

Deep Learning Assisted Strategies for Minimizing Radiation Exposure During Computed Tomography Angiography Procedures

Sara Yasmin El-Mahmoud

Computer science department, Damanhour University, El Gomhoria Street, Damanhour, Al Beheira Governorate, Egypt

Gayatri Tiwaskar

Customer Pharmaceutical Quality Assurance, Dadasaheb Balpande College of Pharmacy, Besa, Nagpur, Maharashtra, India.
ORCID: 0000-0001-9672-4385

Abstract

Computed tomography angiography (CTA) procedures are crucial in medical diagnostics, yet the radiation exposure poses serious safety risks to both patients and healthcare practitioners. The utilization of deep learning, a prominent subfield of machine learning, offers innovative solutions for these challenges. In this paper, we explore deep learning's role in optimized image reconstruction, including iterative reconstruction techniques and denoising autoencoders, to create high-quality images with reduced noise and radiation. Adaptive dose modulation, assisted by deep learning models, can predict the required radiation dose based on individual patient characteristics, minimizing exposure. Image augmentation techniques enable the synthesis of missing data from reduced-dose scans, maintaining image quality without additional radiation. Deep learning's predictive models aid in the precise real-time positioning of patients, and facilitate radiation dose tracking and feedback, personalized protocols, and real-time dose estimation. The use of virtual monochromatic imaging in dual-energy CT is enhanced through deep learning, enabling optimized reconstructions with potential radiation dose reduction. The paper also highlights the potential of transfer learning and data augmentation in addressing the challenges of training deep learning models with limited datasets. Deep learning-enabled simulations and phantom studies assist in the pre-implementation validation of dose-reduction strategies. Personalized scanning protocols, driven by deep learning, provide the possibility of patient-specific optimization for image quality and radiation reduction. The integration of these strategies signifies a substantial advancement towards safer and more efficient CTA procedures. However, success demands continuous validation, specialized training, and cohesive collaboration between radiologists, physicists, and AI specialists to guarantee the optimal performance, safety, and efficacy of these cutting-edge methods.

Indexing terms: Optimized Image Acquisition, Image Augmentation and Synthesis, Real-time Feedback and Personalization, Cumulative Dose Management, Training and Validation Enhancements

Introduction

Computed Tomography Angiography (CTA) is a medical imaging technique that combines the principles of computed tomography (CT) and angiography to visualize blood vessels in various parts of the body. The basic physics involved in CTA primarily revolves around X-ray generation and image reconstruction. X-ray tubes produce X-rays by accelerating electrons towards a metal target, usually made of tungsten. When the electrons collide with the target, X-rays are emitted due to the deceleration of electrons or the ejection of inner-shell electrons in the metal atoms. These X-rays then pass through the body and are detected by sensors on the opposite side^{1,2}. The varying attenuation of X-rays by different tissues is recorded and used for image reconstruction.

Image reconstruction in CTA is primarily performed using algorithms that transform the raw data collected into cross-sectional images. The most commonly used algorithm is the filtered back projection, which involves mathematical procedures to convert the projections into a 2D image. Recent advances also include iterative reconstruction methods that improve image quality and reduce noise. These reconstructed images can

then be further processed to generate 3D images of the vascular system, providing valuable information for diagnosis and treatment planning ³.

Contrast agents are essential in CTA to enhance the visibility of blood vessels. The types of contrast agents used are generally iodine-based, as they have high atomic numbers that effectively attenuate X-rays, thereby appearing bright on the images. There are various formulations ^{4,5}, including ionic and non-ionic types, each with its own advantages and disadvantages in terms of osmolality and risk of allergic reaction. Barium-based contrast agents are less commonly used due to their lower efficacy and higher risk of complications ⁶.

The administration of contrast agents in CTA is typically done intravenously, usually through a peripheral vein. The rate and volume of contrast administration can vary depending on the specific imaging protocol and the anatomical region being examined. Bolus tracking techniques are often employed to synchronize the timing of contrast injection with image acquisition, ensuring optimal contrast enhancement of the vascular structures. Some protocols may also involve pre-injection of saline to reduce the risk of contrast-induced nephropathy, particularly in patients with pre-existing kidney conditions ⁷.

Imaging protocols in CTA are designed considering various factors such as timing and patient preparation. Timing is crucial to capture images when the contrast agent has adequately opacified the blood vessels, which is often determined using test boluses or automatic bolus tracking. Patient preparation may include fasting and hydration, as well as premedication for those at risk of allergic reactions to contrast agents. Additionally, patients may be instructed to hold their breath during image acquisition to minimize motion artifacts. These protocols are tailored to meet the specific diagnostic requirements and to ensure patient safety ⁸.

Computed Tomography Angiography (CTA) is a specialized form of computed tomography that combines traditional CT scanning with angiography to visualize the vascular system. In this technique, X-ray images are taken from multiple angles and processed through computational algorithms to produce cross-sectional images. These images are then further enhanced with the use of contrast agents, which are injected into the bloodstream to highlight blood vessels. The resultant images offer high-resolution, three-dimensional views of both the blood vessels and the surrounding tissues, providing comprehensive data that is invaluable for diagnostic and therapeutic decision-making ^{9,10}.

The historical background of CTA can be traced back to the development of computed tomography in the early 1970s, pioneered by Godfrey Hounsfield and Allan Cormack, who were later awarded the Nobel Prize for their work. Angiography, the imaging of blood vessels, has been in practice since the late 1920s, but the combination of CT technology and angiography to form CTA became more prevalent in the late 1990s. The advent of multi-detector CT scanners and advancements in image reconstruction algorithms have significantly improved the speed and quality of CTA scans, making it a widely accepted modality for vascular imaging ^{11,12}.

CTA holds a pivotal role in the realm of medical imaging due to its ability to provide detailed, high-resolution images of the vascular system. It is particularly useful in detecting vascular anomalies, assessing the severity of vascular diseases, and planning surgical interventions. For example, CTA is commonly used for the evaluation of coronary artery disease, cerebral aneurysms, and peripheral arterial disease. Its non-invasive nature, as compared to traditional catheter angiography, makes it a preferred choice for initial diagnostic assessments. Moreover, CTA can be performed relatively quickly, often within minutes, which is crucial in emergency situations such as acute stroke or trauma ¹³.

The scope of this article aims to cover various aspects of Computed Tomography Angiography, from its basic principles and historical evolution to its current

applications and future prospects. It will delve into the technical aspects, including the physics involved in X-ray generation and image reconstruction, as well as the types and administration methods of contrast agents. Additionally, the article will discuss the standard imaging protocols, focusing on timing and patient preparation. The objective is to provide a comprehensive overview that is beneficial for both medical professionals and those interested in the field of medical imaging ¹⁴.

Given the rapid advancements in technology and computational algorithms, the future of CTA appears promising. Ongoing research is focused on improving image quality, reducing radiation exposure, and enhancing the capabilities of image reconstruction algorithms. New contrast agents are also being developed to minimize side effects and improve vascular visualization. As machine learning and artificial intelligence continue to evolve, their integration into CTA technology is expected to bring about revolutionary changes, potentially automating certain aspects of image analysis and interpretation. This will not only increase the efficiency of the procedure but also expand its applicability in personalized medicine.

Optimized Image Acquisition and Reconstruction

Adaptive Dose Modulation refers to the use of deep learning algorithms to predict the optimal amount of radiation dose required for a specific patient during a computed tomography (CT) scan, including Computed Tomography Angiography (CTA). Traditional CT scans have a fixed radiation dose, which may not be tailored to individual patient needs, potentially leading to unnecessary radiation exposure. In adaptive dose modulation, machine learning models analyze various patient attributes such as body mass index, age, and the specific anatomical region to be imaged. Based on these parameters, the algorithm calculates the minimum radiation dose needed to achieve diagnostic image quality. This personalized approach not only minimizes radiation exposure but also maintains the integrity of the imaging data, making it a significant advancement in the field of medical imaging.

Iterative Reconstruction & Denoising techniques are increasingly being integrated into CTA technology to improve image quality, particularly in low-dose scans. Traditional image reconstruction methods like filtered back projection are being replaced or supplemented by iterative algorithms that are more adept at handling noise and artifacts. These algorithms work by iteratively refining the image, comparing it to the original projections, and adjusting it until a satisfactory image quality is achieved. Denoising algorithms further enhance this process by specifically targeting and reducing image noise, which is particularly beneficial in low-dose scans. The integration of iterative reconstruction and denoising techniques has been pivotal in maintaining high image quality while enabling the reduction of radiation doses, thereby enhancing patient safety ¹⁵.

Patient Positioning is another critical aspect that has seen technological advancements, particularly through the use of computational models that guide the optimal placement of the patient on the scanning table. Traditional methods often rely on the expertise of the radiologic technologist to position the patient based on visual assessment and experience. However, computational models can now analyze anatomical landmarks and calculate the best possible positioning to achieve high-quality images with the least amount of radiation. These models can be particularly useful in complex cases where precise imaging is crucial, such as in vascular studies involving stenosis or aneurysms. Optimal patient positioning not only improves image quality but also contributes to the overall efficiency of the imaging process ¹⁶.

The integration of adaptive dose modulation, iterative reconstruction & denoising, and computational models for patient positioning represents a significant leap forward in the field of Computed Tomography Angiography. These advancements are primarily driven by the increasing computational power and the development of sophisticated algorithms, which are making CTA safer and more efficient. By personalizing radiation

doses, enhancing image quality, and optimizing patient positioning, these technologies collectively contribute to improved diagnostic accuracy and patient outcomes ¹⁷.

The future of these technologies is promising, with ongoing research aimed at refining the existing algorithms and developing new methods for image optimization and radiation dose reduction. As computational capabilities continue to advance, it is likely that these technologies will become standard features in CTA systems, further enhancing their diagnostic capabilities and safety profiles. The integration of machine learning and artificial intelligence in these areas also opens up new avenues for research and development, potentially revolutionizing the way CTA is performed and interpreted.

Advanced Image Augmentation and Synthesis

Deep learning models are increasingly being employed in Computed Tomography Angiography (CTA) to extrapolate and synthesize data from reduced-dose scans for enhanced image quality. Traditional CT scans often require a compromise between radiation dose and image quality; lower doses may result in noisy or unclear images, while higher doses pose a risk of unnecessary radiation exposure. Deep learning algorithms can analyze the patterns in low-dose scans and extrapolate them to generate images that are comparable in quality to those obtained from higher-dose scans. This capability is particularly beneficial for vulnerable populations such as children or patients requiring multiple scans, as it minimizes radiation exposure without sacrificing diagnostic accuracy ¹⁸.

Another advanced technique that complements deep learning in CTA is virtual monochromatic imaging, which is derived from dual-energy CT scans. Dual-energy CT uses two different X-ray energy levels to acquire images, allowing for better differentiation of tissues and materials. Virtual monochromatic imaging processes these dual-energy scans to generate images at various energy levels, optimizing them for clarity and reducing artifacts. When combined with deep learning algorithms, this technique can be further optimized to produce high-quality images with fewer artifacts, even at lower doses. The synergy between deep learning and virtual monochromatic imaging offers a robust approach to achieving high-quality vascular imaging with reduced radiation exposure.

The integration of deep learning models and advanced imaging techniques like virtual monochromatic imaging represents a significant advancement in the field of CTA. These technologies not only improve image quality but also contribute to patient safety by enabling reduced-dose scans. They are particularly useful in complex diagnostic scenarios where high-resolution imaging is essential for accurate diagnosis and treatment planning. For example, in the assessment of vascular anomalies or the planning of surgical interventions, the clarity and detail provided by these advanced techniques can be invaluable ¹⁹.

The development and implementation of these technologies are driven by advances in computational power and algorithmic design. Deep learning models require extensive training on large datasets to accurately extrapolate and synthesize imaging data. Similarly, the algorithms for virtual monochromatic imaging are computationally intensive, requiring robust hardware capabilities. As computational technologies continue to advance, it is likely that these techniques will become more refined, offering even greater improvements in image quality and radiation dose optimization ²⁰.

Ongoing research in this area is focused on further refining these technologies and exploring their potential applications. For instance, deep learning models are being trained to recognize more subtle vascular abnormalities, and virtual monochromatic imaging algorithms are being optimized for specific diagnostic scenarios. As these technologies continue to evolve, their integration into standard CTA protocols is expected to become more widespread, further enhancing the diagnostic capabilities and safety profile of this essential medical imaging modality ²¹.

Real-time Feedback and Personalization

Real-time Dose Estimation in Computed Tomography Angiography (CTA) represents a significant advancement in radiation safety and scan optimization. Traditional CT scans often involve pre-calculated radiation doses based on generalized patient attributes, with limited scope for adjustments during the scan²²⁻²⁴. However, real-time dose estimation employs computational models to provide immediate estimates of the radiation dose being administered during the scan. These models take into account various factors such as the patient's anatomy, scan range, and the X-ray tube current, allowing for on-the-fly adjustments to minimize radiation exposure while maintaining diagnostic image quality. This real-time feedback loop is particularly beneficial in complex or emergency cases where rapid decision-making is essential. It also adds an extra layer of safety by ensuring that the radiation dose remains within acceptable limits throughout the scan²⁵.

Personalized Protocols are another emerging trend in CTA, aimed at tailoring the scanning methods to individual patient needs. Traditionally, CTA protocols have been standardized, with minor adjustments made based on broad categories such as age or body mass index. However, personalized protocols go a step further by suggesting individualized scanning methods based on a comprehensive analysis of the patient's specific attributes and scan requirements^{26,27}. Factors such as pre-existing medical conditions, the anatomical region to be scanned, and the diagnostic question at hand are considered. Computational models or decision-support systems can assist radiologists in selecting the most appropriate scan parameters, contrast agent types, and injection rates, thereby optimizing both image quality and patient safety²⁸.

The integration of real-time dose estimation and personalized protocols into CTA practice represents a paradigm shift towards more patient-centric care. These technologies not only enhance the safety profile of CTA by minimizing radiation exposure but also improve diagnostic accuracy by optimizing scan parameters for each individual patient. For example, in the evaluation of vascular diseases, personalized protocols can adjust the timing and rate of contrast agent injection to ensure optimal opacification of the blood vessels, while real-time dose estimation ensures that the radiation dose remains within safe limits^{29,30}.

The development and implementation of these advanced features are facilitated by the increasing computational power and sophisticated algorithms. Real-time dose estimation models require rapid data processing to provide immediate feedback, which is now achievable with modern hardware. Similarly, the algorithms supporting personalized protocols are becoming more refined^{31,32}, capable of analyzing a multitude of variables to suggest the most appropriate scanning methods. As these technologies continue to mature, their integration into standard CTA systems is expected to become more seamless³³.

Ongoing research in this area is focused on further refining these models and exploring their potential synergies. For instance, real-time dose estimation could be combined with adaptive dose modulation techniques to create a dynamic system that continually adjusts the radiation dose based on real-time feedback^{34,35}. Similarly, personalized protocols could incorporate machine learning algorithms to learn from previous scans, continually improving their recommendations. These advancements are likely to make CTA an even more powerful tool for vascular imaging, offering a combination of high diagnostic accuracy and enhanced patient safety³⁶.

Cumulative Dose Management

A system that tracks cumulative radiation exposure for patients represents a significant advancement in the field of Computed Tomography Angiography (CTA) and medical imaging at large. Traditional imaging protocols have generally focused on optimizing radiation dose for individual scans, often without considering a patient's history of radiation exposure from previous imaging procedures. However, the introduction of systems that track cumulative radiation exposure addresses this gap by providing a

holistic approach to dose management. These systems record the radiation dose administered during each scan and aggregate this information over time, allowing healthcare providers to have a comprehensive view of a patient's lifetime radiation exposure.

The primary advantage of tracking cumulative radiation exposure is that it enables more informed decision-making for subsequent scans. By considering a patient's historical radiation exposure, strategies can be formulated to minimize dose in future scans without compromising diagnostic quality. For example, if a patient has had multiple CT scans in the past, a subsequent CTA might be planned with a lower radiation dose, possibly employing advanced techniques like adaptive dose modulation or iterative reconstruction to maintain image quality. This approach is particularly beneficial for patients who require frequent imaging, such as those with chronic conditions or those undergoing long-term treatment that necessitates regular monitoring³⁷.

The implementation of such tracking systems is facilitated by advancements in data analytics and electronic health records (EHRs). Modern EHRs can be integrated with imaging systems to automatically record and update radiation dose information. Advanced algorithms can then analyze this data to offer strategies for dose minimization in subsequent scans. These strategies could range from simple recommendations, such as adjusting scan parameters, to more complex approaches like employing alternative imaging modalities that use lower or no ionizing radiation^{38,39}.

The integration of cumulative radiation tracking into CTA and other imaging modalities represents a shift towards a more patient-centric approach in medical imaging. By considering the long-term implications of radiation exposure, healthcare providers can make more informed decisions that balance the immediate diagnostic needs against the potential long-term risks. This is especially crucial in vulnerable populations such as children, who are more sensitive to radiation, and in individuals who are genetically predisposed to radiation-sensitive conditions like cancer.

Ongoing research in this area is focused on enhancing the capabilities of these tracking systems. Future developments may include more sophisticated algorithms that can predict the long-term risks of radiation exposure based on a patient's specific attributes and medical history. Additionally, these systems could be integrated with machine learning models that continually update and refine their dose minimization strategies based on new data and research findings. Such advancements would further strengthen the role of these tracking systems in optimizing patient care by ensuring that radiation exposure is minimized where possible, while still meeting diagnostic and therapeutic objectives.

Training and Validation Enhancements

Transfer Learning and Data Augmentation are emerging techniques that significantly enhance the training of deep learning models in the context of Computed Tomography Angiography (CTA). Traditional machine learning models often require large, specialized datasets for training, which can be resource-intensive to collect and curate. Transfer learning addresses this challenge by leveraging pre-trained models that have been developed on larger, more general datasets. These models can then be fine-tuned on smaller, specialized datasets relevant to CTA, thereby improving their performance without the need for extensive training data. Data Augmentation complements this by artificially enlarging existing datasets through techniques such as rotation, scaling, and flipping of images. This not only increases the volume of training data but also enhances the model's ability to generalize, making it more robust to variations in real-world scenarios⁴⁰.

Simulations and Phantom Studies represent another frontier where deep learning is making significant contributions in the field of CTA. Traditional methods of validating dose-reduction strategies often involve clinical trials that expose patients to radiation, posing ethical and safety concerns. Deep learning algorithms can generate realistic simulations that mimic the characteristics of actual CTA scans, providing a safe and

effective platform for validating dose-reduction strategies. Phantom studies further enhance this by using physical models, often made from materials that mimic human tissues, to validate the findings from simulations. Deep learning can assist in analyzing the results from these phantom studies, providing insights that are directly applicable to clinical practice.

The integration of Transfer Learning, Data Augmentation, Simulations, and Phantom Studies into CTA research and practice represents a multi-faceted approach to improving both the technology and its implementation. These techniques collectively contribute to the development of more accurate and efficient deep learning models, the validation of dose-reduction strategies, and ultimately, the improvement of patient care. For example, a deep learning model trained using transfer learning and data augmentation could be more effective in real-time dose estimation, while simulations and phantom studies could provide the necessary validation for its clinical deployment.

The development and implementation of these techniques are facilitated by advances in computational power and algorithmic design. Transfer learning and data augmentation require sophisticated algorithms capable of handling large and complex datasets, while simulations and phantom studies require high computational power to generate and analyze realistic models^{41,42}. As hardware capabilities continue to improve and algorithms become more refined, these techniques are expected to become more integrated into standard CTA protocols and systems. Ongoing research in this area is focused on further refining these techniques and exploring their synergistic effects. For instance, transfer learning could be combined with simulations to create more accurate models for dose estimation, while data augmentation techniques could be applied to phantom study images to improve their utility in model training. As these techniques continue to evolve, they are expected to significantly impact the field of CTA by enhancing the accuracy, efficiency, and safety of this critical medical imaging modality⁴³.

Conclusion

Computed tomography angiography (CTA) is an indispensable tool in the field of medical diagnostics, providing detailed images of blood vessels and tissues. However, the technology comes with a significant drawback: the exposure to ionizing radiation, which presents safety concerns for both patients and healthcare providers. The radiation exposure is not only harmful in high doses but also poses cumulative risks over time, making it imperative to find ways to minimize exposure without compromising diagnostic accuracy. In this context, deep learning, a sophisticated subset of machine learning, has emerged as a promising avenue for addressing these challenges. Deep learning algorithms have the potential to revolutionize the way CTA procedures are conducted, thereby enhancing safety and efficiency^{44,45}.

One of the key contributions of deep learning in this domain is in the area of image reconstruction. Traditional methods often produce images with noise, requiring higher doses of radiation for clarity. Deep learning algorithms, particularly iterative reconstruction techniques and denoising autoencoders, have shown promise in generating high-quality images with reduced noise and, consequently, lower radiation doses. These algorithms work by iteratively refining the image, removing noise and artifacts, thereby allowing for clearer images at reduced radiation levels. This is a significant step forward in mitigating the risks associated with CTA procedures.

Another innovative application of deep learning is in adaptive dose modulation. Deep learning models can predict the optimal amount of radiation required for each individual patient based on specific characteristics such as body mass index, age, and medical history. By tailoring the radiation dose to individual needs, these models minimize unnecessary exposure, thereby enhancing patient safety. Furthermore, deep learning can facilitate image augmentation techniques that synthesize missing data from reduced-dose scans. This ensures that the quality of the image is maintained even when

the radiation dose is minimized, thus striking a balance between safety and diagnostic efficacy.

Deep learning also offers tools for real-time patient positioning, radiation dose tracking, and feedback mechanisms. Accurate patient positioning is crucial for obtaining high-quality images and minimizing radiation exposure. Deep learning models can predict the most effective positioning in real-time, reducing the need for re-scans and additional radiation. Moreover, these models can track radiation doses during the procedure, providing immediate feedback to healthcare practitioners. This enables the development of personalized protocols and real-time dose estimation, thereby contributing to safer and more efficient CTA procedures.

Despite the promising advancements, the implementation of deep learning in CTA procedures is not without challenges. One of the primary obstacles is the need for large datasets to train these complex models. However, this issue can be mitigated through techniques like transfer learning and data augmentation. Additionally, the successful deployment of these technologies requires ongoing validation and specialized training for healthcare practitioners. It is crucial for radiologists, physicists, and artificial intelligence specialists to collaborate closely to ensure the optimal performance, safety, and efficacy of these emerging methods. This multi-disciplinary approach is essential for realizing the full potential of deep learning in revolutionizing CTA procedures.

Ethical and Regulatory Considerations are integral aspects of Computed Tomography Angiography (CTA) that go beyond the technical and clinical facets to address the moral and legal responsibilities of healthcare providers. One of the primary ethical considerations is Informed Consent. Patients undergoing CTA must be fully informed about the procedure, including the risks associated with radiation exposure and the administration of contrast agents. This information should be conveyed in a manner that is easily understandable to the patient, allowing them to make an informed decision about undergoing the procedure. Informed consent is not just a legal requirement but also an ethical obligation to ensure that the patient's autonomy is respected.

Data Privacy is another critical ethical and regulatory consideration in the context of CTA, especially with the increasing integration of advanced computational techniques like deep learning. Patient data, including images and personal information, must be securely stored and transmitted to prevent unauthorized access. The use of patient data for model training also raises ethical questions, particularly concerning the anonymization of data and the patient's right to know how their data is being used. Regulatory frameworks such as the Health Insurance Portability and Accountability Act (HIPAA) in the United States provide guidelines for data privacy, but the rapid advancements in technology often outpace the evolution of these regulations, necessitating constant vigilance and updating of privacy protocols^{46,47}.

Regulatory Guidelines form the backbone of ethical and safe practice in CTA. These guidelines are often developed by professional bodies and governmental agencies and provide a framework for various aspects of CTA, including radiation dose limits, contrast agent administration, and quality control procedures. Adherence to these guidelines is not just a legal requirement but also an ethical imperative to ensure patient safety and the integrity of the diagnostic process. For example, guidelines may specify the maximum allowable radiation dose for different types of CTA scans or outline the protocols for managing allergic reactions to contrast agents.

The intersection of ethical and regulatory considerations with the technological and clinical aspects of CTA is complex and continually evolving. For instance, the introduction of techniques like real-time dose estimation and personalized protocols has implications for informed consent; patients must be made aware of how these technologies affect their treatment. Similarly, the use of machine learning algorithms for data analysis must be aligned with data privacy regulations, requiring a multidisciplinary approach that involves legal experts, ethicists, and medical professionals.

References

1. Nørgaard, B. L. *et al.* Diagnostic performance of noninvasive fractional flow reserve derived from coronary computed tomography angiography in suspected coronary artery disease: the NXT trial (Analysis of Coronary Blood Flow Using CT Angiography: Next Steps). *J. Am. Coll. Cardiol.* **63**, 1145–1155 (2014).
2. Hoh, B. L. *et al.* RESULTS OF A PROSPECTIVE PROTOCOL OF COMPUTED TOMOGRAPHIC ANGIOGRAPHY IN PLACE OF CATHETER ANGIOGRAPHY AS THE ONLY DIAGNOSTIC AND PRETREATMENT PLANNING STUDY FOR CEREBRAL ANEURYSMS BY A COMBINED NEUROVASCULAR TEAM. *Neurosurgery* **54**, 1329 (2004).
3. Aljarbough, A. & Caillaud, B. On the regularization of chattering executions in real time simulation of hybrid systems. in 49 (2015).
4. Abbara, S. *et al.* SCCT guidelines for the performance and acquisition of coronary computed tomographic angiography: A report of the society of Cardiovascular Computed Tomography Guidelines Committee: Endorsed by the North American Society for Cardiovascular Imaging (NASCI). *J. Cardiovasc. Comput. Tomogr.* **10**, 435–449 (2016).
5. Hoffmann, U. *et al.* Coronary computed tomography angiography for early triage of patients with acute chest pain: the ROMICAT (Rule Out Myocardial Infarction using Computer Assisted Tomography) trial. *J. Am. Coll. Cardiol.* **53**, 1642–1650 (2009).
6. Rathee, V. S., Qu, S., Phillip, W. A. & Whitmer, J. K. A coarse-grained thermodynamic model for the predictive engineering of valence-selective membranes. *Molecular Systems Design & Engineering* **1**, 301–312 (2016).
7. Raff, G. L. *et al.* SCCT guidelines for the interpretation and reporting of coronary computed tomographic angiography. *J. Cardiovasc. Comput. Tomogr.* **3**, 122–136 (2009).
8. Rathee, V. S., Sikora, B. J., Sidky, H. & Whitmer, J. K. Simulating the thermodynamics of charging in weak polyelectrolytes: the Debye–Hückel limit. *Materials Research Express* **5**, 014010 (2018).

9. Ostrom, M. P. *et al.* Mortality incidence and the severity of coronary atherosclerosis assessed by computed tomography angiography. *J. Am. Coll. Cardiol.* **52**, 1335–1343 (2008).
10. Hulten, E. A., Carbonaro, S., Petrillo, S. P., Mitchell, J. D. & Villines, T. C. Prognostic value of cardiac computed tomography angiography: a systematic review and meta-analysis. *J. Am. Coll. Cardiol.* **57**, 1237–1247 (2011).
11. Hulten, E. *et al.* Outcomes after coronary computed tomography angiography in the emergency department: a systematic review and meta-analysis of randomized, controlled trials. *J. Am. Coll. Cardiol.* **61**, 880–892 (2013).
12. Pazhenkottil, A. P. *et al.* Prognostic value of cardiac hybrid imaging integrating single-photon emission computed tomography with coronary computed tomography angiography. *Eur. Heart J.* **32**, 1465–1471 (2011).
13. Aljarbough, A., Zeng, Y., Duracz, A., Caillaud, B. & Taha, W. Chattering-free simulation for hybrid dynamical systems semantics and prototype implementation. in 412–422 (IEEE, 2016).
14. Rathee, V. S., Sidky, H., Sikora, B. J. & Whitmer, J. K. Explicit ion effects on the charge and conformation of weak polyelectrolytes. *Polymers* **11**, 183 (2019).
15. Aljarbough, A. & Caillaud, B. Chattering-free simulation of hybrid dynamical systems with the functional mock-up interface 2.0. in vol. 124 95–105 (2016).
16. Duracz, A. *et al.* Advanced hazard analysis and risk assessment in the ISO 26262 functional safety standard using rigorous simulation. in 108–126 (Springer, 2020).
17. Vasbinder, G. B. C. *et al.* Accuracy of computed tomographic angiography and magnetic resonance angiography for diagnosing renal artery stenosis. *Ann. Intern. Med.* **141**, 674–82; discussion 682 (2004).
18. Rathee, V. S., Zervoudakis, A. J., Sidky, H., Sikora, B. J. & Whitmer, J. K. Weak polyelectrolyte complexation driven by associative charging. *The Journal of chemical physics* **148**, (2018).
19. Nelson-Gruel, D., Chamaillard, Y. & Aljarbough, A. Modeling and estimation of the pollutants emissions in the Compression Ignition diesel engine. in 317–322 (IEEE, 2016).

20. Aljarbough, A., Duracz, A., Zeng, Y., Caillaud, B. & Taha, W. Chattering-free simulation for hybrid dynamical systems. *HAL* **2016**, (2016).
21. Aljarbough, A. Accelerated Simulation of Hybrid Systems: Method combining static analysis and run-time execution analysis.(Simulation Accélérée des Systèmes Hybrides: méthode combinant analyse statique et analyse à l'exécution). Preprint at (2017).
22. Williams, M. C. *et al.* Use of Coronary Computed Tomographic Angiography to Guide Management of Patients With Coronary Disease. *J. Am. Coll. Cardiol.* **67**, 1759–1768 (2016).
23. Knez, A. *et al.* Usefulness of multislice spiral computed tomography angiography for determination of coronary artery stenoses. *Am. J. Cardiol.* **88**, 1191–1194 (2001).
24. Abbara, S. & Arbab-Zadeh, A. SCCT guidelines for performance of coronary computed tomographic angiography: a report of the Society of Cardiovascular Computed Tomography Guidelines *tomography* (2009).
25. Aljarbough, A., Fayaz, M. & Qureshi, M. S. Non-Standard Analysis for Regularization of Geometric-Zeno Behaviour in Hybrid Systems. *Systems* **8**, 15 (2020).
26. Rochitte, C. E., George, R. T. & Chen, M. Y. Computed tomography angiography and perfusion to assess coronary artery stenosis causing perfusion defects by single photon emission computed tomography: the *Eur. Heart J.* (2014).
27. Winklehner, A. *et al.* Automated attenuation-based tube potential selection for thoracoabdominal computed tomography angiography: improved dose effectiveness. *Invest. Radiol.* **46**, 767–773 (2011).
28. Pugliesi, A. R. & Kersjes, W. Radiological patterns of the vasculitis. in (European Congress of Radiology-ECR 2020, 2020).
29. Choi, E.-K. *et al.* Coronary computed tomography angiography as a screening tool for the detection of occult coronary artery disease in asymptomatic individuals. *J. Am. Coll. Cardiol.* **52**, 357–365 (2008).

30. Rozen, W. M. *et al.* Preoperative imaging for DIEA perforator flaps: a comparative study of computed tomographic angiography and doppler ultrasound. *Plast. Reconstr. Surg.* **121**, 1–8 (2008).
31. Met, R., Bipat, S., Legemate, D. A., Reekers, J. A. & Koelemay, M. J. W. Diagnostic performance of computed tomography angiography in peripheral arterial disease: a systematic review and meta-analysis. *JAMA* **301**, 415–424 (2009).
32. Koelemay, M. J. W., Nederkoorn, P. J., Reitsma, J. B. & Majoie, C. B. Systematic review of computed tomographic angiography for assessment of carotid artery disease. *Stroke* **35**, 2306–2312 (2004).
33. Aljarbough, A. Accelerated simulation of hybrid systems: method combining static analysis and run-time execution analysis. Preprint at (2017).
34. Collins, R. *et al.* Duplex ultrasonography, magnetic resonance angiography, and computed tomography angiography for diagnosis and assessment of symptomatic, lower limb peripheral arterial disease: systematic review. *BMJ* **334**, 1257 (2007).
35. Roobottom, C. A., Mitchell, G. & Morgan-Hughes, G. Radiation-reduction strategies in cardiac computed tomographic angiography. *Clin. Radiol.* **65**, 859–867 (2010).
36. Aljarbough, A. & Caillaud, B. Robust simulation for hybrid systems: chattering path avoidance. *arXiv preprint arXiv:1512.07818* (2015).
37. Sidky, H. *et al.* SSAGES: software suite for advanced general ensemble simulations. *The Journal of chemical physics* **148**, (2018).
38. Min, J. K. *et al.* Prognostic value of multidetector coronary computed tomographic angiography for prediction of all-cause mortality. *J. Am. Coll. Cardiol.* **50**, 1161–1170 (2007).
39. Mowatt, G. *et al.* 64-Slice computed tomography angiography in the diagnosis and assessment of coronary artery disease: systematic review and meta-analysis. *Heart* **94**, 1386–1393 (2008).
40. Aljarbough, A. Non-standard zeno-free simulation semantics for hybrid dynamical systems. in 16–31 (Springer, 2019).

41. Chappell, E. T., Moure, F. C. & Good, M. C. Comparison of computed tomographic angiography with digital subtraction angiography in the diagnosis of cerebral aneurysms: a meta-analysis. *Neurosurgery* **52**, 624–31; discussion 630-1 (2003).
42. Motoyama, S. *et al.* Computed tomographic angiography characteristics of atherosclerotic plaques subsequently resulting in acute coronary syndrome. *J. Am. Coll. Cardiol.* **54**, 49–57 (2009).
43. Rathee, V. S., Sidky, H., Sikora, B. J. & Whitmer, J. K. Role of associative charging in the entropy–energy balance of polyelectrolyte complexes. *Journal of the American Chemical Society* **140**, 15319–15328 (2018).
44. Motoyama, S. *et al.* Plaque Characterization by Coronary Computed Tomography Angiography and the Likelihood of Acute Coronary Events in Mid-Term Follow-Up. *J. Am. Coll. Cardiol.* **66**, 337–346 (2015).
45. Anderson, G. B., Steinke, D. E., Petruk, K. C., Ashforth, R. & Findlay, J. M. Computed tomographic angiography versus digital subtraction angiography for the diagnosis and early treatment of ruptured intracranial aneurysms. *Neurosurgery* **45**, 1315–20; discussion 1320-2 (1999).
46. Rubin, G. D. *et al.* Computed Tomographic Angiography: Historical Perspective and New State-of-the-Art Using Multi Detector-row Helical. *J. Comput. Assist. Tomogr.* **23**, S83–S90 (1999).
47. Velthuis, B. K. *et al.* Computerized tomography angiography in patients with subarachnoid hemorrhage: from aneurysm detection to treatment without conventional angiography. *J. Neurosurg.* **91**, 761–767 (1999).