Clinical Validation of a Deep Learning Model for Segmenting and Quantifying Intracranial and Ventricular Volumes on Computed Tomography

Bruna Garbes Gonçalves Pinto^{1*†}, Tayran Milá Mendes Olegário^{1†}, Pedro Vinicius Alves Silva¹, Gabriel Monteiro Ferracioli¹, Artur José Marques Paulo¹, Klaus Schumacher¹, Mateus Trinconi Cunha^{1, 2}, Henrique Min Ho Lee¹, Mariana Athaniel Silva Rodrigues¹, Felipe Campos Kitamura³, Joselisa Peres Queiroz de Paiva¹, Rafael Maffei Loureiro¹

*Corresponding author(s). E-mail(s): bruna.goncalves@einstein.br; †These authors contributed equally to this work.

1 Supplementary Material

The stages of generating the pre-segmentation masks of the intracranial volume include:

- 1. **Image size verification:** If the image has less than 500 slices, it is resized to a new image with 500 slices.
- 2. Application of Gaussian filter: A Gaussian filter with a parameter $\sigma=1$ is applied to the image to reduce noise, uniformizing some image regions and thus facilitating the segmentation process.
- 3. **Thresholding application:** A mask is generated by applying a threshold value to the image. The mask includes all voxels with values between -15 and 50 Hounsfield units.

^{1*}Image Department, Hospital Israelita Albert Einstein, São Paulo, Brazil.
²Sunnybrook Health Sciences Centre, University of Toronto, Toronto, Canada.

^{3*}Departament of Diagnostic Imaging, Universidade Federal de São Paulo, São Paulo, Brazil.

- 4. **Erosion operation application:** An erosion operation is applied to the resulting mask from the previous step to "erode" the sides of the mask and thus disconnect the various structures present in the mask.
- 5. **Selection of the largest connected region:** then the largest connected region of the mask is selected. This largest connected region is expected to correspond to part of the intracranial volume region.
- 6. **Dilation operation application:** A dilation operation is then applied to the largest connected region to "compensate" for previously applied erosion operation. It is expected that after applying this operation, only the intracranial volume region will be obtained. This dilation will only affect voxels with values between -50HU and 150HU.
- 7. Cutting the bottom region of the mask: Finally, a heuristic is applied that traverses the image from bottom to top and calculates the number of connected regions in the axial slices. If only one connected region appears in the slice, it is excluded; the heuristic is terminated if more than one connected region appears.

 ${\bf Table~1} \ \ {\bf Kruskal\text{-}Wallis~and~Effect~Sizes~for~Volumes~by~Patient~Sex}$

	P-Value	η^2	Magnitude
Intracranial Volume - Ground truth	< 0.001	0.38	Large
Intracranial Volume - DeepCTE3D	< 0.001	0.36	Large
Ventricular Volume - Ground truth	< 0.001	0.02	Small
Ventricular Volume - Deep CTE3D $$	0.02	0.01	Small

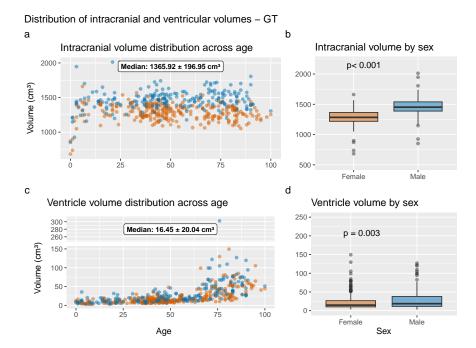
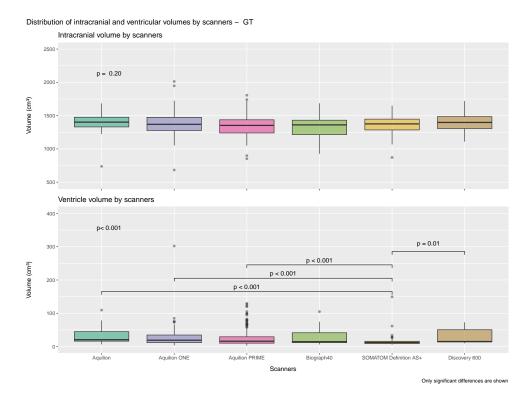


Fig. 1 Volume distributions of ground-truth segmentations. a) Intracranial volume by patient age. b) Intracranial volume by patient sex. c) Ventricular volume by patient age. d) Ventricular volumes by patient sex.

Table 2 Kruskal-Wallis and Effect Sizes for Volumes by Equipment

	P-Value	η^2	Magnitude
Intracranial Volume - GT	0.20	< 0.01	Small
Intracranial Volume - DeepCTE3D	0.25	< 0.01	Small
Ventricular Volume - GT	< 0.001	0.03	Small
Ventricular Volume - DeepCTE3D	< 0.001	0.05	Small



 ${\bf Fig.~2} \ \ {\bf Intracranial~and~ventricular~volumes~by~scanner~model~-~Ground~Truth.}$

Distribution of intracranial and ventricular volumes - DeepCTE3D

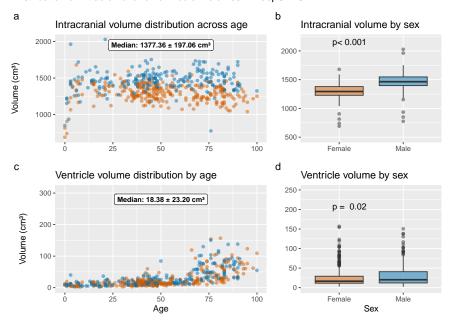
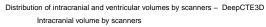


Fig. 3 Volume distributions of DeepCTE3D segmentations. a) Intracranial volume by patient age. b) Intracranial volume by patient sex. c) Ventricular volume by patient age. d) Ventricular volume by patient sex.



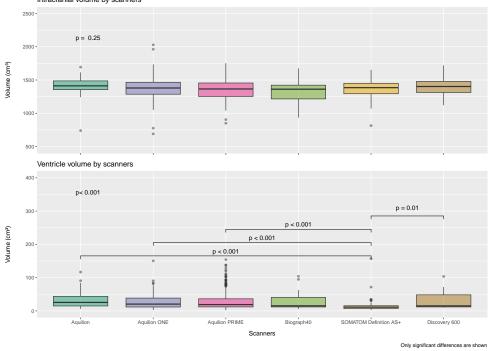


Fig. 4 Volumes by scanner model - DeepCTE3D.

 ${\bf Table~3}~$ Kruskal-Wallis and Effect Sizes for Relative Differences by Patient Sex and Scanners

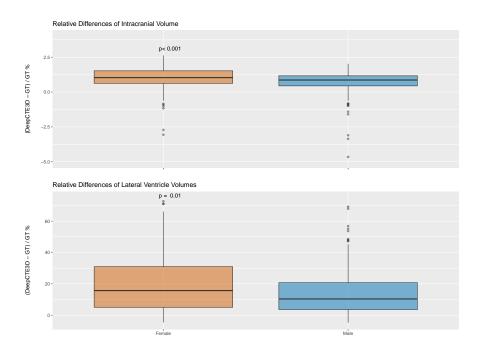
Structure	Group	P-Value	η^2	Magnitude
Intracranial volumes Intracranial volumes Lateral ventricles volumes Lateral ventricle volumes	Patient Sex	<0.001	0.03	Small
	Scanner	<0.001	0.12	Moderate
	Patient Sex	0.03	0.02	Small
	Scanner	<0.001	0.23	Large

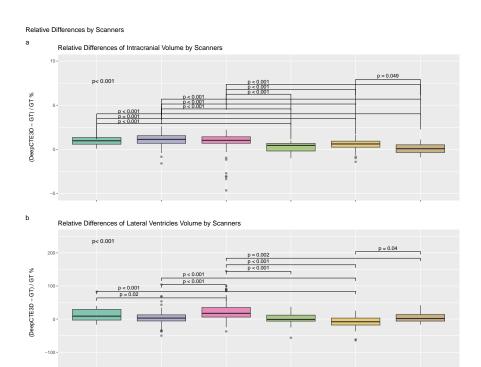
 ${\bf Table~4}~{\bf Effect~Sizes~for~Intracranial~Relative~Differences~Between~Scanner~Type~Pairwise~Comparisons$

Scanner A	Scanner B	P-Value	r	Magnitude
Aquilion	Biograph40	< 0.001	0.49	Moderate
Aquilion	SOMATOM Definition AS+	< 0.001	0.35	Moderate
Aquilion	Discovery 600	< 0.001	0.55	Large
Aquilion ONE	Biograph40	< 0.001	0.35	Moderate
Aquilion ONE	SOMATOM Definition AS+	< 0.001	0.37	Moderate
Aquilion ONE	Discovery 600	< 0.001	0.33	Moderate
Aquilion PRIME	Biograph40	< 0.001	0.28	Small
Aguilion PRIME	SOMATOM Definition AS+	< 0.001	0.31	Moderate
Aquilion PRIME	Discovery 600	< 0.001	0.27	Small
SOMATOM Definition AS+	Discovery 600	0.049	0.24	Small

 ${\bf Table~5} \ \ {\bf Effect~Sizes~for~Lateral~Ventricle~Relative~Differences~Between~Scanner~Type~Pairwise~Comparisons.}$

Scanner A	Scanner B	P-Value	r	Magnitude
Aquilion Aquilion ONE Aquilion ONE Aquilion ONE Aquilion PRIME Aquilion PRIME	Aquilion PRIME SOMATOM Definition AS+ Aquilion PRIME SOMATOM Definition AS+ Biograph40 SOMATOM Definition AS+	0.02 <0.001 <0.001 <0.001 <0.001 <0.001	0.15 0.44 0.41 0.29 0.23 0.50	Small Moderate Moderate Small Small Large
Aquilion PRIME SOMATOM Definition AS+	Discovery 600 Discovery 600	= 0.002 0.04	0.19 0.26	Small Small

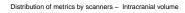


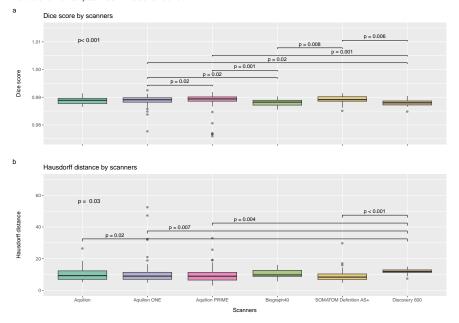


 $\textbf{Fig. 6} \ \ \text{Relative differences by scanners} \ \ \text{a)} \ \ \text{Relative differences for intracranial volume b)} \ \ \text{Relative Difference for lateral ventricle volumes}.$

 ${\bf Table~6}~$ Kruskal-Wallis and Effect Sizes for Metrics by Patient Sex

	P-Value	η^2	Magnitude
Intracranial Dice	0.03	0.01	Small
Intracranial Hausdorff	0.15	<0.01	Small
Ventricular Dice	<0.001	0.02	Small
Ventricular Hausdorff	0.82	<0.01	Small

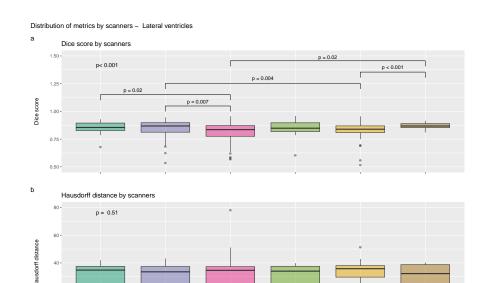




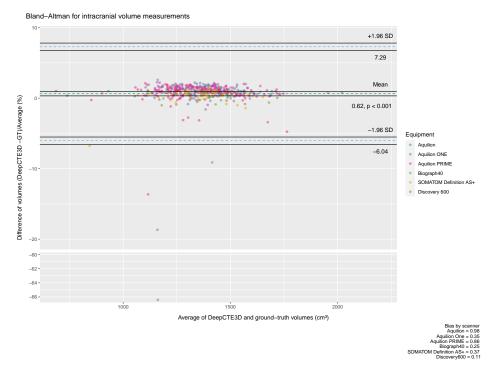
 ${\bf Fig.~7~~Dice~and~Hausdorff~distributions~of~intracranial~segmentation~across~scanner~models.~a)~Dice~Score~across~models.~b)~Hausdorff~Distance~across~models$

 ${\bf Table~7}~{\rm Kruskal\text{-}Wallis~and~Effect~Sizes~for~Metrics~by~Equipment}$

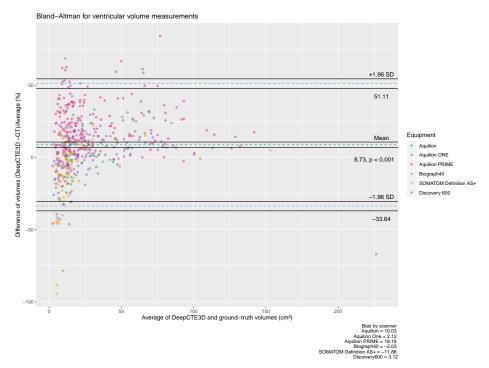
	P-Value	η^2	Magnitude
Intracranial Dice Intracranial Hausdorff Ventricular Dice	< 0.001 0.03 < 0.001	0.04 0.02 0.04	Small Small
Ventricular Hausdorff	0.51	< 0.04	Small



 ${\bf Fig.~8}\ \ {\rm Dice~and~Hausdorff~distributions~of~ventricular~segmentation~across~scanner~models.~a)~{\rm Dice~Score~across~equipment.~b)~Hausdorff~Distance~across~scanner~models}$



 ${\bf Fig.~9}~{\rm Bland\text{-}Altman~plot~assessing~agreement~between~DeepCTE3D\text{-}generated~and~ground\text{-}truth~intracranial~volumes~colored~by~scanner~type.}$



 ${\bf Fig.~10~~Bland-Altman~plot~assessing~agreement~between~DeepCTE3D-generated~and~ground-truth~ventricular~volumes~colored~by~scanner~type.}$

 ${\bf Table~8} \ \ {\bf Bias~and~Effect~Size~by~Equipment~for~Intracranial~Volumes}.$

Scanner A	Scanner B	P-Value	r	Magnitude
Aquilion	Biograph40	< 0.001	0.49	Moderate
Aquilion	SOMATOM Definition AS+	0.001	0.35	Moderate
Aquilion	Discovery 600	< 0.001	0.55	Large
Aquilion ONE	Biograph40	< 0.001	0.35	Moderate
Aquilion ONE	SOMATOM Definition AS+	< 0.001	0.37	Moderate
Aquilion ONE	Discovery 600	< 0.001	0.33	Moderate
Aquilion PRIME	Biograph40	< 0.001	0.28	Small
Aquilion PRIME	SOMATOM Definition AS+	< 0.001	0.31	Moderate
Aquilion PRIME	Discovery 600	< 0.001	0.27	Small
SOMATOM Definition AS+	Discovery 600	0.049	0.24	Small

 ${\bf Table~9} \ \ {\bf Bias~and~Effect~Size~by~Equipment~for~Lateral~Ventricles~Volumes}.$

Scanner A	Scanner B	P-Value	r	Magnitude
Aquilion Aquilion ONE Aquilion ONE Aquilion PRIME Aquilion PRIME	Aquilion PRIME SOMATOM Definition AS+ Aquilion PRIME SOMATOM Definition AS+ Biograph40 SOMATOM Definition AS+	0.02 < 0.001 < 0.001 < 0.001 < 0.001 < 0.001	0.15 0.44 0.41 0.29 0.23 0.50	Small Moderate Moderate Small Small Large
Aquilion PRIME SOMATOM Definition AS+	Discovery 600 Discovery 600	0.002 0.04	0.19 0.26	Small Small

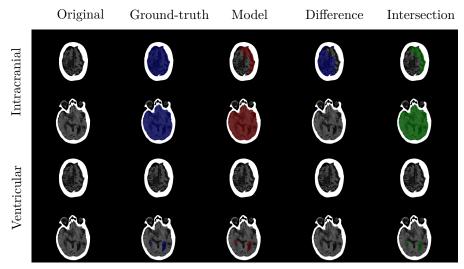


Fig. 11 Example 1 of poor model performance: This case exhibits an angulated visualization of the head, a large frontoparietal encephalomalacia on the right side, and asymmetric ventricular ectasia. Segmentation errors occur predominantly in the high convexity on the same side as the encephalomalacia but also extend into regions with normal brain parenchyma. In the ventricular areas, the model overflowed the boundaries of the lateral ventricles.

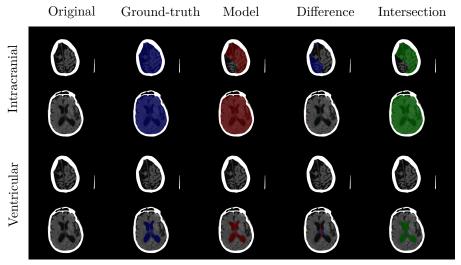


Fig. 12 Example 2 of poor model performance: This case exhibits a focal frontoparietal subarachnoid enlargement on the right side, moderate ventricular ectasia, and a mild degree of parenchymal atrophy. Segmentation errors occur predominantly in the high convexity on the same side as the enlargement but also extend into regions with normal brain parenchyma.

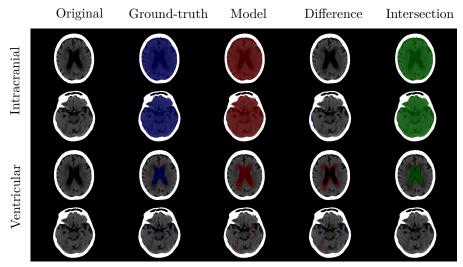


Fig. 13 Example 3 of poor model performance: This case exhibits a greater degree of white matter disease, a moderate degree of parenchymal atrophy with proportional ventricular ectasia, and a small thalamic lacune (not shown). Segmentation errors predominantly occur in the ventricular spaces, overflowing the boundaries, with some subarachnoid spaces also being misclassified (on the right side). The lacune was erroneously segmented as part of the ventricular space.