3D-D SA. With appropriate acquisition protocols and reformatting algorithms, HR-CBCT provides explicit insight into the most detailed anatomy, showing, in effect, essentially the level of resolution available to the neurosurgeon under the operating microscope.

We will demonstrate in the following paragraphs the capability of HR-CBCT acquired during angiography to aid the neurointerventionalist in different pathological conditions. All the images were acquired during routine clinical care as per neurointerventionalist preference.

Aneurysms

Apart from its common use for post-implant evaluation after aneurysm treatment \(^{13}\) (figure 1), which can also be done noninvasively with intravenous contrast injection, \(^{14}\) HR-CBCT has found utility in several aspects of pre-treatment evaluation and planning—for example, in the identification of very small aneurysms, particularly in the posterior circulation. The limited sensitivity of 2D D SA in patients with perimesencephalic hemorrhage is well known and it is our experience that HR-CBCT provides a significant and critically important increase in resolution enabling detection of very small aneurysms (figure 2). We recommend HR-CBCT acquisition when there is a particularly high suspicion for aneurysm and the images acquired with the traditional 2D- or 3D-D SA are not definitively answering the clinical question. Similarly, HR-CBCT has been found useful in identifying traumatic (figure 3) or mycotic (figure 4) aneurysms when the 2D-D SA failed in obviously pointing to the disease. Another role of HR-CBCT in the work-up of brain aneurysms is to evaluate the detailed features of a target aneurysm before treatment,
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such as: small lobulations in the aneurysm fundus, better understanding of complex neck aneurysms (figure 5), or, as importantly, the presence of small perforators arising from the aneurysm neck—specifically of paramount importance in cases of parent vessel sacrifice such as in dissecting vertebral artery aneurysms (figure 7).15 16

Brain arteriovenous malformations

In brain arteriovenous malformations (AVMs), HR-CBCT has been used for localization purposes, either for surgical17 or stereotactic radiation18 planning. In our practice it has also found a role for enhanced understanding of the vascular microanatomy of an AVM. Generally speaking, this is not a common scenario, since most of the information is usually gathered with 2D-, 3D- or 4D-D SA imaging. But the images provided by HR-CBCT do contain more data and as such may be additionally useful in certain situations—for example, when a precise localization is needed either preoperatively or to better correlate with the rest of the cross-sectional imaging during intervention. HR-CBCT has proved particularly useful for anatomical understanding in posterior fossa brainstem or cerebellar AVMs, areas in which the size of the vessels—normal or pathologic—and their superimposition makes the distinction of different branches often challenging (figures 8–9). We
have also found it useful to obtain CBCT with microcatheter injection in certain scenarios, such as when trying to understand a smaller AVM compartment. The obvious limitation of HR-CBCT for AVM is that the length of the protocol results in opacification of arteries and veins with similar intensity which may confound easy differentiation of these vascular structures. Sometimes, however, this can translate into a better understanding of the AVM’s venous outflow and adjacent normal pial veins not involved in primary drainage of the lesion (figure 8).

Cranial dural arteriovenous fistula

The role of HR-CBCT in dural arteriovenous fistula (dAVF) has been demonstrated already in multiple articles and it is currently used by many groups. One obvious reason to prefer CBCT to 3D-DSA is to understand the angioarchitecture of the fistula in more detail, namely the vessels supplying it and adjacent dangerous anastomoses (figures 10–11). A careful evaluation of the images provided may lead to identification of the venous pouch which can be superselected as a venous target. The different openings in the wall of the sinus can be identified, counted and targeted as needed (figure 10). Specifically for cavernous dAVF, HR-CBCT proved useful for mapping the preferred transvenous route in the case of occluded inferior petrosal sinus (IPS) since the long acquisition makes it possible to visualize the distal IPS in the late venous phase or proved useful to visualize the occluded superior petrosal sinus as the preferred route.

Stroke

Traditional utilizations of non-contrast CBCT, either before the thrombectomy as a substitute of the traditional CT scanner or during/after the thrombectomy to rule out a hemorrhage, are very well detailed elsewhere. The roles of HR-CBCT angiography during stroke thrombectomy are infrequent. They include the evaluation of collaterals or the visualization of vessels distally to a point of occlusion, through an acquisition obtained during intra-arterial injection from the ascending aorta. HR-CBCT has also proved very useful to understand the underlying condition in situations of refractory or recurrent occlusion, being often able to allow the identification of intracranial atherosclerosis or dissection by visualizing the flap or the false lumen (figure 12).

Miscellaneous

Generally, HR-CBCT has found a role in many other conditions as well, such as in the case of carotid-cavernous fistula (CCF) where the images obtained helped to understand the size of the hole and the potential venous pathways (figure 13). For vasospasm, this technique proved useful to identify precisely the foci of stenosis, offering insights into the pathophysiology of the condition. For middle meningeal artery embolization procedures in the setting of subdural hematomas, as well as for tumor embolization procedures, it can provide information on the collateral pathways and the arterial supply. For spinal vascular malformations, especially spinal dAVF, HR-CBCT has...
proved very useful in localizing the fistula and treatment planning by identifying in detail the radiculopial and radiculomedullary vessels in relation to the dAVF feeders.29 30

Exploration of normal anatomy
The images obtained to evaluate neurovascular diseases proved to be particularly rich in information to collaterally understand the normal neurovascular anatomy. While this is not the topic of this review, we point to other papers31–34 to further understand the role of HR-CBCT for this purpose.

Figure 11 Elderly woman with cavernous dAVF. Common carotid artery (A) and selective ascending pharyngeal artery (B) 2D-DSA lateral projections showing a cavernous dAVF with supply from multiple ECA and ICA branches and exclusive cortical venous drainage, without ophthalmic or petrosal sinus drainage. The HR-CBCT was performed reconstructed in full VOI (C) and smaller VOI (D) also with volume rendered reformat (E), demonstrating well the primary fistula pouch (arrow in D) as well as showing the existence of a hypoplastic, partially thrombosed inferior petrosal sinus (arrow in C). Based on the HR-CBCT images, the decision was made to attempt a catheterization through the inferior petrosal sinus, which was successful, leading to fistula cure with a selective pouch coiling (G) and sparing the rest of the cavernous sinus, as shown in post-embolization common carotid artery angiography lateral projection subtracted (F) and non-subtracted (G). 2D-DSA, two-dimensional digital subtraction angiography; dAVF, dural arteriovenous fistula; ECA, external carotid artery; HR-CBCT, high resolution cone beam CT; ICA, internal carotid artery; VOI, volume of interest.

Figure 12 Patient with history of trauma and left MCA inferior division ischemic infarct, presenting for angiogram to evaluate the intracranial vasculature. While 2D-DSA (A) clearly showed a stenosis of the inferior division of the left MCA, it is only by performing HR-CBCT with small VOI reconstruction (B–D), corresponding to the inset of figure A, that the anatomy of the vessel could be interpreted as being related to an intracranial dissection, with the false lumen terminating in a blind pouch (dotted arrows) and compressing the true lumen of the inferior division MCA, hence causing the stenosis (white arrows). 2D-DSA, two-dimensional digital subtraction angiography; HR-CBCT, high resolution cone beam CT; MCA, middle cerebral artery; VOI, volume of interest.

Figure 13 Direct traumatic cavernous fistula with lack of antegrade flow in the ipsilateral ICA (A) and collateral supply from the contralateral ICA (B). HR-CBCT performed through right ICA injection (C, D) showed very well the communication between the carotid artery and the cavernous sinus—schematized in the inset with the internal carotid artery (in red) and the cavernous sinus (in blue). Notice in D the filling of the contralateral cavernous sinus from the CCF through the intercavernous sinus with the filling defect of the left cavernous sinus structures such as the ICA and the cranial nerves (arrows), CCF; carotid-cavernous fistula; HR-CBCT, high resolution cone beam CT; ICA, internal carotid artery.

Limitations
Limitations of the HR-CBCT technique include the length of acquisition which carries with it: (1) the requirement for complete paralysis; (2) a vascular phase limitation making it difficult to visualize only the arterial or only the venous phase; (3) the amount of contrast given, since the contrast opacification has to be even throughout the acquisition, a longer acquisition means a larger amount of contrast; (4) the amount of radiation, since longer acquisitions usually result in larger amounts of radiation.

Paralysis. While HR-CBCT can be obtained during any routine angiography, it is our experience that patient movement during the acquisition limits greatly the quality of the images obtained. If a high level of resolution is needed for diagnostic purposes, general anesthesia with breath hold should be used.

Vascular phase. The best images are provided using 20s HR-CBCT, but acquiring a volume over 20s makes it impossible to be able to separate the visualization of the arteries from veins, the so-called temporal resolution, typically thought of as a major advantage of catheter angiography. Most of the time this limitation is secondary since the distinction of the anatomic structures can be made by following the branches to their bigger parent vessels, but when trying to distinguish small perforators such as basilar perforators or white matter perforators, the task remains daunting. We believe that technological evolution in this perspective with faster CBCT acquisition with preserved resolution (such as for Siemens, the 8s DynaCT on Icono vs the 20s option on Artis Q) will result resolutory from this perspective.
A 6 s protocol may be enough but a faster one (4 s) may be even more ideal, given the time of arterial to venous flow in the brain.

Amount of contrast. An acquisition of 20 s necessitates an injection of 40–70 mL of diluted contrast depending on the territory injected. It is certainly reasonable to try and limit the acquisition for situations of clinical interest. While contrast material-induced nephropathy is rare after cerebral angiography (possibly due to separation of total contrast material dose in time intervals, with small amounts administered during separate injections), we recommend against indiscriminate volumetric acquisition of multiple territories. It is possible that a relatively larger injection for CBCT angiography may increase the risk, but in our experience we did not have any such occurrence in patients with normal renal function. We have also not encountered any neurologic effects from high volume contrast injections.

Radiation. The radiation doses with current generation angiographic machines are remarkably lower than those with older generation equipment and a 20 s CBCT results in approximately 200 mGy (reference air kerma). Regardless, rotational angiography is associated with reduced radiation doses compared with 2D-DSA for a few reasons: the acquisition often allows for diminished utilization of 2D-DSA making the final total dose reduced; furthermore, the radiation is dispersed over multiple different projections, so even if the overall mGy is higher, the contribution of a CBCT acquisition to the maximum skin dose during a procedure is low. Certainly it is good practice to have the team temporarily leave the angio suite during the CBCT acquisition to limit the radiation exposure to personnel.

CONCLUSION

HR-CBCT angiography, which is available with any contemporary fluoroscopy machinery, is critically useful in certain clinical scenarios to improve the understanding of neurovascular anatomy and pathology, as well as guiding therapeutic decision-making. It is rapidly becoming an essential tool for the modern neurointerventionalist.

REFERENCES


