



Stanford
MEDICINE

Advancements of Artificial Intelligence in Proton Therapy

Serdar Charyyev, PhD

August 24, 2024

OVERVIEW

1

INTRODUCTION

2

AI in PLANNING

4

AI in DELIVERY

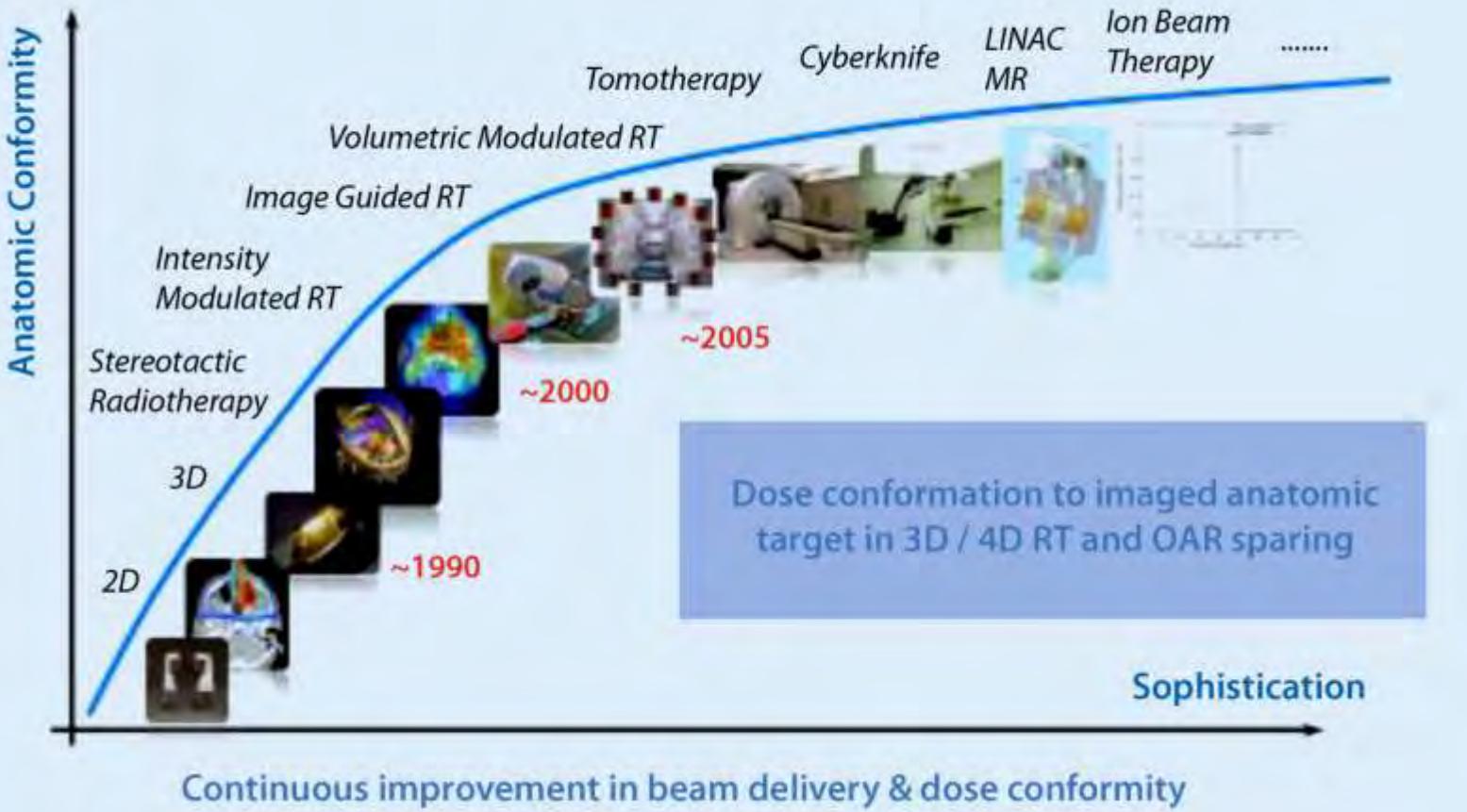
5

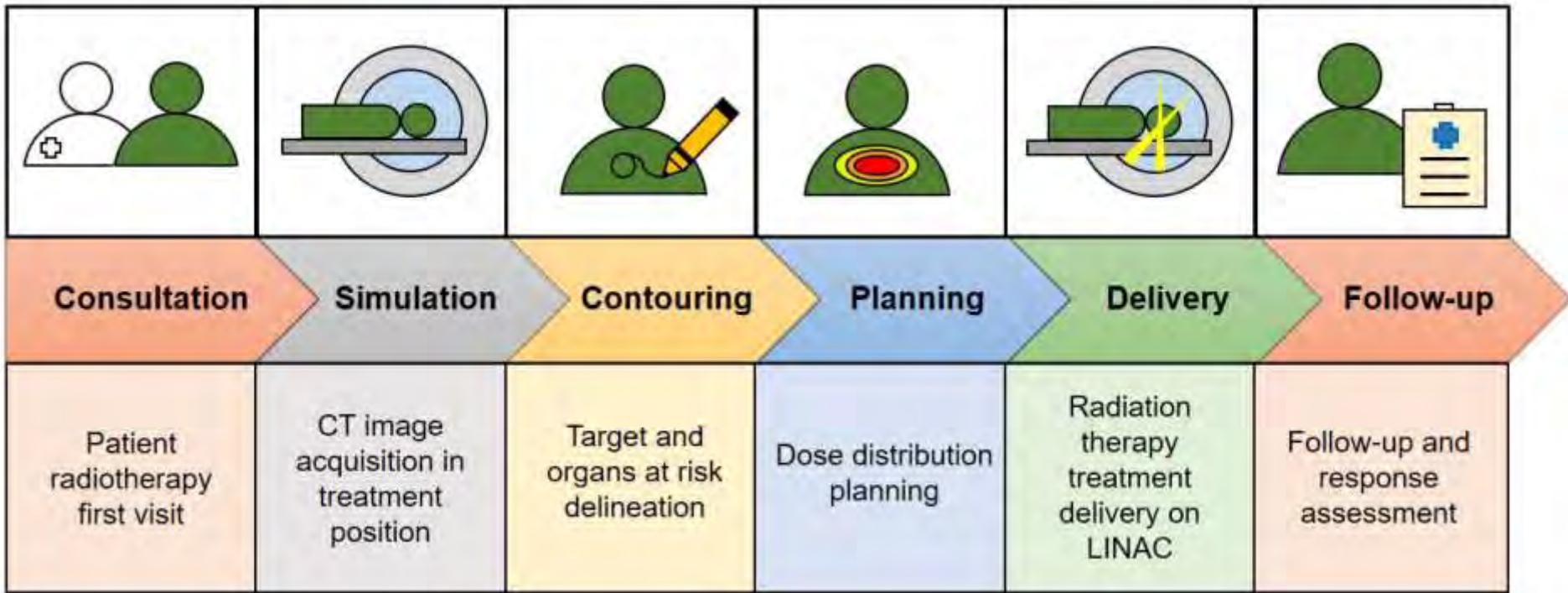
CONCLUSIONS



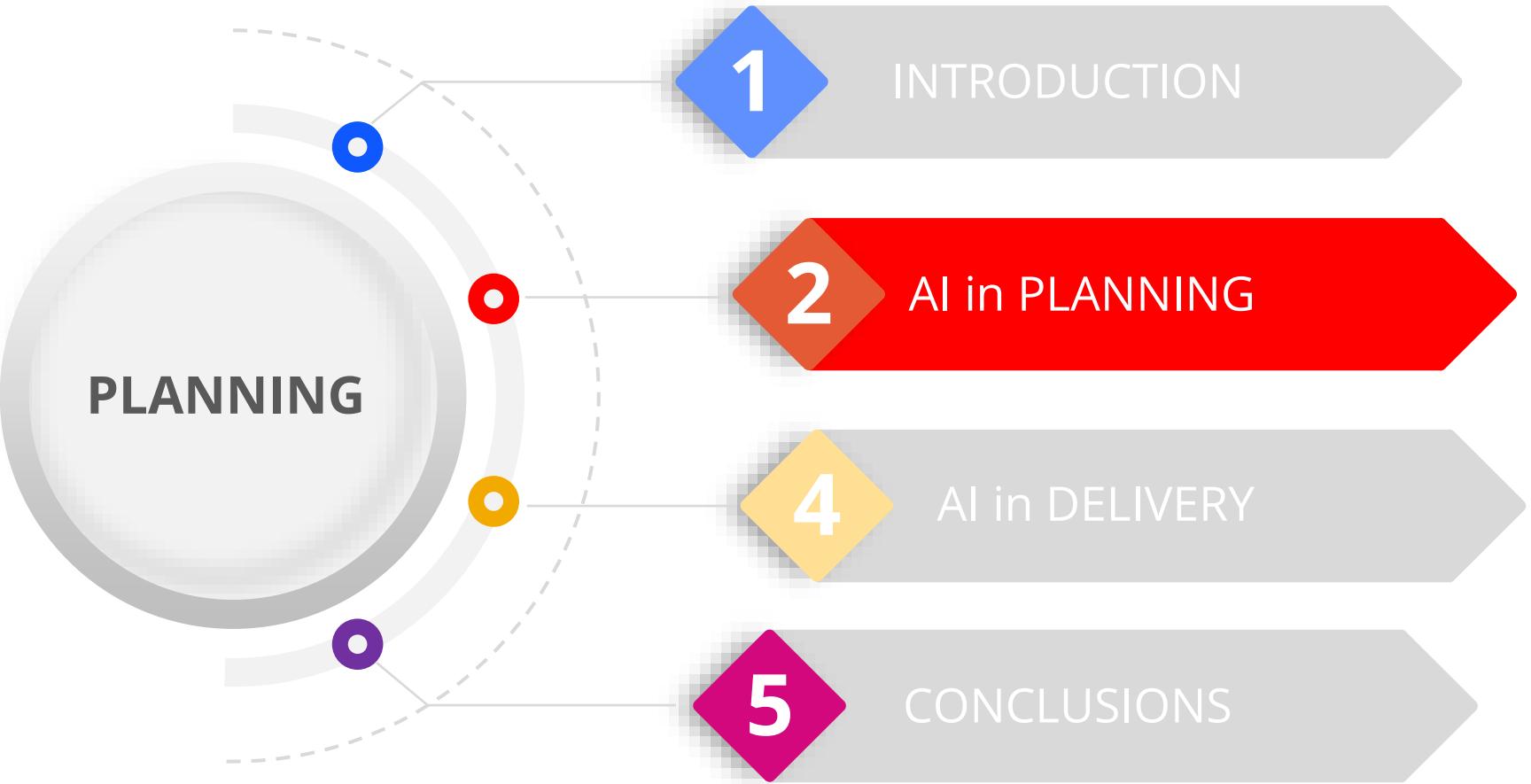
Stanford
MEDICINE

Anatomic Conformity





Marvaso et al., *Applied Sciences*, 12(22), 2022



PLANNING

1

INTRODUCTION

2

AI in PLANNING

4

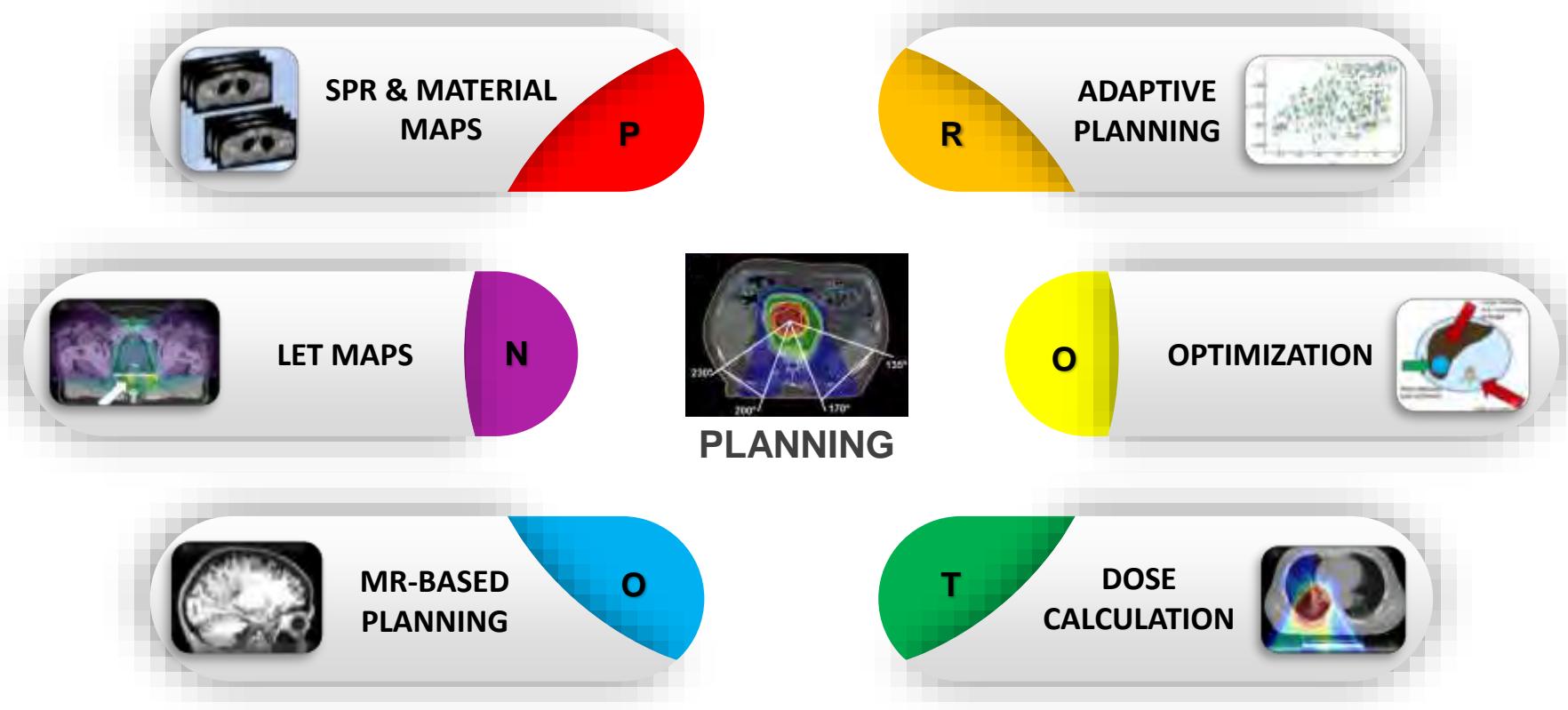
AI in DELIVERY

5

CONCLUSIONS

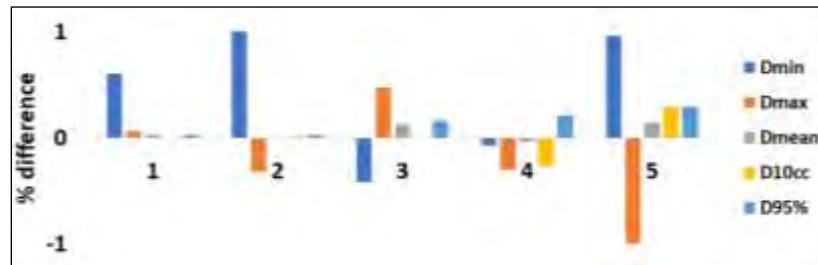
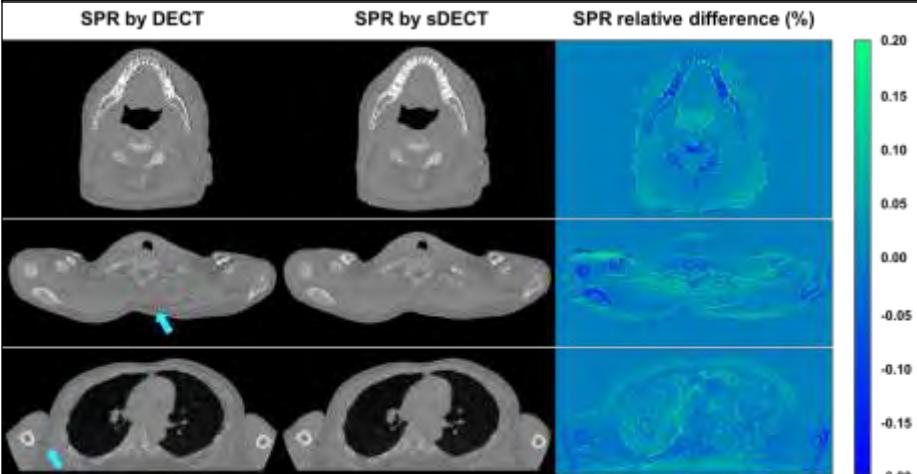
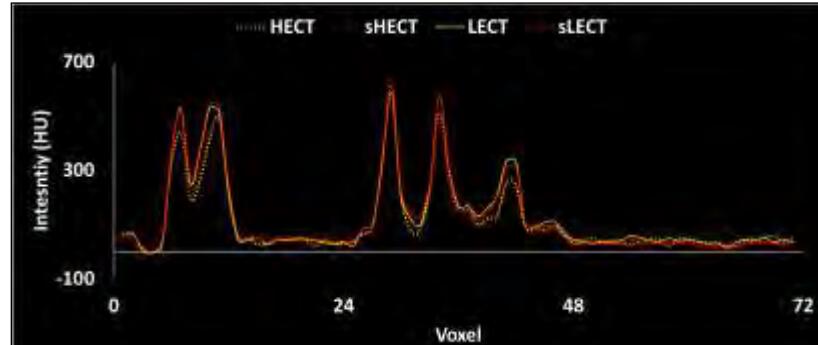
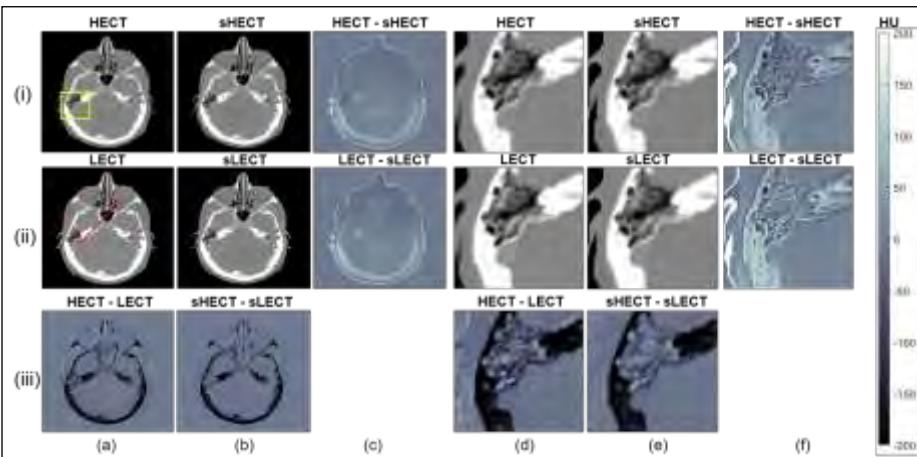


Stanford
MEDICINE



**Stanford
MEDICINE**

SPR & MATERIAL MAPS

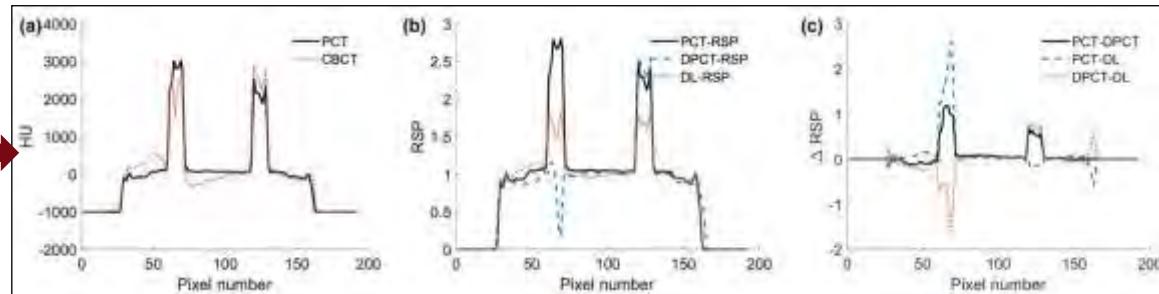
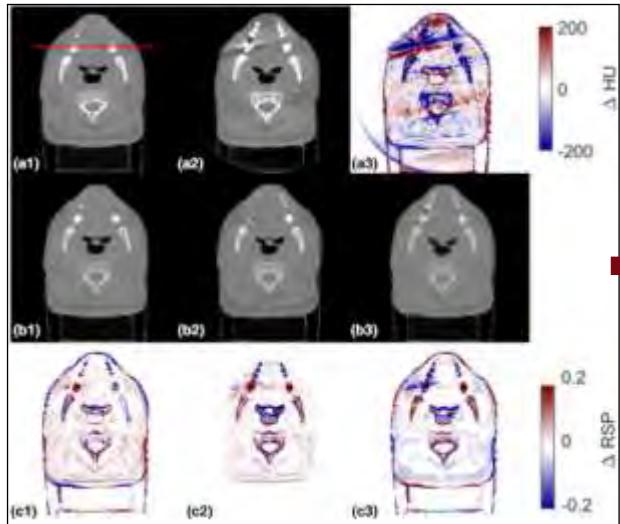


Charyev et al., BJR, 94, 2021
Wang et al., IJPT, 7, 2021

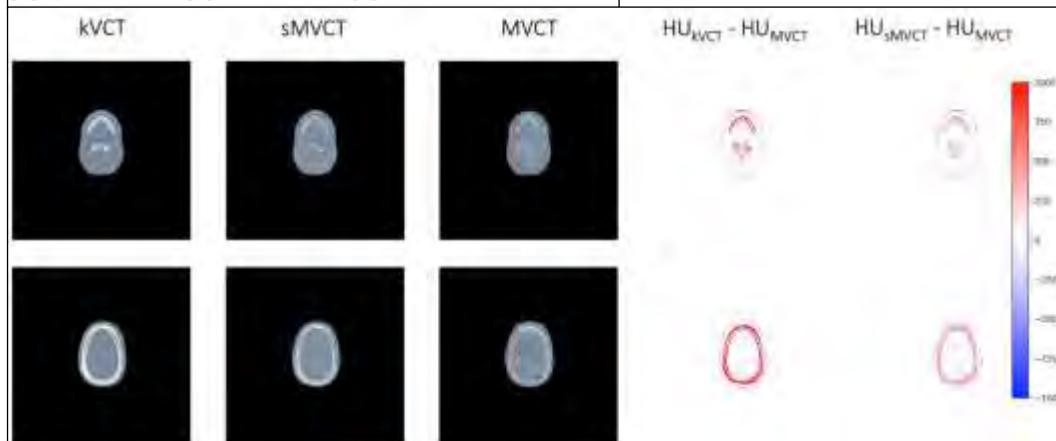


Stanford
MEDICINE

SPR & MATERIAL MAPS



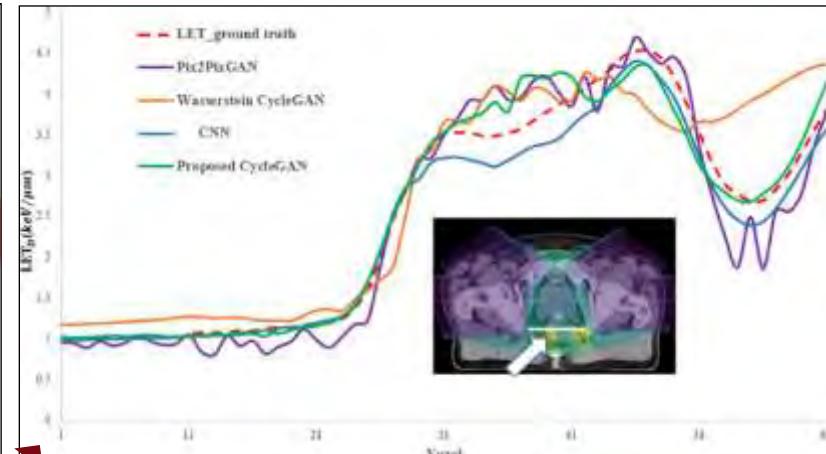
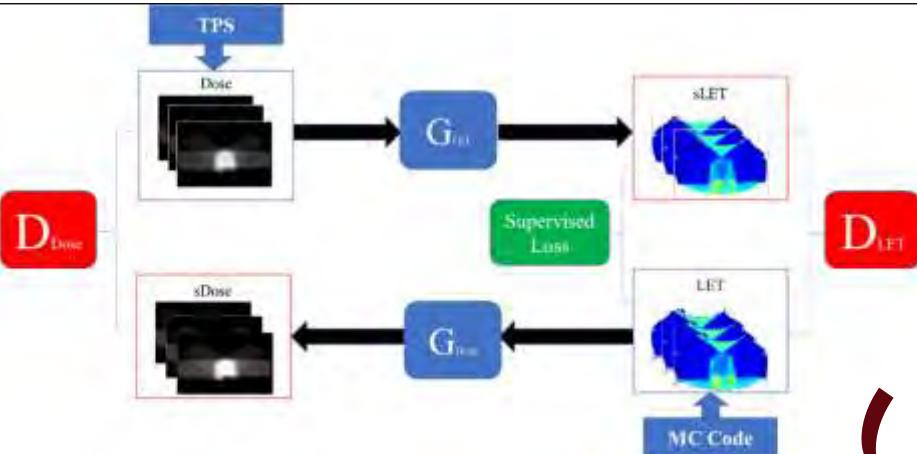
Harms et al., *Med Phys*, 47, 2020



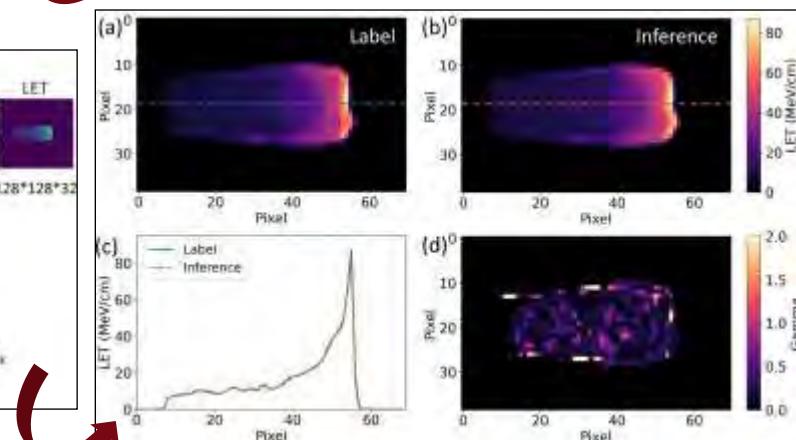
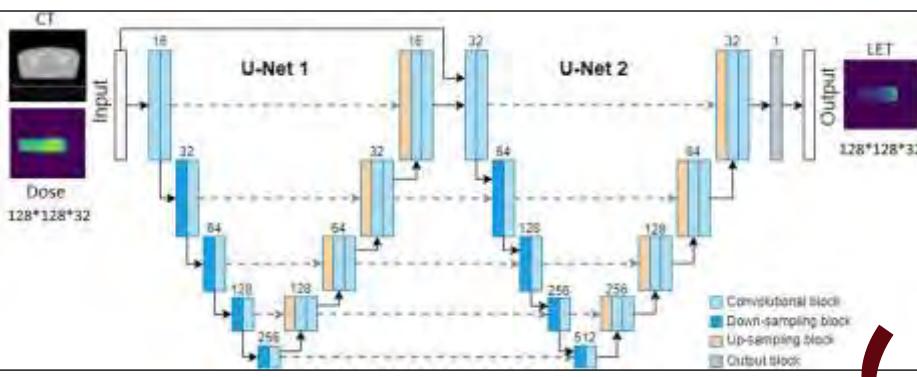
Scholey et al., *PMB*, 67, 2022

- [1] Y. Gao et al., "Single energy CT-based mass density and relative stopping power estimation for proton therapy using deep learning method," *Front. Oncol.*, vol. 13, 2023, Art. no. 1278180.
- [2] S. Charyev et al., "Learning-based synthetic dual energy CT imaging from single energy CT for stopping power ratio calculation in proton radiation therapy," 2020, *Br. J. Radiol.*, 95(1129), doi: 10.1259/bjr.20210644
- [3] T. Wang et al., "Learning-based stopping power mapping on dual energy CT for proton radiation therapy," 2021, *Int. J. Part. Ther.*, 7(3), doi: 10.14338/IJPT-D-20-00020.1
- [4] Y. Liu, L. Zhou, and H. Peng, "Machine learning based oxygen and carbon concentration derivation using dual-energy CT for PETbased dose verification in proton therapy," *Med. Phys.*, vol. 49, no. 5, pp. 3347–3360, 2022, doi: 10.1002/mp.15581.
- [5] C.-W. Chang et al., "Dual-energy CT based mass density and relative stopping power estimation for proton therapy using physics-informed deep learning," *Phys. Med. Biol.* vol. 67, no. 11, 2022, Art. no. 115010, doi: 10.1088/1361-6560/ac6ebc.
- [6] C.-W. Chang et al., "Validation of a deep learning-based material estimation model for Monte Carlo dose calculation in proton therapy," *Phys. Med. Biol.*, vol. 67, no. 21, 2022, Art. no. 215004.
- [7] J. Harms et al., "Cone-beam CT-derived relative stopping power map generation via deep learning for proton radiotherapy," *Med. Phys.*, vol. 47, no. 9, pp. 4416–4427, 2020.
- [8] J. Scholey et al., "Improved accuracy of relative electron density and proton stopping power ratio through CycleGAN machine learning," *Phys. Med. Biol.*, vol. 67, no. 10, 2022, Art. no. 105001.
- [9] C.-W. Chang et al., "Multimodal imaging-based material mass density estimation for proton therapy using supervised deep learning," *Br. J. Radiol.*, vol. 96, no. 1152, 2023, Art. no. 20220907.

LET MAPS



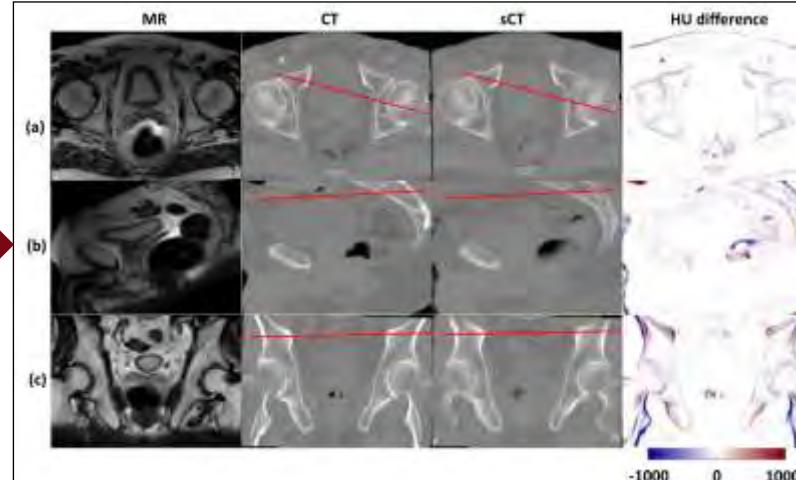
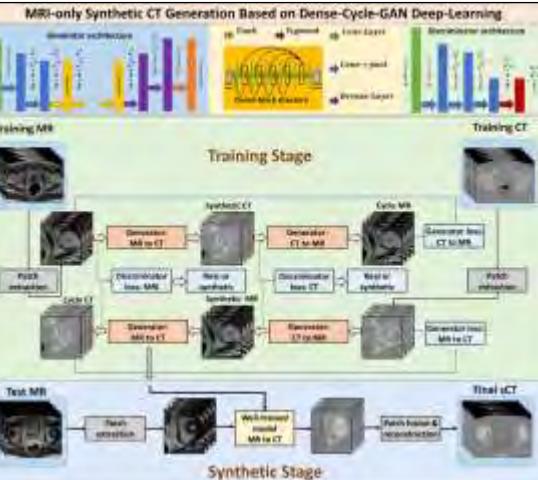
Gao et al., PMB, 69, 2024



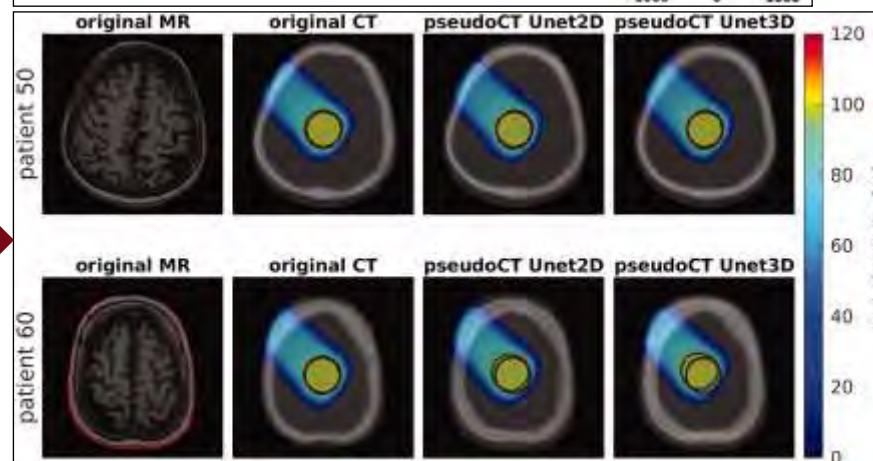
Tang et al., PMB, 69, 2024

- [1] Y. Gao et al., "Deep learning-based synthetic dose-weighted LET map generation for intensity modulated proton therapy," *Phys. Med. Biol.*, vol. 69, no. 2, Art. no. 025004, 2024.
- [2] X. Tang et al., "Deep learning based linear energy transfer calculation for proton therapy," *Phys. Med. Biol.*, vol. 69, no. 11, 2024, doi: 10.1088/1361-6560/ad4844.
- [3] F. Pirlepesov et al., "Three-dimensional dose and LETD prediction in proton therapy using artificial neural networks," *Med. Phys.*, vol. 49, no. 12, pp. 7417–7427, Dec. 2022.
- [4] S. Starke et al., "A deep-learning-based surrogate model for Monte-Carlo simulations of the linear energy transfer in primary brain tumor patients treated with proton-beam radiotherapy," *Phys. Med. Biol.*, vol. 69, no. 14, Art. no. 145007, 2024.
- [5] P. Stasica et al., "Single proton LET characterization with the Timepix detector and artificial intelligence for advanced proton therapy treatment planning," *Phys. Med. Biol.*, vol. 68, no. 10, Art. no. 105001, 2023.

MR-BASED PLANNING



Liu et al., PMB, 64, 2019

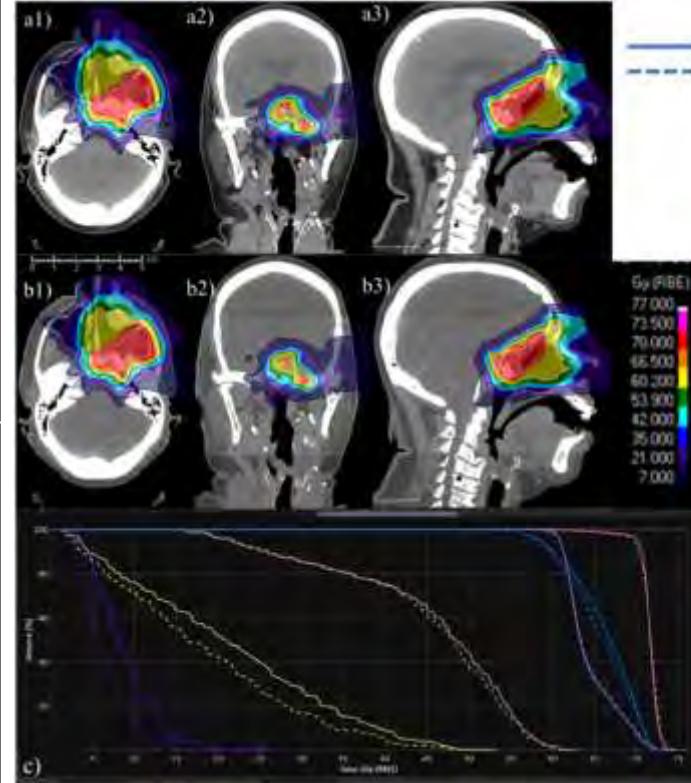
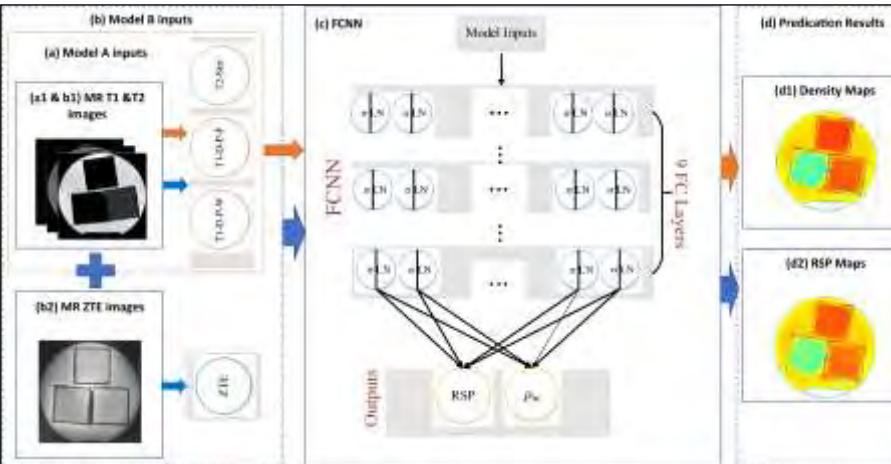


Neppl et al., Acta Oncologica, 58, 2019

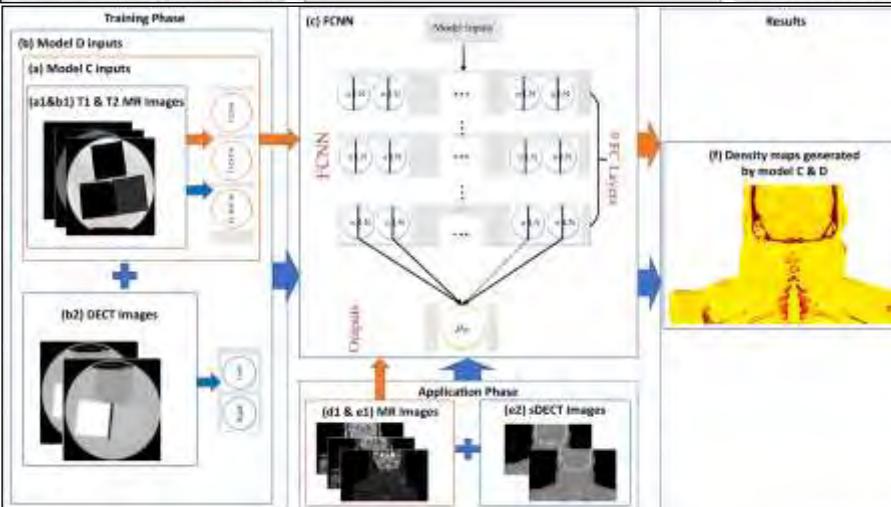


Stanford
MEDICINE

MR-BASED PLANNING



Clinical TPCT
MRI & sDECT



[1] Y. Liu et al., "MRI-based treatment planning for proton radiotherapy: Dosimetric validation of a deep learning-based liver synthetic CT generation method," *Phys. Med. Biol.*, vol. 64, no. 14, 2019, Art. no. 145015.

[2] Y. Liu et al., "Evaluation of a deep learning-based pelvic synthetic CT generation technique for MRI-based prostate proton treatment planning," *Phys. Med. Biol.*, vol. 64, no. 20, 2019, Art. no. 205022.

[3] S. Kazemifar et al., "Dosimetric evaluation of synthetic CT generated with GANs for MRI-only proton therapy treatment planning of brain tumors," *J. Appl. Clin. Med. Phys.*, vol. 21, no. 5, pp. 76–86, 2020.

[4] M. F. Spadea et al., "Deep Convolution Neural Network (DCNN) multiplane approach to synthetic CT generation from MR images—Application in brain proton therapy," *Int. J. Radiat. Oncol. Biol. Phys.*, vol. 105, no. 3, pp. 495–503, 2019.

[5] S. Neppl et al., "Evaluation of proton and photon dose distributions recalculated on 2D and 3D Unet-generated pseudoCTs from T1-weighted MR head scans," *Acta Oncologica*, vol. 58, no. 10, pp. 1429–1434, 2019.

[6] M. C. Florkow et al., "Deep learning-enabled MRI-only photon and proton therapy treatment planning for paediatric abdominal tumours," *Radiother. Oncol.*, vol. 153, pp. 220–227, Dec. 2020, doi: 10.1016/j.radonc.2020.09.056.

[7] M. Maspero et al., "Deep learning-based synthetic CT generation for paediatric brain MR-only photon and proton radiotherapy," *Radiother. Oncol.*, vol. 153, pp. 197–204, Dec. 2020.

[8] G. Shafai-Erfani et al., "MRI-based proton treatment planning for base of skull tumors," *Int. J. Partic. Ther.*, vol. 6, no. 2, pp. 12–25, 2019.

[9] J. M. Wolterink, A. M. Dinkla, M. H. F. Savenije, P. R. Seevinck, C. A. T. van den Berg, and I. Işgum, "Deep MR to CT synthesis using unpaired data," in *Simulation and Synthesis in Medical Imaging (SASHIMI) (Lecture Notes in Computer Science)*, S. Tsafaris, A. Gooya, A. Frangi, and J. Prince, Eds., vol. 10557. Cham, Switzerland: Springer, 2017. [Online]. Available: https://doi.org/10.1007/978-3-319-68127-6_2

[10] Zimmermann L, Buschmann M, Herrmann H, Heilemann G, Kuess P, Goldner G, Nyholm T, Georg D, Nesvacil N. An MR-only acquisition and artificial intelligence based image-processing protocol for photon and proton therapy using a low field MR. *Med Phys.* 2021 Feb;31(1):78-88. doi: 10.1016/j.zemedi.2020.10.004. Epub 2021 Jan 15. PMID: 33455822.

[11] Zimmermann L, Knäsl B, Stock M, Lütgendorf-Caucig C, Georg D, Kuess P. An MRI sequence independent convolutional neural network for synthetic head CT generation in proton therapy. *Z Med Phys.* 2022 May;32(2):218-227. doi: 10.1016/j.zemedi.2021.10.003. Epub 2021 Dec 15. PMID: 34920940; PMCID: PMC9948837.

[12] Chen S, Peng Y, Qin A, Liu Y, Zhao C, Deng X, Deraniyagala R, Stevens C, Ding X. MR-based synthetic CT image for intensity-modulated proton treatment planning of nasopharyngeal carcinoma patients. *Acta Oncol.* 2022 Nov;61(11):1417-1424. doi: 10.1080/0284186X.2022.2140017. Epub 2022 Oct 28. PMID: 36305424.

[13] Liu, Ruirui, et al. "Synthetic dual-energy CT for MRI-only based proton therapy treatment planning using label-GAN." *Physics in Medicine & Biology* 66.6 (2021): 065014.

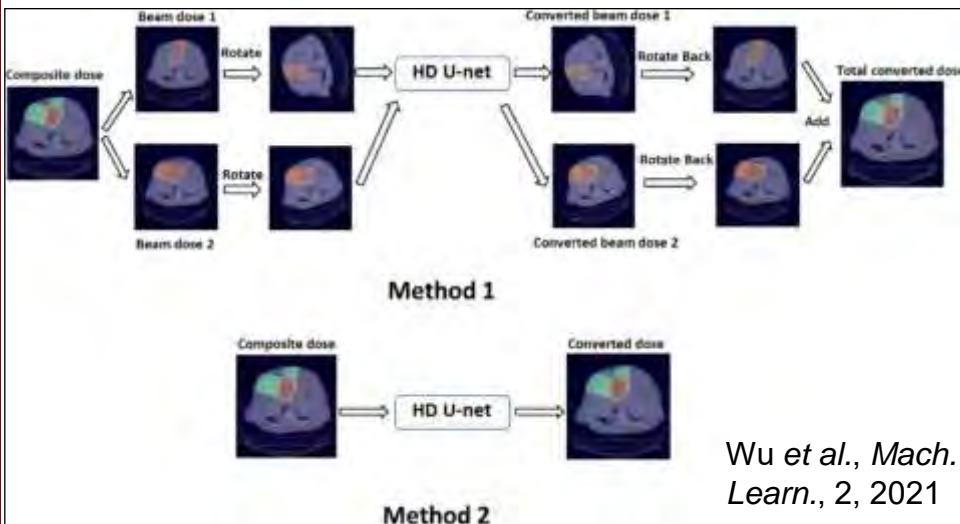
[14] Li X, Bellotti R, Meier G, Bachtiary B, Weber D, Lomax A, Buhmann J, Zhang Y. Uncertainty-aware MR-based CT synthesis for robust proton therapy planning of brain tumour. *Radiother Oncol.* 2024 Feb;191:110056. doi: 10.1016/j.radonc.2023.110056. Epub 2023 Dec 15. PMID: 38104781.

[15] Gao, Yuan, et al. "MRI-only based material mass density and relative stopping power estimation via deep learning for proton therapy: a preliminary study." *Scientific Reports* 14.1 (2024): 11166.

[16] Wang, Chuang, et al. "Toward MR-only proton therapy planning for pediatric brain tumors: Synthesis of relative proton stopping power images with multiple sequences and development of an online quality assurance tool." *Medical physics* 49.3 (2022): 1559-1570.



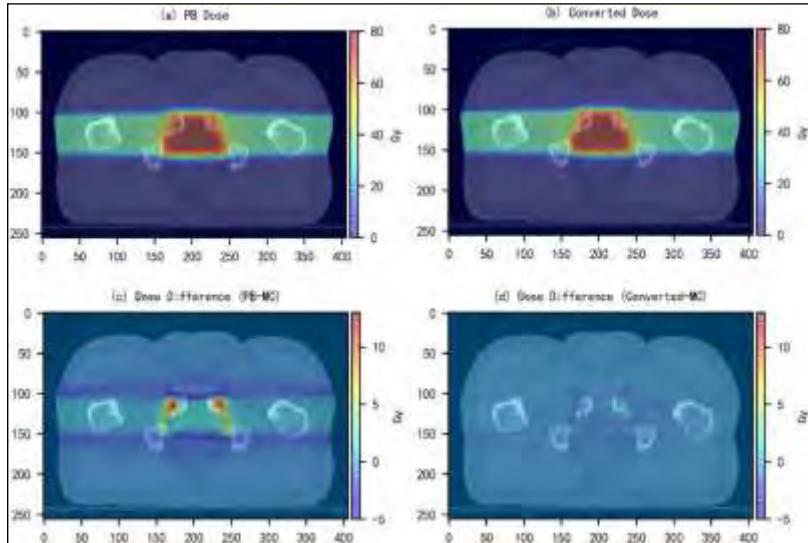
DOSE CALCULATION



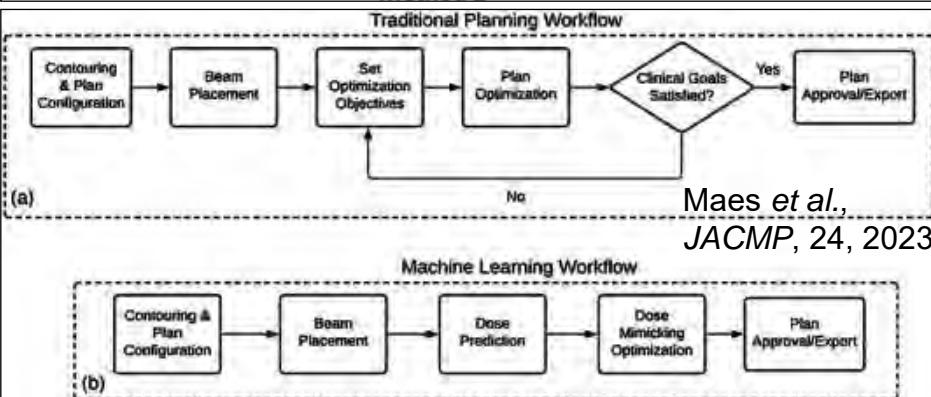
Method 1

Method 2

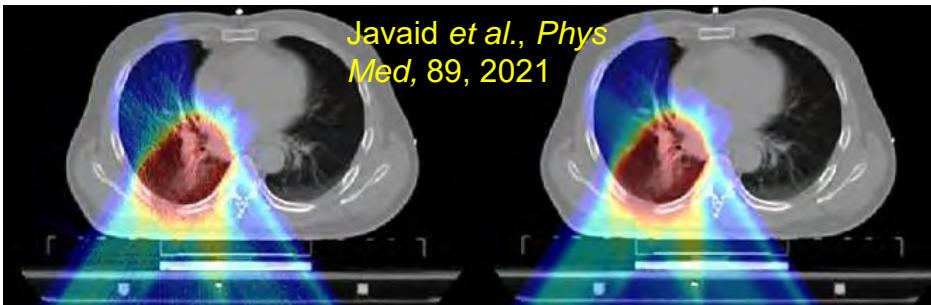
Wu et al., *Mach. Learn.*, 2, 2021



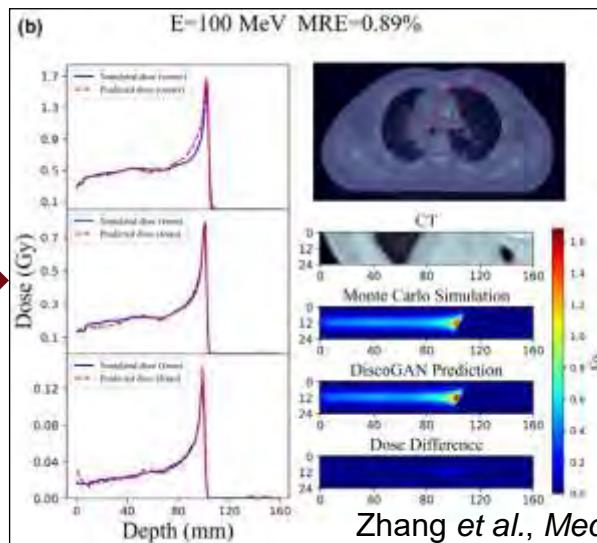
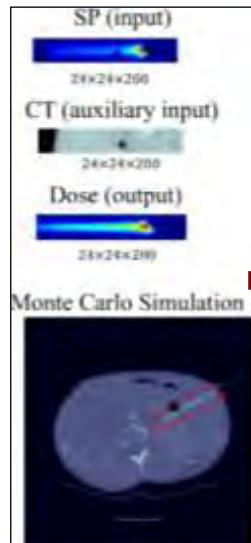
Javaid et al., *Phys. Med.*, 89, 2021



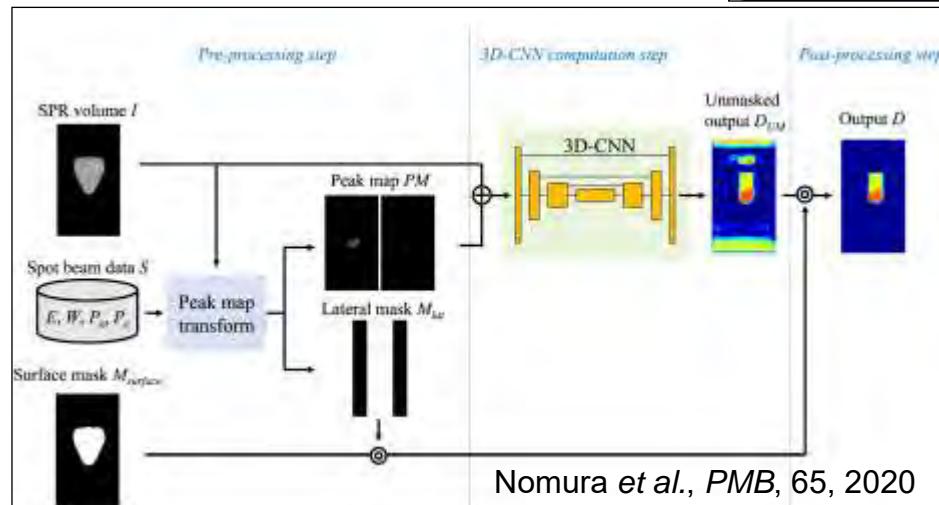
Maes et al.,
JACMP, 24, 2023



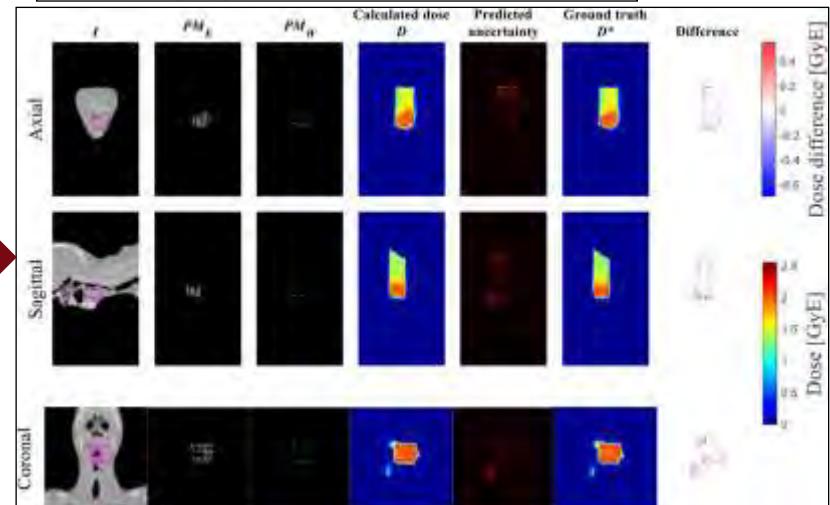
DOSE CALCULATION



Zhang et al., Med Phys, 48, 2021

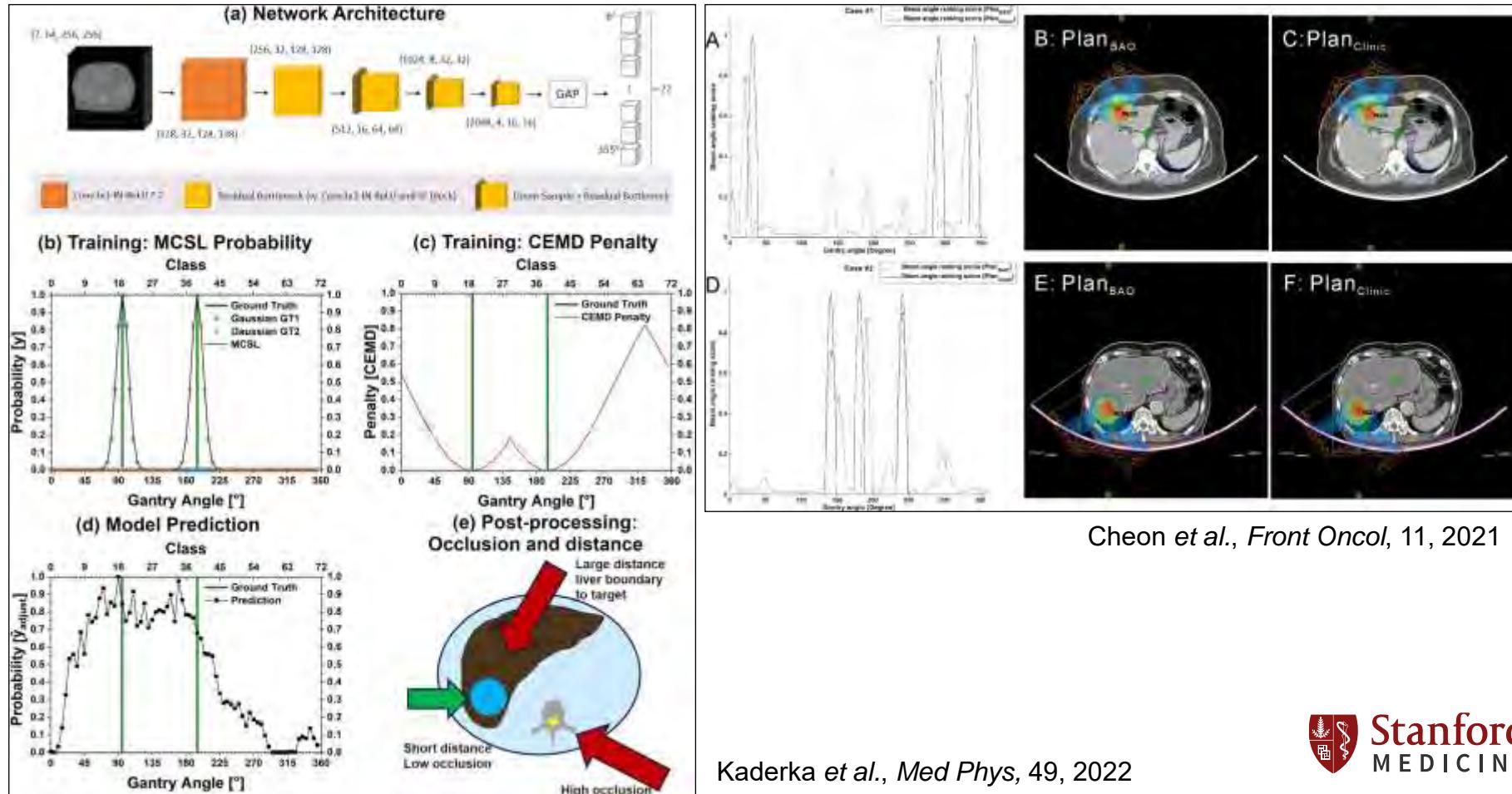


Nomura et al., PMB, 65, 2020



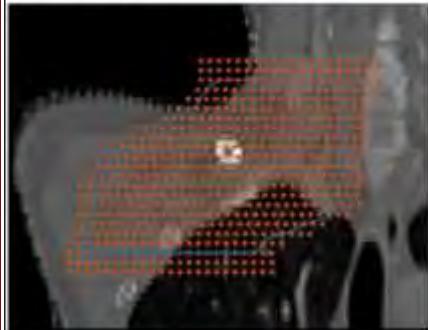
- [1] C. Wu et al., "Improving proton dose calculation accuracy by using deep learning," *Mach. Learn. Sci. Technol.*, vol. 2, no. 1, 2020, Art. no. 15017.
- [2] Y. Nomura, J. Wang, H. Shirato, S. Shimizu, and L. Xing, "Fast spot-scanning proton dose calculation method with uncertainty quantification using a three-dimensional convolutional neural network," *Phys. Med. Biol.*, vol. 65, no. 21, 2020, Art. no. 215007.
- [3] A. Neishabouri, N. Wahl, A. Mairani, U. Köthe, and M. Bangert, "Long short-term memory networks for proton dose calculation in highly heterogeneous tissues," *Med. Phys.*, vol. 48, pp. 1893–1908, Apr. 2021.
- [4] D. Nguyen et al., "3D radiotherapy dose prediction on head and neck cancer patients with a hierarchically densely connected U-net deep learning architecture," *Phys. Med. Biol.*, vol. 64, no. 6, 2019, Art. no. 65020.
- [5] X. K. Zhang, Z. Hu, G. Zhang, Y. Zhuang, Y. Wang, and H. Peng, "Dose calculation in proton therapy using a discovery cross-domain generative adversarial network (DiscoGAN)," *Med. Phys.*, vol. 48, no. 5, pp. 2646–2660, 2021, doi: 10.1002/mp.14781.
- [6] Chen M, Pang B, Zeng Y, Xu C, Chen J, Yang K, Chang Y, Yang Z. Evaluation of an automated clinical decision system with deep learning dose prediction and NTCP model for prostate cancer proton therapy. *Phys Med Biol*. 2024 May 30;69(11). doi: 10.1088/1361-6560/ad48f6. PMID: 38718814.
- [7] Wang X, Hao Y, Duan Y, Yang D. A deep learning approach to remove contrast from contrast-enhanced CT for proton dose calculation. *J Appl Clin Med Phys*. 2024 Feb;25(2):e14266. doi: 10.1002/acm2.14266. Epub 2024 Jan 25. PMID: 38269961; PMCID: PMC10860532.
- [8] Pastor-Serrano O, Perkó Z. Millisecond speed deep learning based proton dose calculation with Monte Carlo accuracy. *Phys Med Biol*. 2022 May 9;67(10). doi: 10.1088/1361-6560/ac692e. PMID: 35447605.
- [9] Wang W, Chang Y, Liu Y, Liang Z, Liao Y, Qin B, Liu X, Yang Z. Feasibility study of fast intensity-modulated proton therapy dose prediction method using deep neural networks for prostate cancer. *Med Phys*. 2022 Aug;49(8):5451-5463. doi: 10.1002/mp.15702. Epub 2022 May 19. PMID: 35543109; PMCID: PMC10358316.
- [10] Zhang L, Holmes JM, Liu Z, Vora SA, Sio TT, Vargas CE, Yu NY, Keole SR, Schild SE, Bues M, Li S, Liu T, Shen J, Wong WW, Liu W. Beam mask and sliding window-facilitated deep learning-based accurate and efficient dose prediction for pencil beam scanning proton therapy. *Med Phys*. 2024 Feb;51(2):1484-1498. doi: 10.1002/mp.16758. Epub 2023 Sep 25. PMID: 37748037.
- [11] Maes D, Holmstrom M, Helander R, Saini J, Fang C, Bowen SR. Automated treatment planning for proton pencil beam scanning using deep learning dose prediction and dose-mimicking optimization. *J Appl Clin Med Phys*. 2023 Oct;24(10):e14065. doi: 10.1002/acm2.14065. Epub 2023 Jun 19. PMID: 37334746; PMCID: PMC10562035.
- [12] Javaid U, Souris K, Huang S, Lee JA. Denoising proton therapy Monte Carlo dose distributions in multiple tumor sites: A comparative neural networks architecture study. *Phys Med*. 2021 Sep;89:93-103. doi: 10.1016/j.ejmp.2021.07.022. Epub 2021 Aug 3. PMID: 34358755.
- [13] Guerreiro F, Seravalli E, Janssens GO, Maduro JH, Knopf AC, Langendijk JA, Raaymakers BW, Kontaxis C. Deep learning prediction of proton and photon dose distributions for paediatric abdominal tumours. *Radiother Oncol*. 2021 Mar;156:36-42. doi: 10.1016/j.radonc.2020.11.026. Epub 2020 Nov 29. PMID: 33264639.
- [14] Mentzel F, Kröninger K, Lerch M, Nackenhorst O, Rosenfeld A, Tsui AC, Weingarten J, Hagenbuchner M, Guatelli S. Small beams, fast prediction: a comparison of machine learning dose prediction models for proton minibeam therapy. *Med Phys*. 2022 Dec;49(12):7791-7801. doi: 10.1002/mp.16066. Epub 2022 Nov 10. PMID: 36309820.

OPTIMIZATION

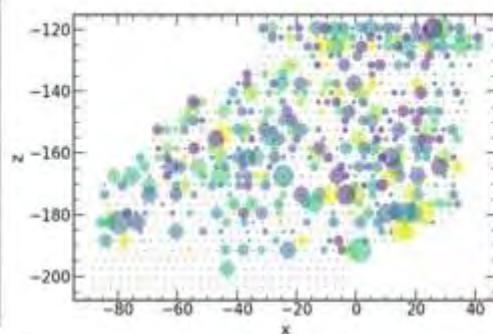


ADAPTIVE PLANNING

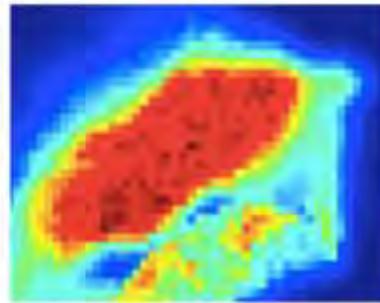
PTV Contour + Spot Pattern



Spot Weight Reference



Dose Map



Training Phase

Train Dose-map



Residual Dose-map

3D-ResNet Backbone



Loss 123*13

Residual weight

Resize

Spot pattern and Train weights

Spot pattern and Reference weights

Estimated Residual weight

Resize

Spot pattern and Fine-tuned weights

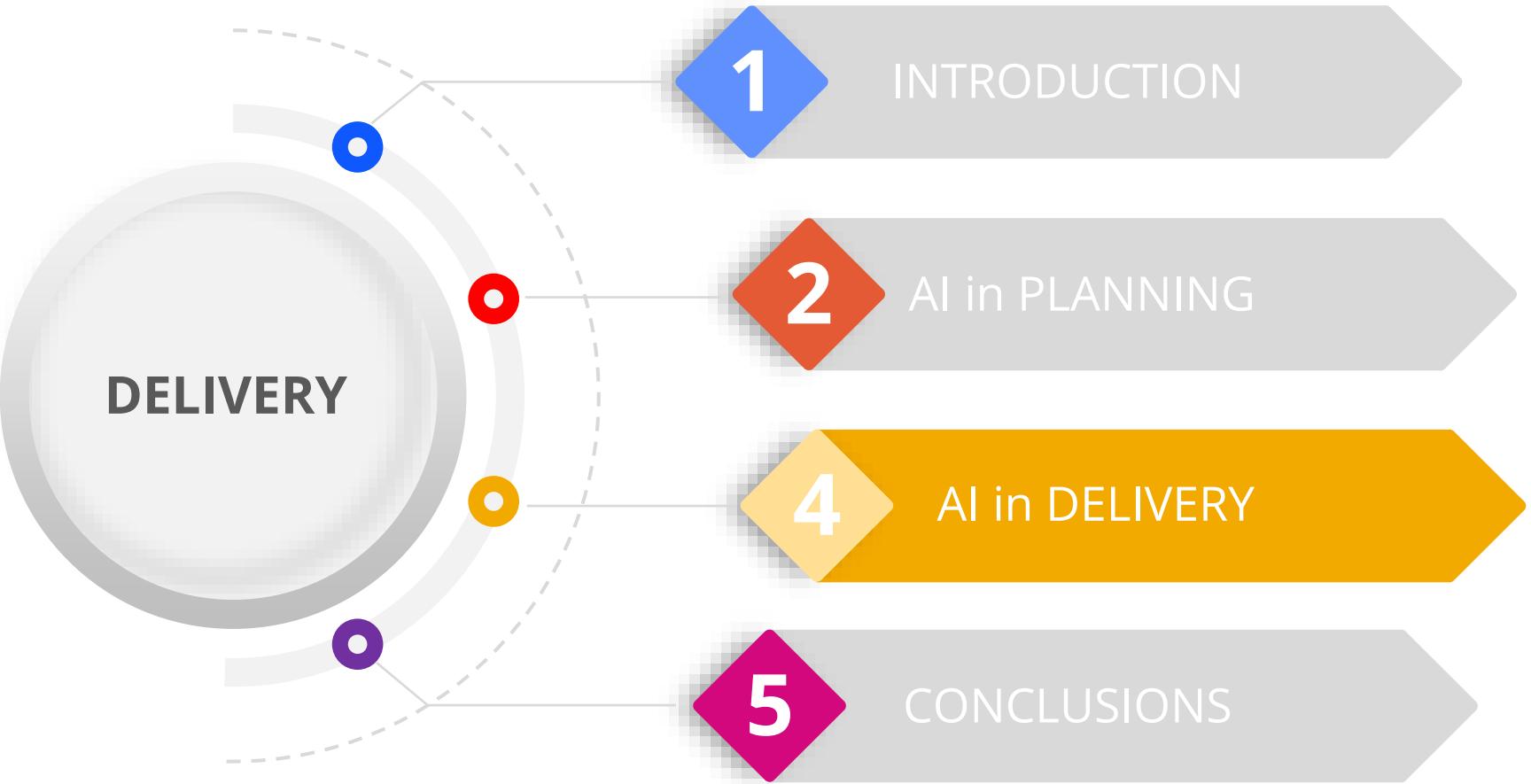
Testing Phase

Zhang et al., PMB, 67, 2022



Stanford
MEDICINE

- [1] Vazquez I, Gronberg MP, Zhang X, Court LE, Zhu XR, Frank SJ, Yang M. A deep learning-based approach for statistical **robustness evaluation** in proton therapy treatment planning: a feasibility study. *Phys Med Biol.* 2023 Apr 26;68(9). doi: 10.1088/1361-6560/accc08. PMID: 37040785.
- [2] Viar-Hernandez D, Molina-Maza JM, Vera-Sánchez JA, Perez-Moreno JM, Mazal A, Rodriguez-Vila B, Malpica N, Torrado-Carvajal A. Enhancing adaptive proton therapy through **CBCT** images: **Synthetic head and neck CT generation** based on 3D vision transformers. *Med Phys.* 2024 Jul;51(7):4922-4935. doi: 10.1002/mp.17057. Epub 2024 Apr 3. PMID: 38569141.
- [3] Pang B, Si H, Liu M, Fu W, Zeng Y, Liu H, Cao T, Chang Y, Quan H, Yang Z. Comparison and evaluation of different deep learning models of **synthetic CT generation from CBCT** for nasopharynx cancer adaptive proton therapy. *Med Phys.* 2023 Nov;50(11):6920-6930. doi: 10.1002/mp.16777. Epub 2023 Oct 6. PMID: 37800874.
- [4] Zhang G, Zhou L, Han Z, Zhao W, Peng H. SWFT-Net: a deep learning framework for efficient **fine-tuning spot weights** towards adaptive proton therapy. *Phys Med Biol.* 2022 Dec 9;67(24). doi: 10.1088/1361-6560/aca517. PMID: 36541496.
- [5] Zhang G, Chen X, Dai J, Men K. A plan verification platform for online adaptive proton therapy using deep learning-based **Monte-Carlo denoising**. *Phys Med.* 2022 Nov;103:18-25. doi: 10.1016/j.ejmp.2022.09.018. Epub 2022 Oct 3. PMID: 36201903.
- [6] Wang C, Uh J, Merchant TE, Hua CH, Acharya S. Facilitating **MR-Guided** Adaptive Proton Therapy in Children Using Deep Learning-Based **Synthetic CT**. *Int J Part Ther.* 2021 Jun 25;8(3):11-20. doi: 10.14338/IJPT-20-00099.1. PMID: 35127971; PMCID: PMC8768893.
- [7] de Koster RJC, Thummerer A, Scandurra D, Langendijk JA, Both S. Technical note: **Evaluation of deep learning based synthetic CTs** clinical readiness for dose and NTCP driven head and neck adaptive proton therapy. *Med Phys.* 2023 Dec;50(12):8023-8033. doi: 10.1002/mp.16782. Epub 2023 Oct 13. PMID: 37831597.
- [8] Thummerer A, Seller Oriá C, Zaffino P, Meijers A, Guterres Marmitt G, Wijsman R, Seco J, Langendijk JA, Knopf AC, Spadea MF, Both S. Clinical suitability of deep learning based **synthetic CTs** for adaptive proton therapy of lung cancer. *Med Phys.* 2021 Dec;48(12):7673-7684. doi: 10.1002/mp.15333. Epub 2021 Nov 16. PMID: 34725829; PMCID: PMC9299115.
- [8] Kaushik S, Ödén J, Sharma DS, Fredriksson A, Toma-Dasu I. Generation and evaluation of anatomy-preserving **virtual CT** for online adaptive proton therapy. *Med P9ys.* 2024 Mar;51(3):1536-1546. doi: 10.1002/mp.16941. Epub 2024 Jan 17. PMID: 38230803.
- [10] Lalonde A, Winey B, Verburg J, Paganetti H, Sharp GC. Evaluation of **CBCT scatter correction** using deep convolutional neural networks for head and neck adaptive proton therapy. *Phys Med Biol.* 2020 Dec 4;65(24):10.1088/1361-6560/ab9fcf. doi: 10.1088/1361-6560/ab9fcf. Erratum in: *Phys Med Biol.* 2021 Jun 30;66(13). doi: 10.1088/1361-6560/ac0cc2. PMID: 32580174; PMCID: PMC8920050.
- [11] Thummerer A, Seller Oriá C, Zaffino P, Visser S, Meijers A, Guterres Marmitt G, Wijsman R, Seco J, Langendijk JA, Knopf AC, Spadea MF, Both S. Deep learning-based **4D-synthetic CTs** from sparse-view CBCTs for dose calculations in adaptive proton therapy. *Med Phys.* 2022 Nov;49(11):6824-6839. doi: 10.1002/mp.15930. Epub 2022 Aug 27. PMID: 35982630; PMCID: PMC10087352.
- [12] Kaderka R, Liu KC, Liu L, VanderStraeten R, Liu TL, Lee KM, Tu YE, MacEwan I, Simpson D, Urbanic J, Chang C. Toward **automatic beam angle selection** for pencil-beam scanning proton liver treatments: A deep learning-based approach. *Med Phys.* 2022 Jul;49(7):4293-4304. doi: 10.1002/mp.15676. Epub 2022 May 11. PMID: 35488864.
- [13] Cheon W, Ahn SH, Jeong S, Lee SB, Shin D, Lim YK, Jeong JH, Youn SH, Lee SU, Moon SH, Kim TH, Kim H. **Beam Angle Optimization** for Double-Scattering Proton Delivery Technique Using an Eclipse Application Programming Interface and Convolutional Neural Network. *Front Oncol.* 2021 Sep 14;11:707464. doi: 10.3389/fonc.2021.707464. PMID: 345112 PMCID: PMC8471903.



DELIVERY

1

INTRODUCTION

2

AI in PLANNING

4

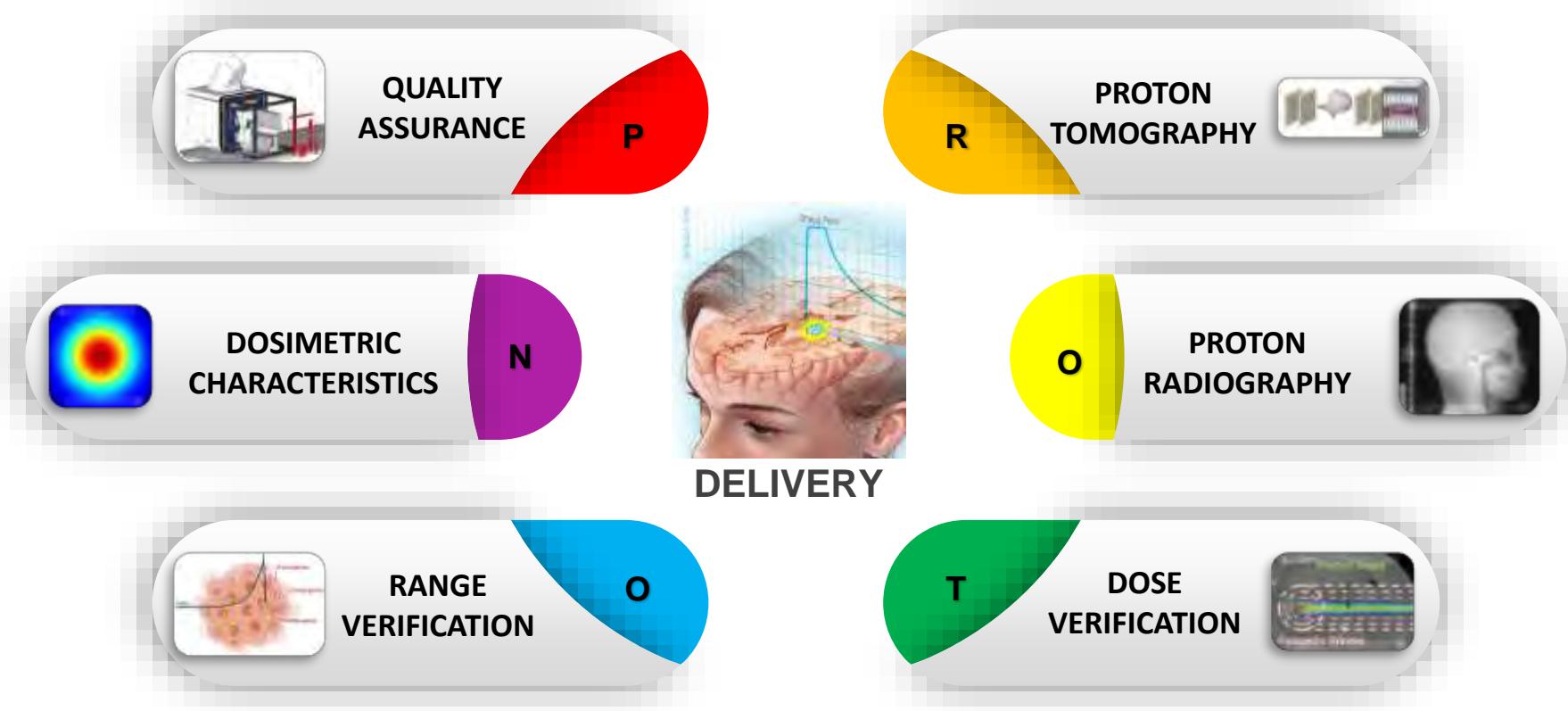
AI in DELIVERY

5

CONCLUSIONS

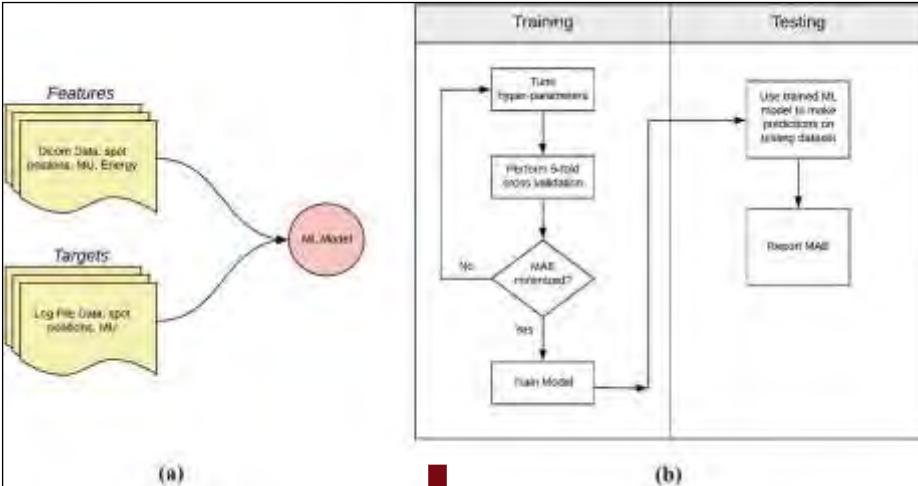
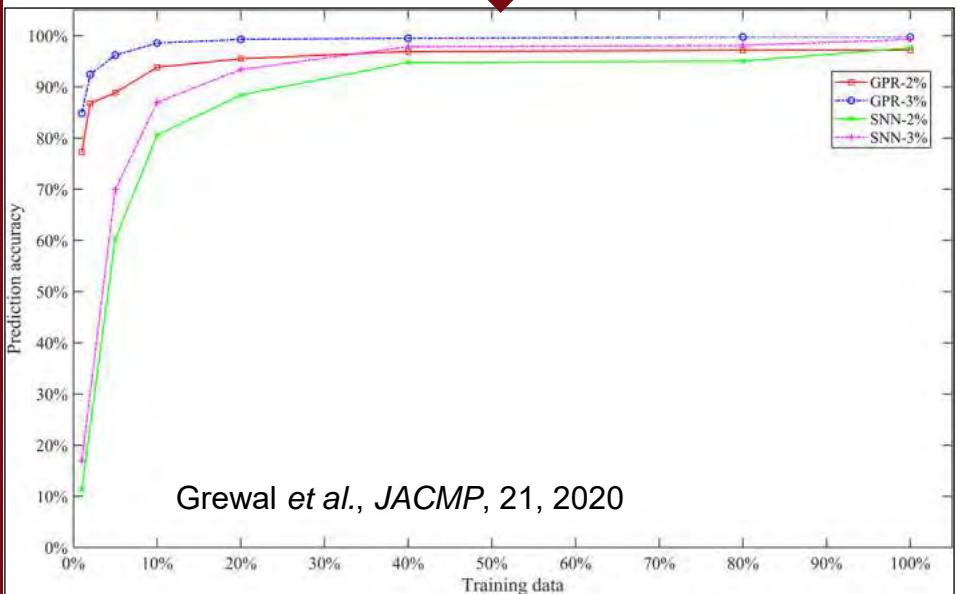
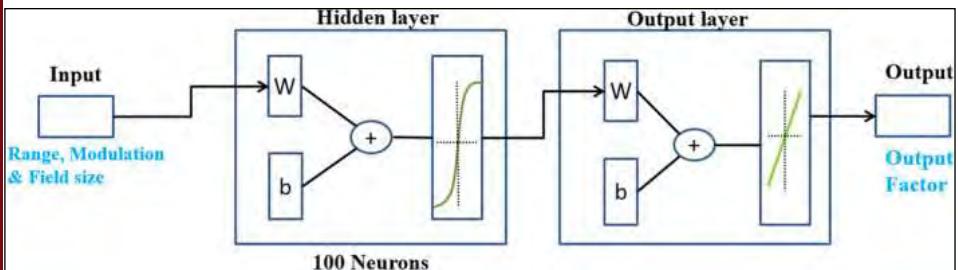


Stanford
MEDICINE

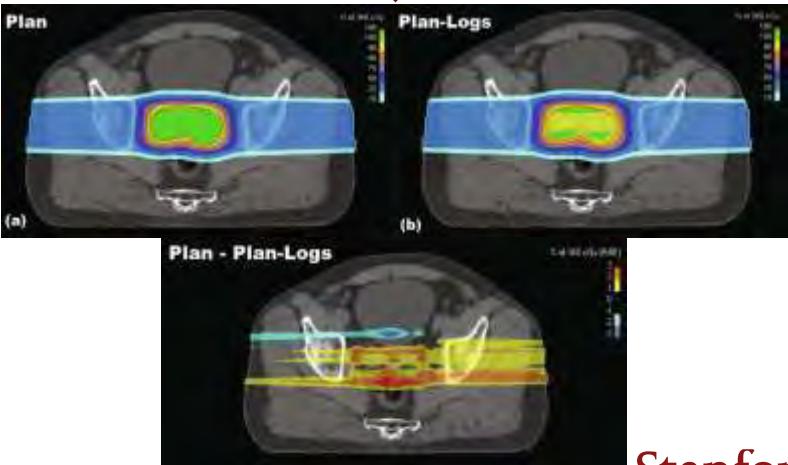


**Stanford
MEDICINE**

QA & DOSIMETRY

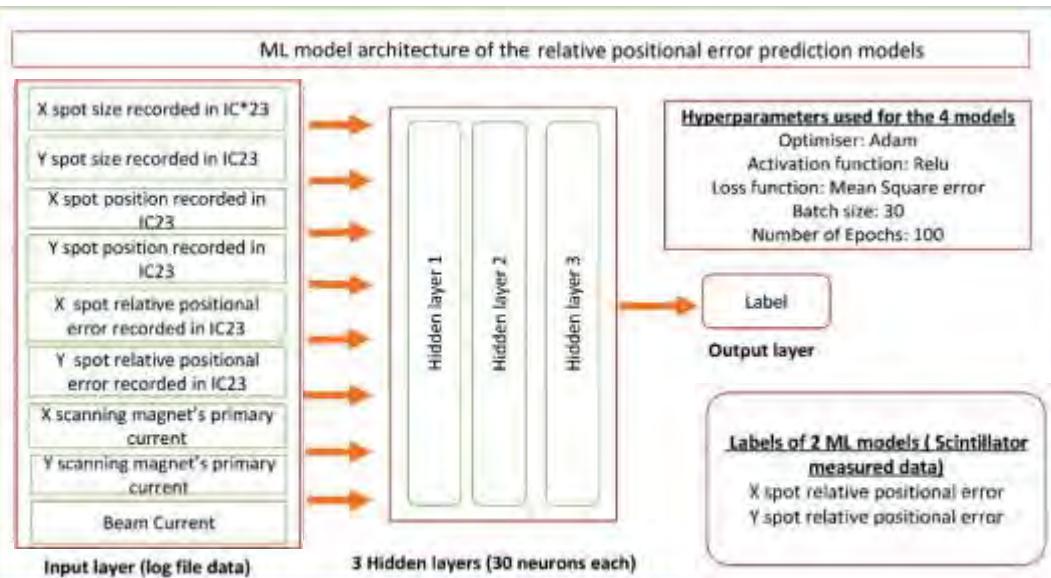


(a) (b)

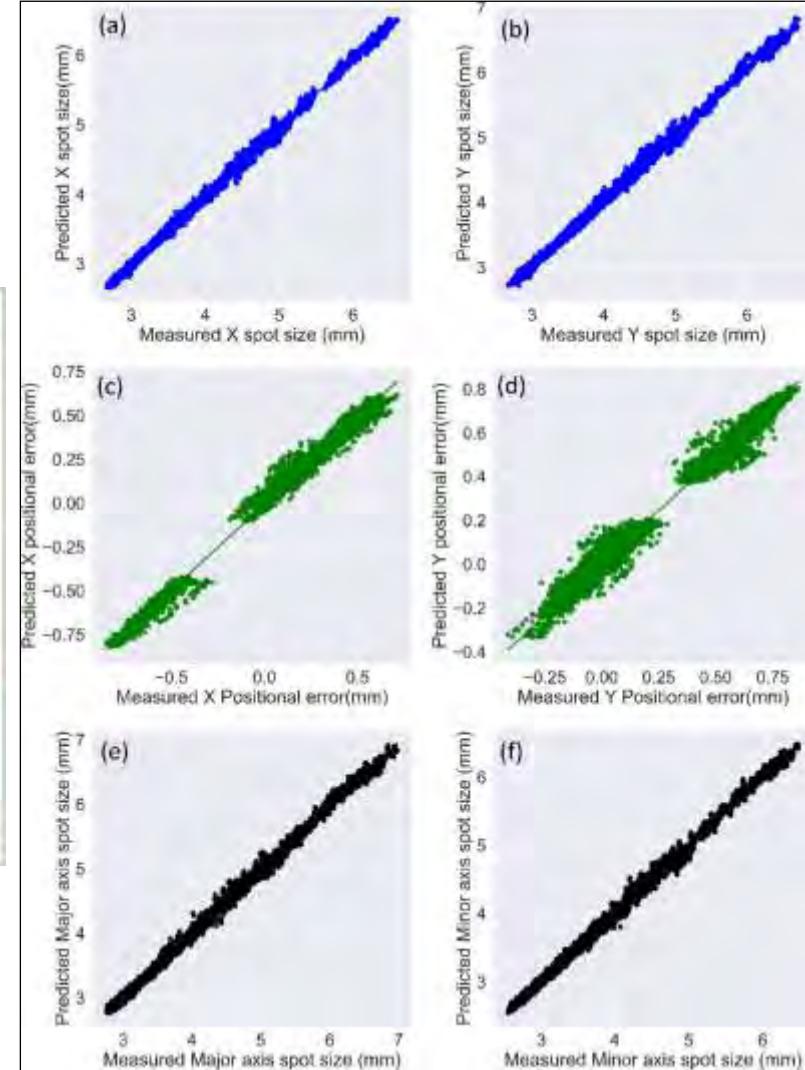


Maes et al., Physica Medica, 78, 2020

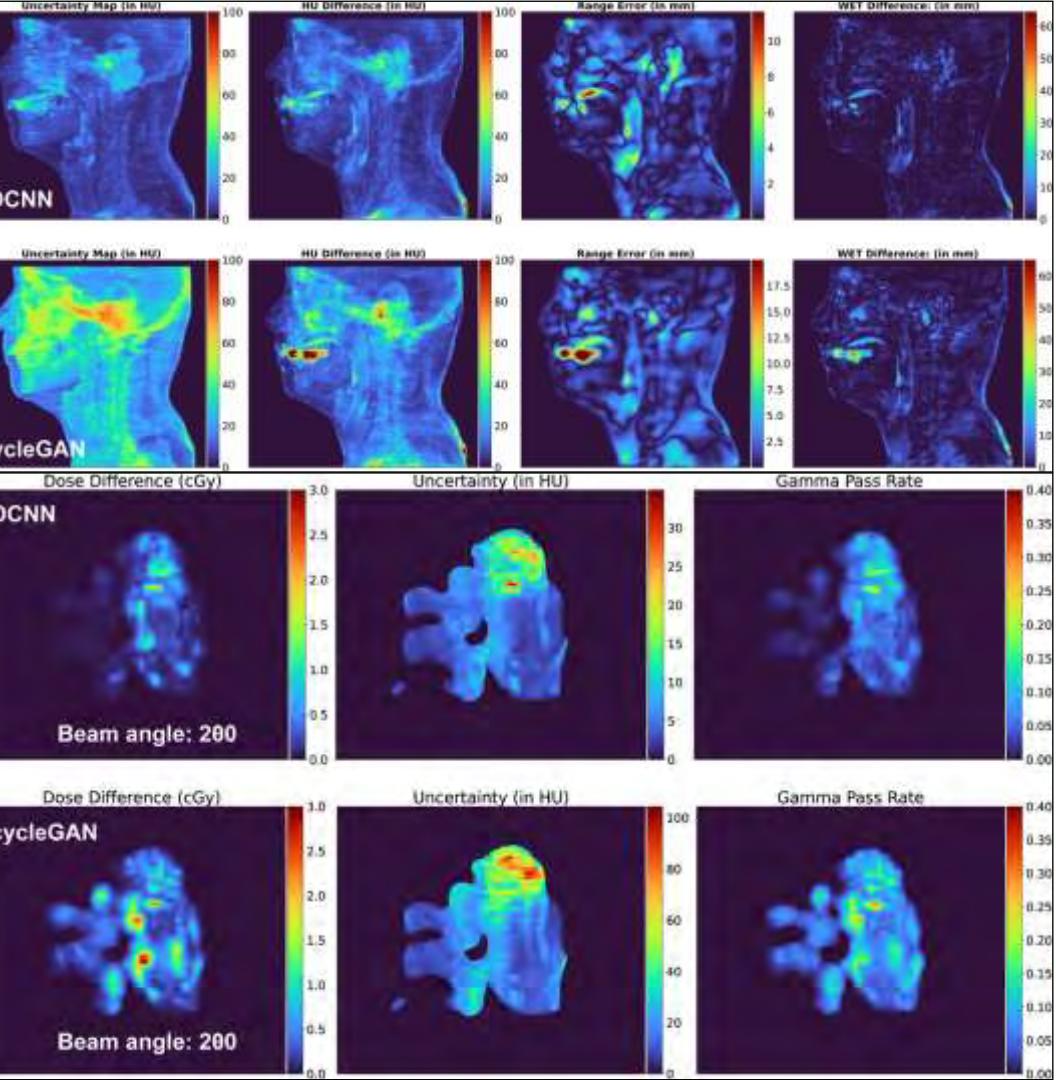
QA & DOSIMETRY



Ranjith et al., BPEX, 10, 2024



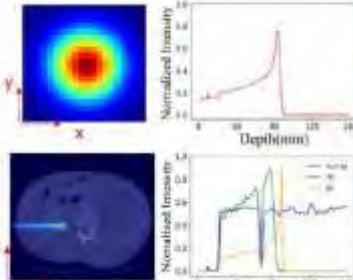
QA & DOSIMETRY



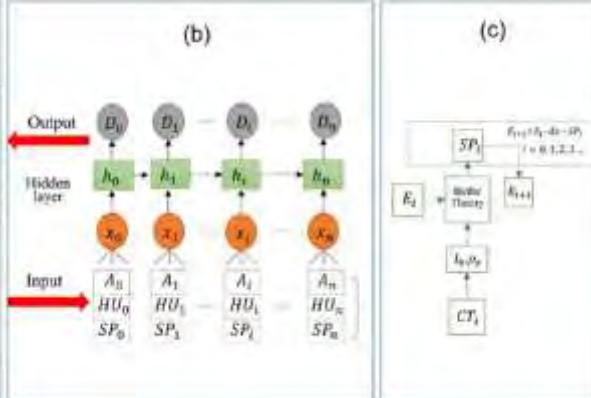
- [1] B. Sun et al., "A machine learning approach to the accurate prediction of monitor units for a compact proton machine", *Med Phys*, vol. 45, pp. 2243-2251, May 2018.
- [2] H. S. Grewal, M. S. Chacko, S. Ahmad and H. Jin, "Prediction of the output factor using machine and deep learning approach in uniform scanning proton therapy", *J. Appl. Clin. Med. Phys.*, vol. 21, pp. 128-134, Jul. 2020.P
- [3] D. Maes et al., "A machine learning-based framework for delivery error prediction in proton pencil beam scanning using irradiation log-files", *Physica Medica*, vol. 78, pp. 179-186, Oct. 2020.
- [4] Fleury E, Herault J, Spruijt K, Kouwenberg J, Angellier G, Hofverberg P, Horwacik T, Kajdrowicz T, Pignol JP, Hoogeman M, Trnková P. A generalized model for monitor units determination in ocular proton therapy using machine learning: A proof-of-concept study. *Phys Med Biol.* 2024 Feb 12;69(4). doi: 10.1088/1361-6560/ad1d68. PMID: 38211314.
- [5] Ranjith CP, Krishnan M, Raveendran V, Chaudhari L, Laskar S. An artificial neural network based approach for predicting the proton beam spot dosimetric characteristics of a pencil beam scanning technique. *Biomed Phys Eng Express.* 2024 Apr 22;10(3). doi: 10.1088/2057-1976/ad3ce0. PMID: 38652667.
- [6] Galapon AV Jr, Thummerer A, Langendijk JA, Wagenaar D, Both S. Feasibility of Monte Carlo dropout-based uncertainty maps to evaluate deep learning-based synthetic CTs for adaptive proton therapy. *Med Phys.* 2024 Apr;51(4):2499-2509. doi: 10.1002/mp.16838. Epub 2023 Nov 13. PMID: 37956266.

RANGE & DOSE VERIFICATION

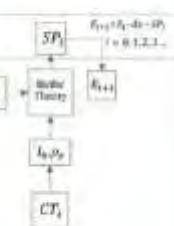
(a)



(b)

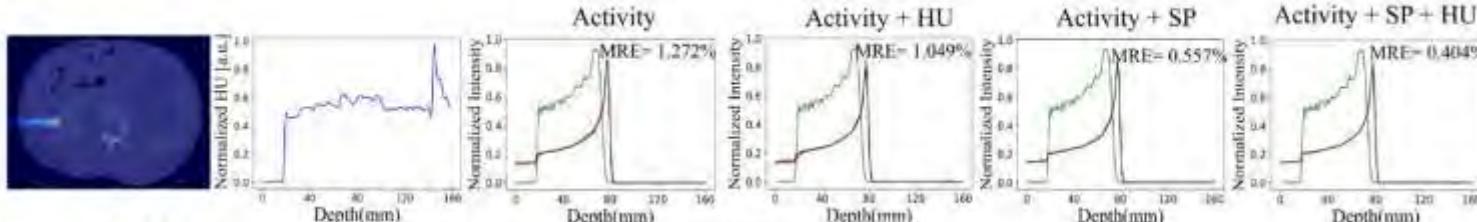


(c)

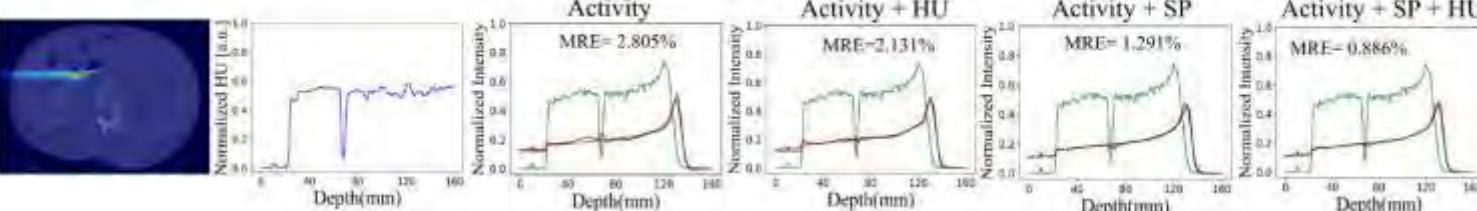


Hu et al., PMB, 65, 2020

(a) E = 90 MeV

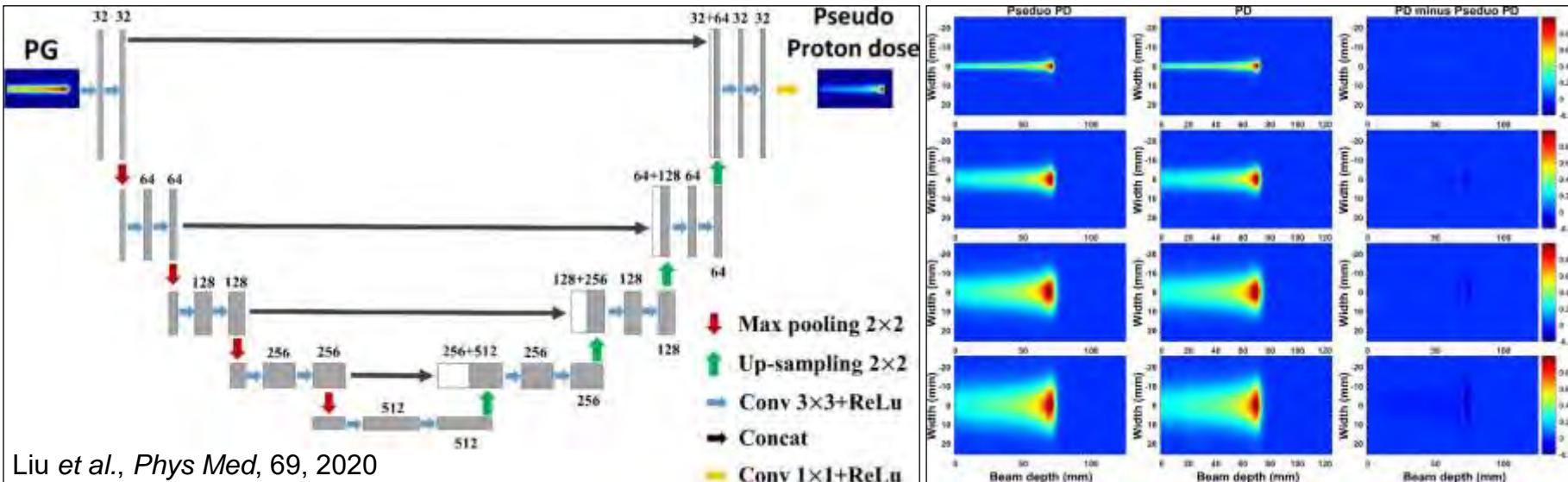


(b) E = 125 MeV

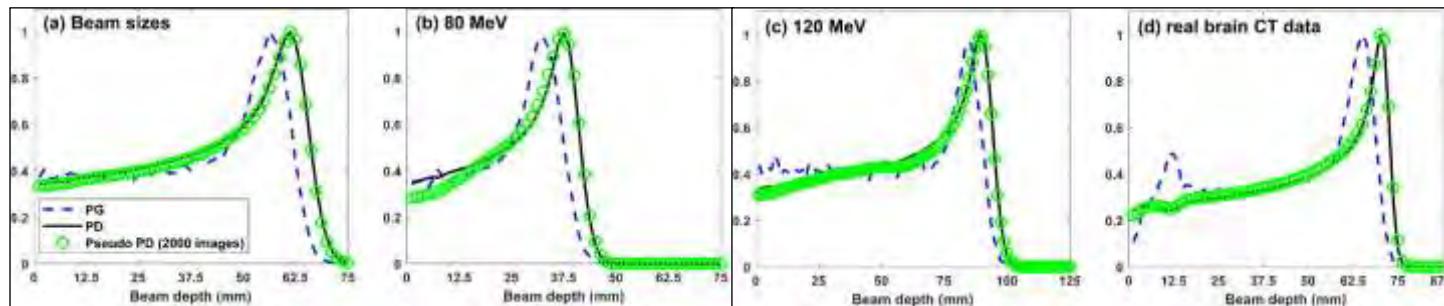


Stanford
MEDICINE

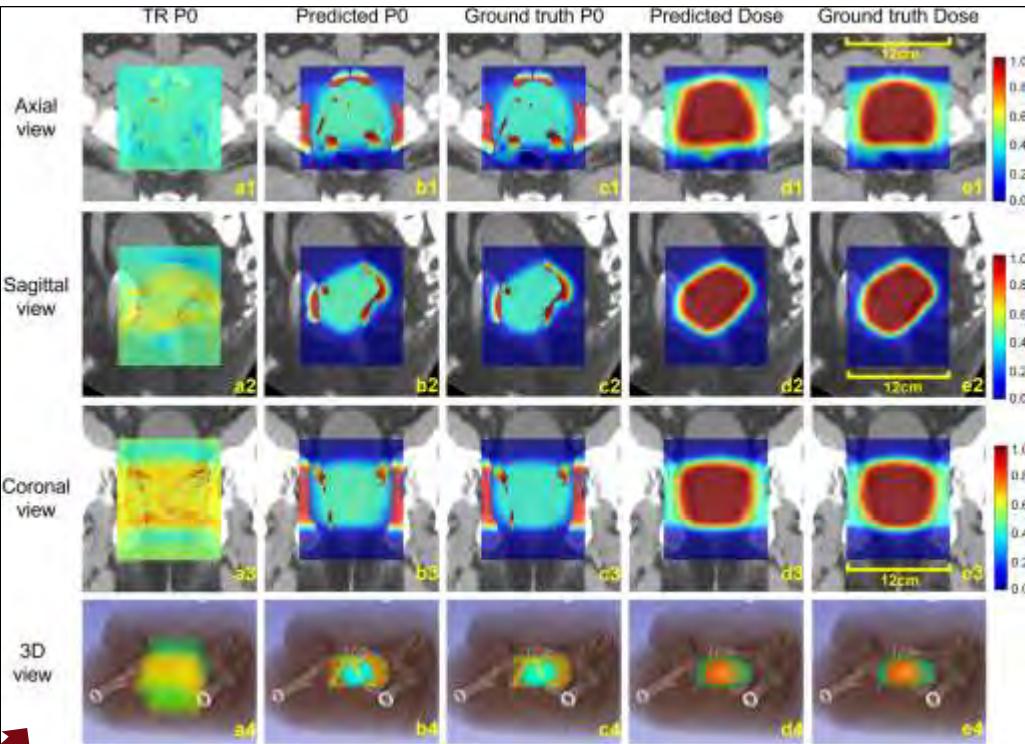
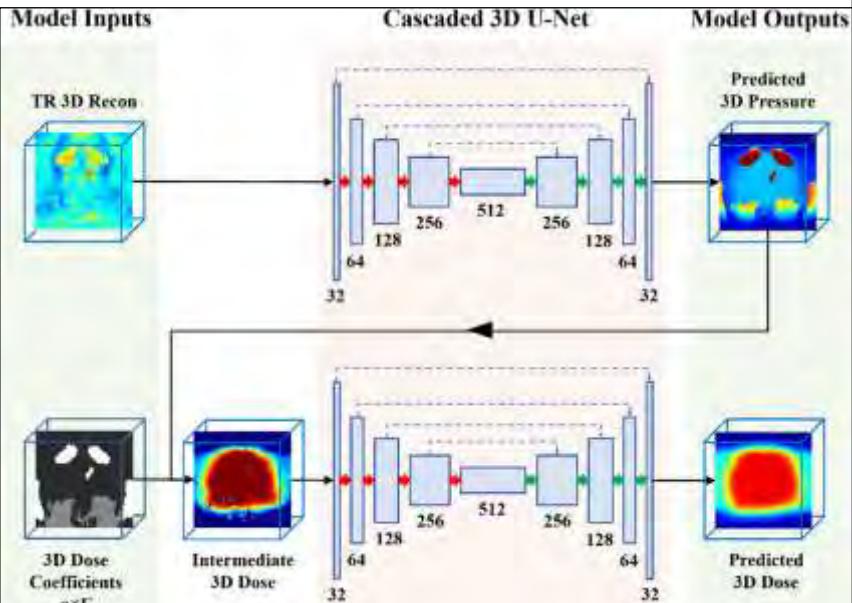
RANGE & DOSE VERIFICATION



Liu et al., Phys Med, 69, 2020



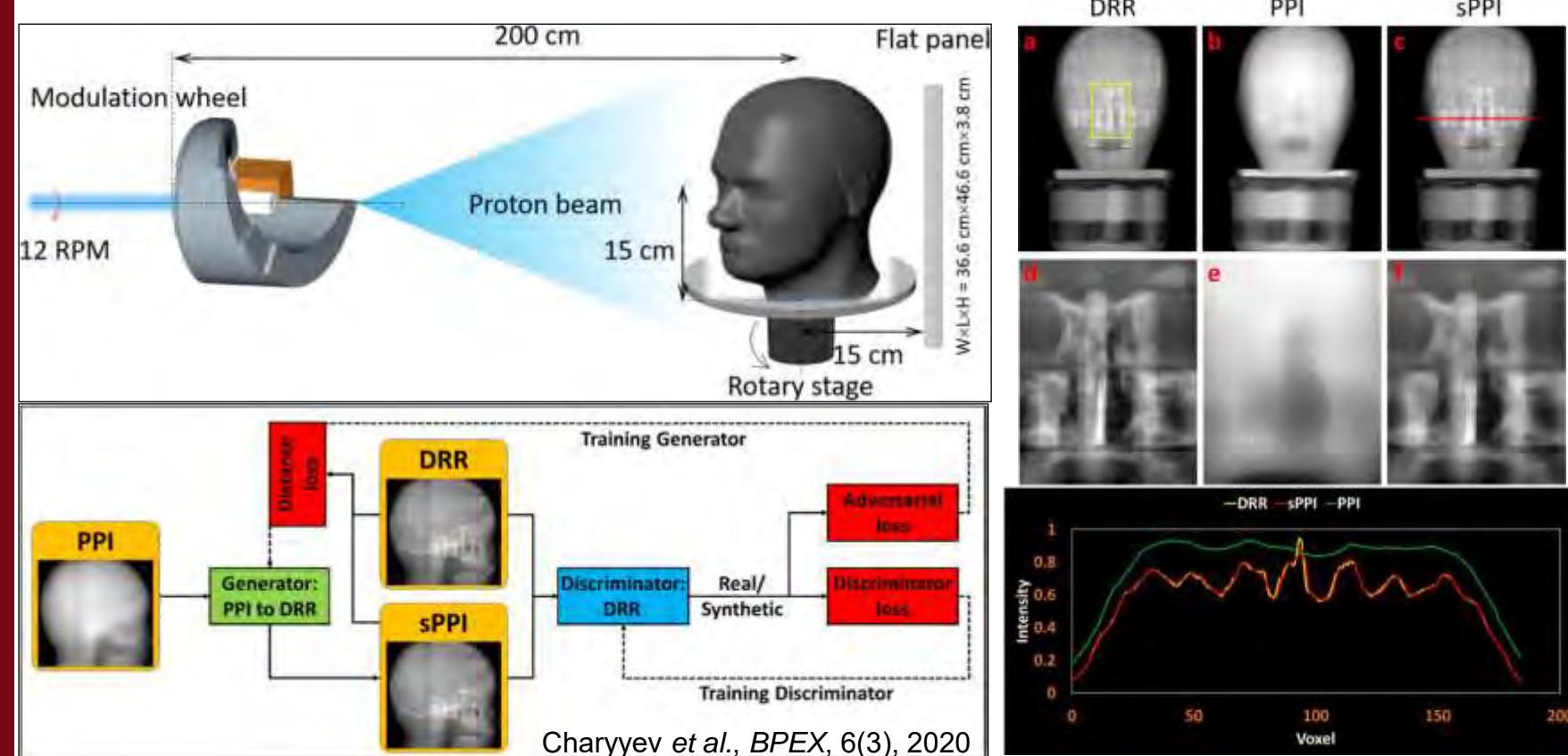
RANGE & DOSE VERIFICATION



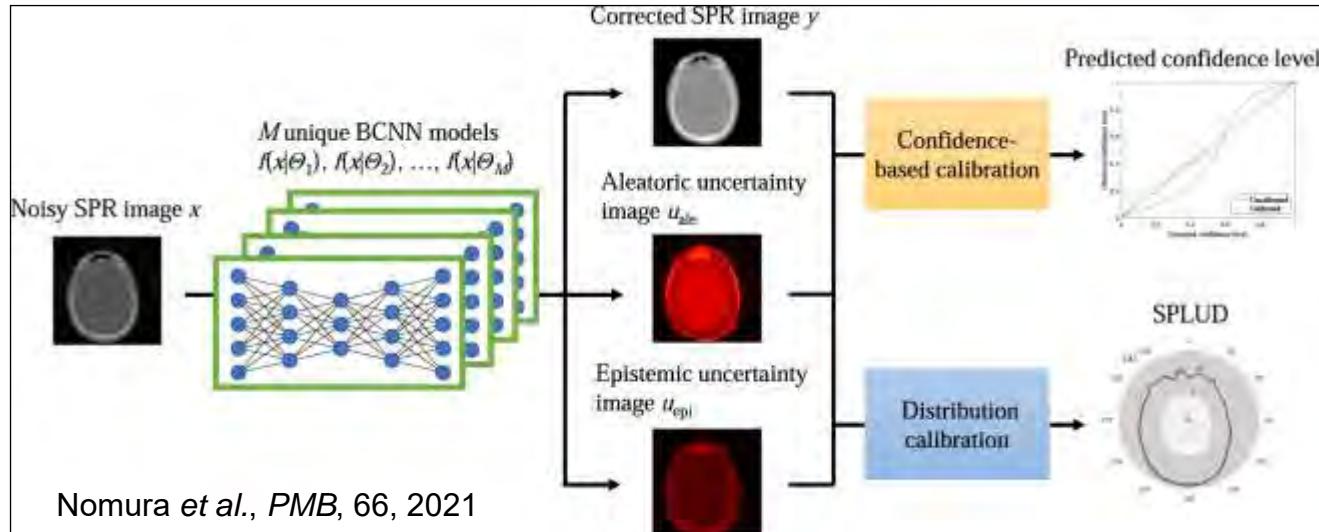
Jiang et al., PMB, 67, 2022

- [1] Z. Li, Y. Wang, Y. Yu, K. Fan, L. Xing and H. Peng, "Machine learning approaches for range and dose verification in proton therapy using proton-induced positron emitters", Med. Phys., vol. 46, no. 12, pp. 5748-5757, 2019.
- [2] C. Liu, Z. Li, W. Hu, L. Xing and H. Peng, "Range and dose verification in proton therapy using proton-induced positron emitters and recurrent neural networks (RNNs)", Phys. Med. Biol., vol. 64, no. 17, 2019.
- [3] Z. Hu, G. Li, X. Zhang, K. Ye, J. Lu and H. Peng, "A machine learning framework with anatomical prior for Online dose verification using positron emitters and PET in proton therapy", Phys. Med. Biol., vol. 65, no. 18, 2020.
- [4] S. Ma, Z. Hu, K. Ye, X. Zhang, Y. Wang and H. Peng, "Feasibility study of patient-specific dose verification in proton therapy utilizing positron emission tomography (PET) and generative adversarial network (GAN)", Med. Phys., vol. 47, no. 10, pp. 5194-5208, 2020.
- [5] P. Gueth et al., "Machine learning-based patient specific prompt-gamma dose monitoring in proton therapy", Phys. Med. Biol., vol. 58, no. 13, pp. 4563-4577, 2013.
- [6] C.-C. Liu and H.-M. Huang, "A deep learning approach for converting prompt gamma images to proton dose distributions: A Monte Carlo simulation study", Phys. Med., vol. 69, pp. 110-119, Jan. 2020.
- [7] S. Yao, Z. Hu, Q. Xie, Y. Yang and H. Peng, "Further investigation of 3D dose verification in proton therapy utilizing acoustic signal wavelet decomposition and machine learning", Biomed. Phys. Eng. Exp., vol. 8, Nov. 2021.
- [8] S. Yao et al., "Feasibility study of range verification based on proton-induced acoustic signals and recurrent neural network", Phys. Med. Biol., vol. 65, Nov. 2020.
- [9] Z. Jiang et al., "3-D in-vivo dose verification in prostate proton therapy with deep learning-based proton-acoustic imaging", Phys. Med. Biol., vol. 67, no. 21, 2022.
- [10] Lang Y, Jiang Z, Sun L, Tran P, Mossahebi S, Xiang L, Ren L. Patient-specific deep learning for 3D protoacoustic image reconstruction and dose verification in proton therapy. Med Phys. 2024 Jul 9. doi: 10.1002/mp.17294. Epub ahead of print. PMID: 38980065.
- [11] Tian, Liheng, and Armin Lühr. "Proton range uncertainty caused by synthetic computed tomography generated with deep learning from pelvic magnetic resonance imaging." Acta Oncologica 62.11 (2023): 1461-1469.

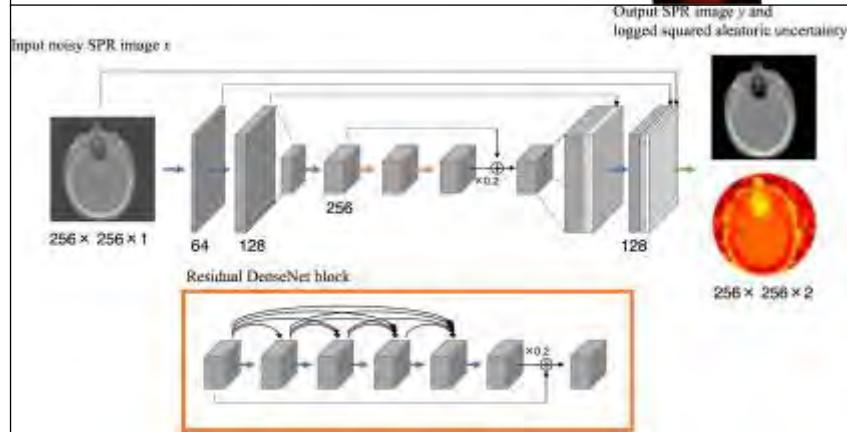
RADIOGRAPHY & TOMOGRAPHY



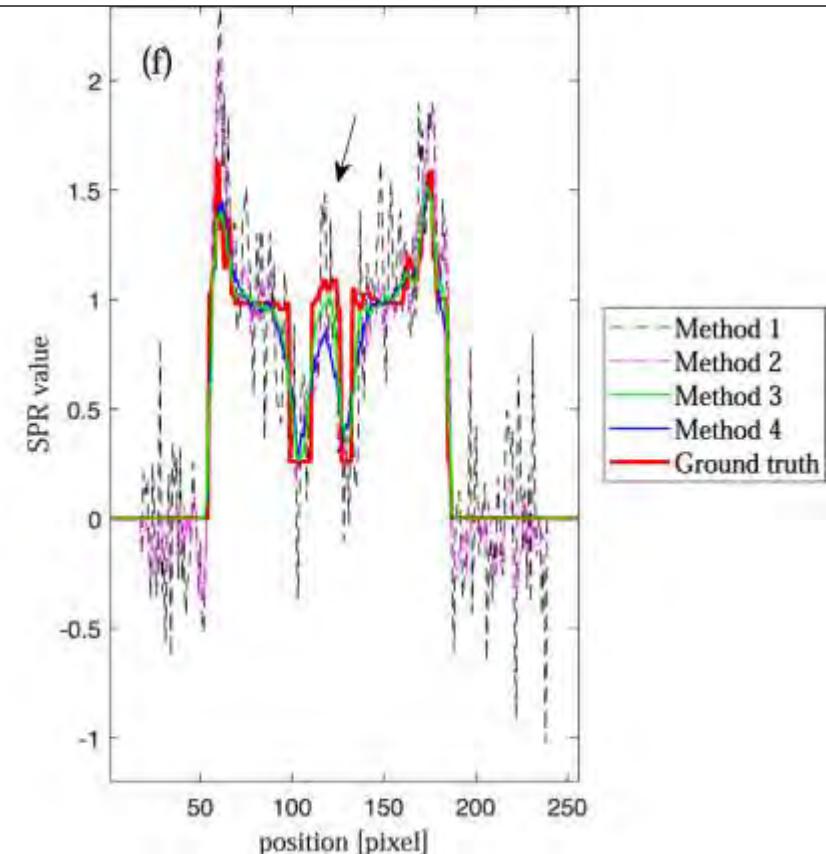
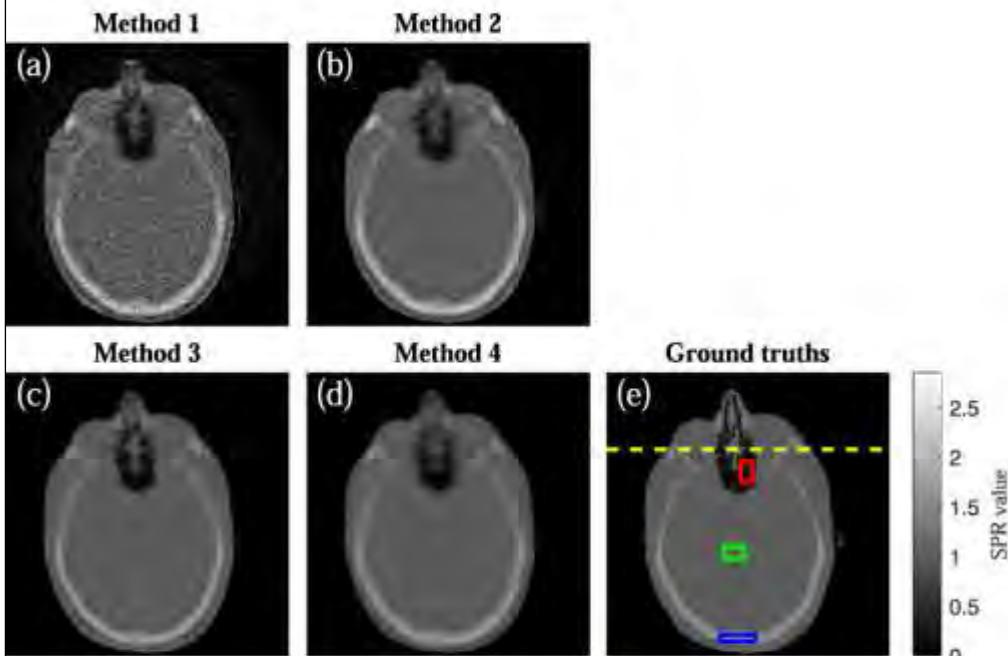
RADIOGRAPHY & TOMOGRAPHY



Nomura et al., PMB, 66, 2021



RADIOGRAPHY & TOMOGRAPHY



- [1] Sheng, Cong, et al. "A denoising method based on deep learning for proton radiograph using energy resolved dose function." *Physics in Medicine & Biology* 69.2 (2024): 025015.
- [2] Alaka, B. G., et al. "A comparative study of machine-learning approaches in proton radiography using energy-resolved dose function." *Physica Medica* 106 (2023): 102525.
- [3] Ackernley T, Casse G, Cristoforetti M. Proton path reconstruction for proton computed tomography using neural networks. *Phys Med Biol.* 2021 Apr 6;66(7). doi: 10.1088/1361-6560/abf00f. PMID: 33735852.
- [4] Nomura Y, Tanaka S, Wang J, Shirato H, Shimizu S, Xing L. Calibrated uncertainty estimation for interpretable proton computed tomography image correction using Bayesian deep learning. *Phys Med Biol.* 2021 Mar 16;66(6):065029. doi: 10.1088/1361-6560/abe956. PMID: 33626513.
- [5] van der Heyden B, Cohilis M, Souris K, de Freitas Nascimento L, Sterpin E. Artificial intelligence supported single detector multi-energy proton radiography system. *Phys Med Biol.* 2021 May 4;66(10). doi: 10.1088/1361-6560/abe918. PMID: 33621962.
- [6] Charyev S, Lei Y, Harms J, Eaton B, McDonald M, Curran WJ, Liu T, Zhou J, Zhang R, Yang X. High quality proton portal imaging using deep learning for proton radiation therapy: a phantom study. *Biomed Phys Eng Express.* 2020 Apr 27;6(3):035029. doi: 10.1088/2057-1976/ab8a74. PMID: 33438674.

CONCLUSIONS

1

INTRODUCTION

2

AI in PLANNING

4

AI in DELIVERY

5

CONCLUSIONS



Stanford
MEDICINE

- We are in a golden age of progress in the field of AI.
- Proton therapy is perfectly suited to benefit from AI to enhance accuracy and efficiency.
- Specifically, AI can benefit during treatment planning; dose calculation; quality assurance; image guidance; and delivery.
- There are challenges and pitfalls.



Stanford
MEDICINE