

# Evaluating the Clinical Utility of Artificial Intelligence Assistance and its Explanation on the Glioma Grading Task

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## **Outline** A phase 2 clinical user study of AI on glioma grading

- 1. Study motivation**
- 2. Study design:** material, participants, procedure
- 3. Result:** doctor+AI > doctor?

### **Disclosure statement**

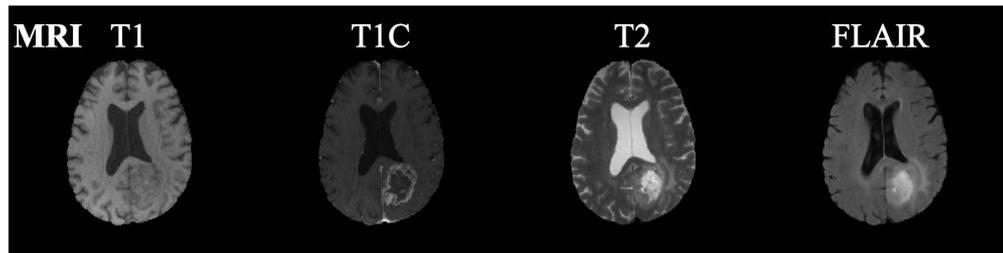
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All authors declare no financial or non-financial competing interests.

## Motivation

**Glioma:** most common primary tumor of brain and spine

Initial investigation includes CT or MRI for diagnosis, tumor grading, treatment planning, progression tracking, prognosis, etc.



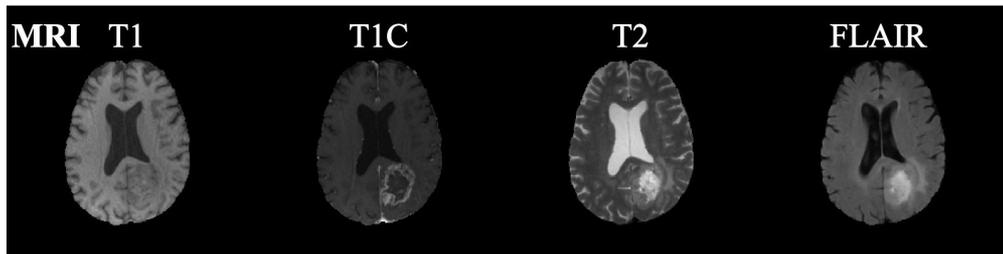
[1] Artificial intelligence in glioma imaging: challenges and advances. J Neural Eng. 2020

[2] Evaluating artificial intelligence in medicine: phases of clinical research. JAMIA Open. 2020.

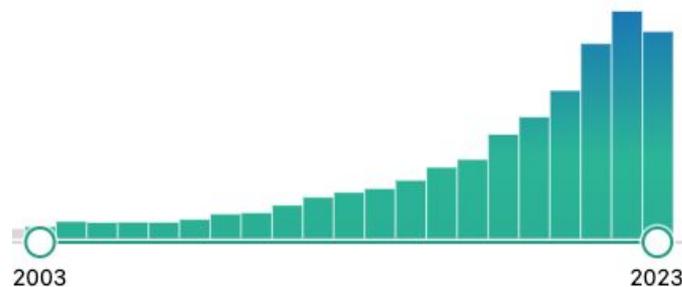
## Motivation

**Glioma:** most common primary tumor of brain and spine

Initial investigation includes CT or MRI for diagnosis, tumor grading, treatment planning, progression tracking, prognosis, etc.



Artificial intelligence (AI) may be a potential tool to assist doctors in glioma imaging tasks, as it is trained to recognize patterns from a large amount of data



PubMed keywords: glioma + AI  
[https://bit.ly/AI\\_in\\_glioma\\_imaging](https://bit.ly/AI_in_glioma_imaging) [1]

[1] Artificial intelligence in glioma imaging: challenges and advances. J Neural Eng. 2020

[2] Evaluating artificial intelligence in medicine: phases of clinical research. JAMIA Open. 2020.

# Motivation

To safeguard the use of AI in clinical setting:

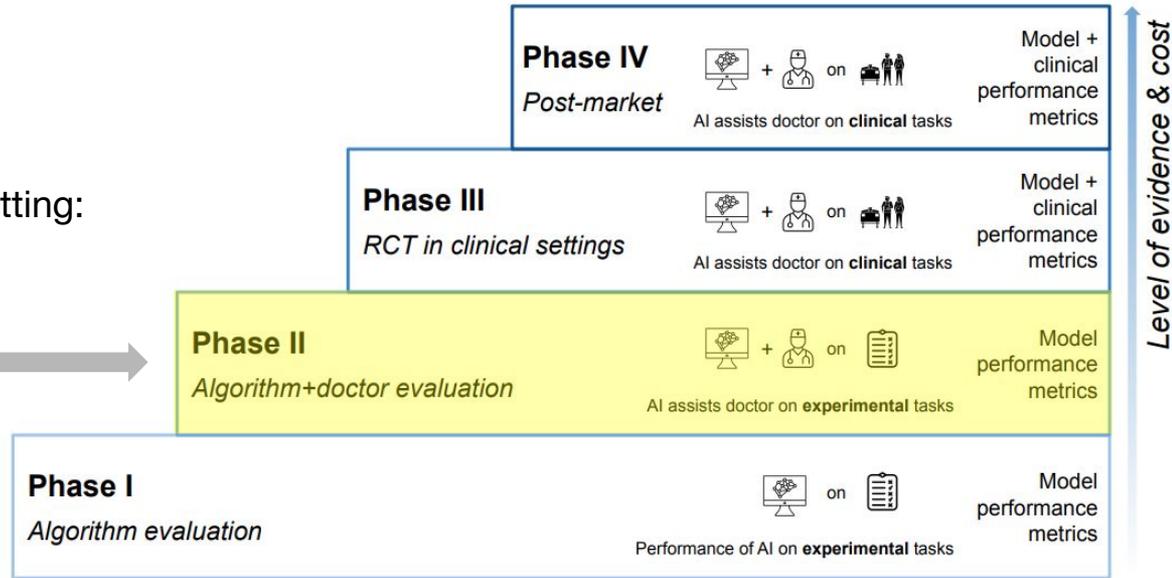
## 1. Clinical evaluation

Phase 2 clinical study



## 2. eXplainable AI (XAI)

Being able to explain decisions to clinical users



The four phases of evaluating the clinical utility of AI in glioma imaging [1].

[1] Artificial intelligence in glioma imaging: challenges and advances. J Neural Eng. 2020

[2] Evaluating artificial intelligence in medicine: phases of clinical research. JAMIA Open. 2020.

## Research question

**Doctor + AI > Doctor ?**

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If so,

**Doctor + AI > max(Doctor, AI) ?**



Complementary  
doctor-AI  
performance

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Suggestion Explanation

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Complementary  
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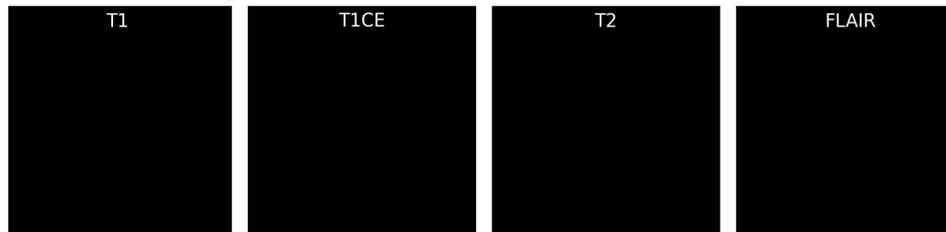
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# Glioma grading on MRI Data

BraTS 20'  
Dataset <sup>1</sup>  
n=369

## Grade 2-3 glioma

n=76



— Tumor mask  
contour

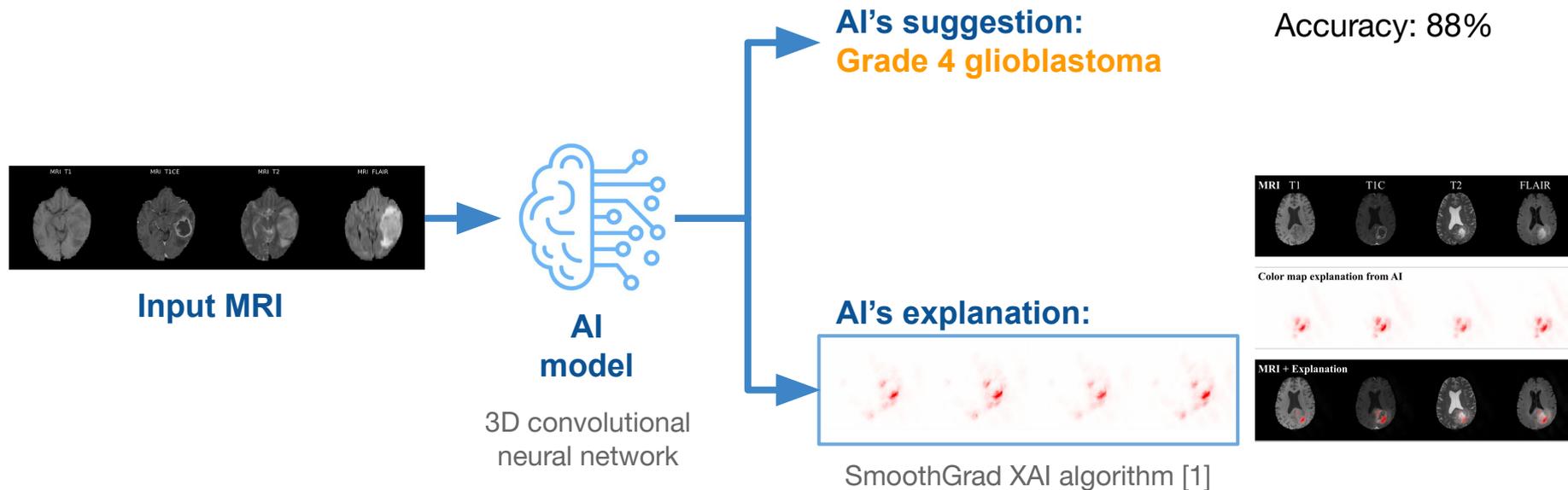
## Grade 4 glioblastoma

n=293



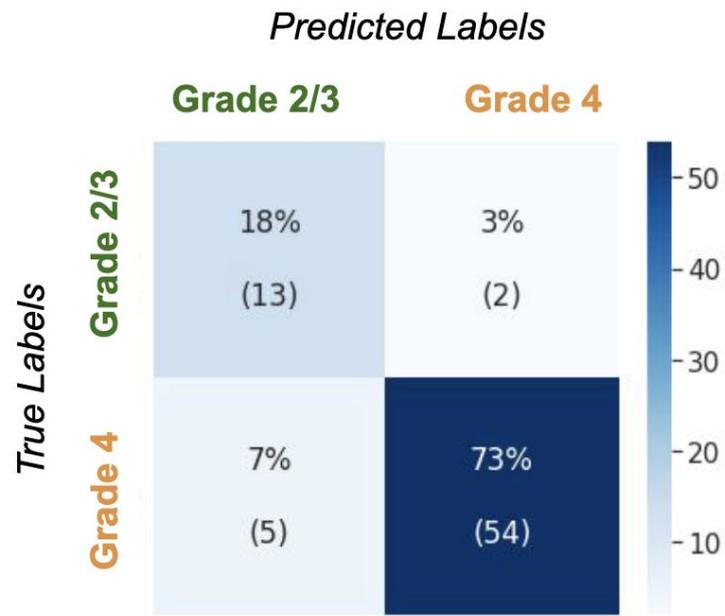
[1] The Multimodal Brain Tumor Image Segmentation Benchmark (BRATS). Menze, et al., IEEE TMI 2015.

# Glioma grading on MRI AI assistance: a suggestion + an explanation



[1] Smoothgrad: removing noise by adding noise, 2017. Arxiv: 1706.03825

# Glioma grading on MRI AI model performance



	Sensitivity	Specificity	Positive Predictive Value	F1 score	Number of class in the test set
Grade 2 or 3 glioma	0.87	0.92	0.72	0.79	15
Grade 4 glioblastoma	0.92	0.87	0.96	0.94	59

# Candidates of 16 post-hoc heatmap explanation methods on the glioma task

## Gradient based

Grad-CAM



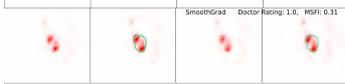
Gradient



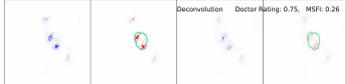
Input x Gradient



SmoothGrad



Deconvolution



Guided Backpropagation



Guided Grad-CAM



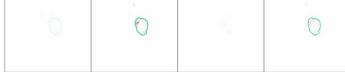
Integrated Gradient



DeepLIFT



Gradient SHAP



## Perturbation based

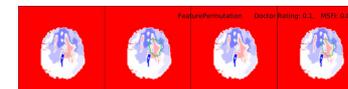
Occlusion



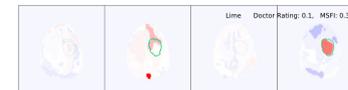
Feature Ablation



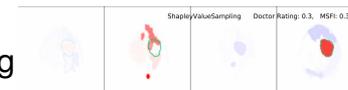
Feature Permutation



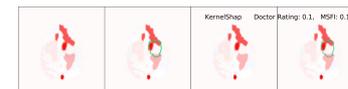
LIME



Shapley Value Sampling

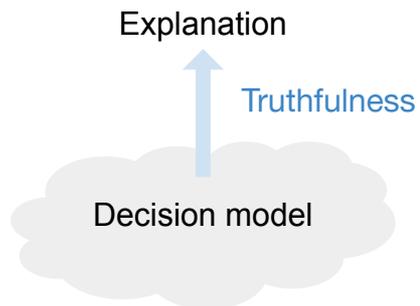


Kernel SHAP



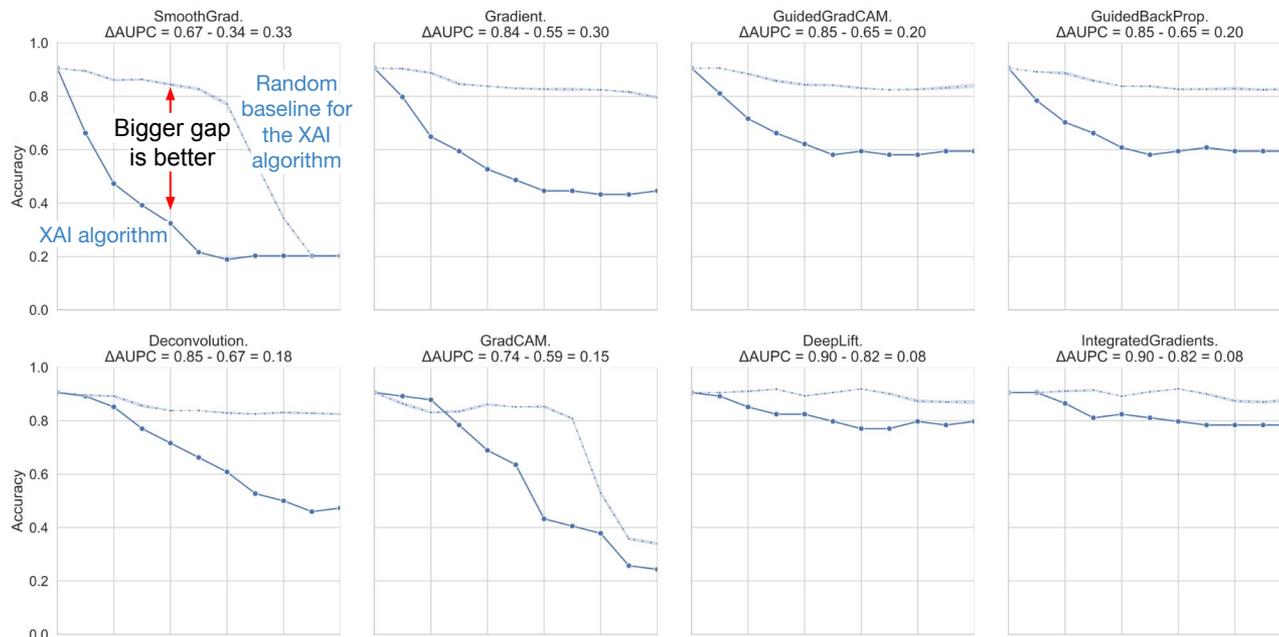
# The AI explanation method was selected for being relatively truthful to AI decision-making

## Gradual feature removal experiment



### Assumption:

Removing important features will cause AI performance to drop.



.....

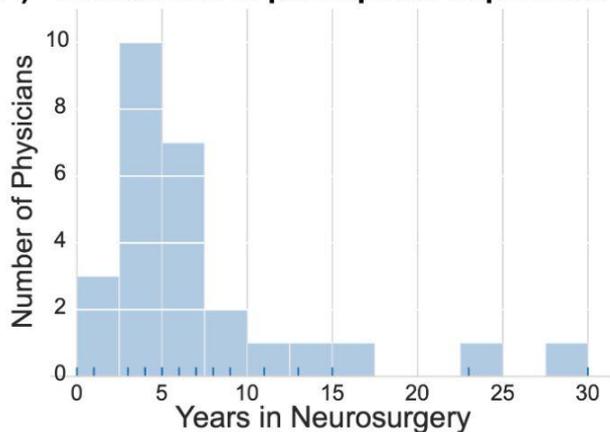
# Participants

- **Inclusion criteria**
  - Neurosurgeon, radiologist, or neuro-radiologist;
  - Attending, fellow, or resident
- **Eligibility screening** by:
  - survey screening questions, and
  - **DR** task accuracy > 0.55

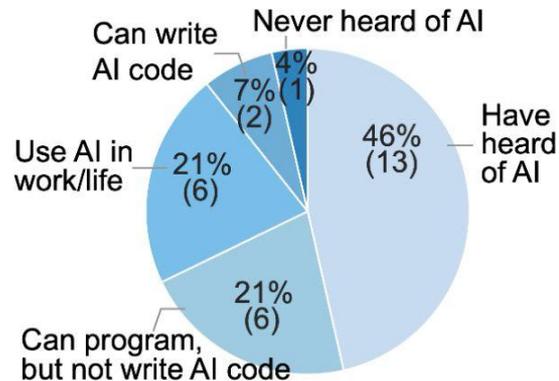
# Participants

- 35 participants (recruitment rate 15%) in neurosurgery
- 12 attending, 2 fellows, 21 residents
- Years of practicing neurosurgery:  $7.1 \pm 6.5$
- Female: male = 7:19; age:  $34.7 \pm 8.2$

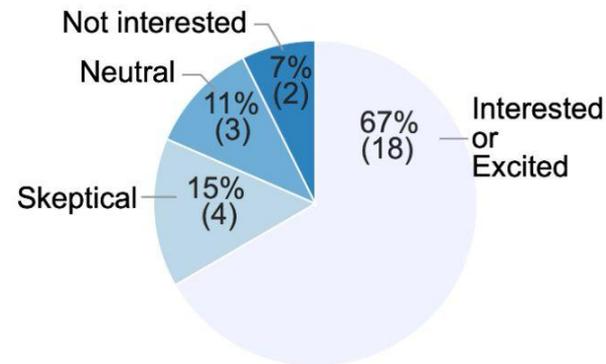
**(A) Distribution of participants' experience**



**(B) Familiarity with AI**



**(C) Attitude toward AI**



## Study design

- National online **survey**, 25 MRI, 30-40 min.

Participants gave judgment at three conditions:



Grade 4  
tumor



Grade 4  
tumor



1. DR

2. DR+AI

3. DR+XAI

Doctor only

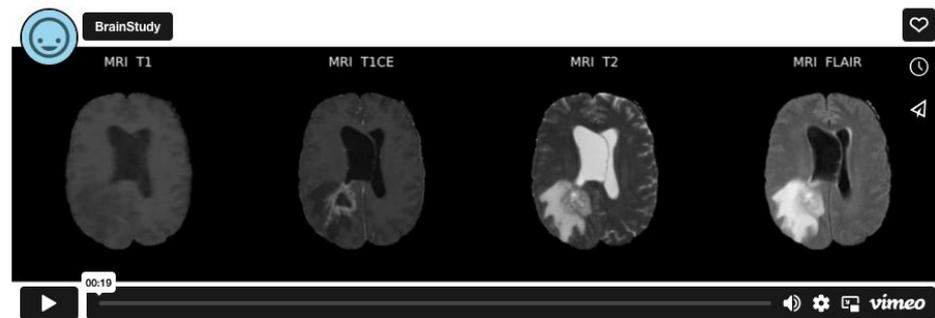
Doctor assisted  
by AI **prediction**

Doctor assisted  
by AI prediction &  
**explanation**

Next, you will use this AI to assist you on tumor grading for 25 new patients' MRIs.

Each 3D MR image is presented as a video. You can click the triangle button to play the video and read the MRI.

\* 8.



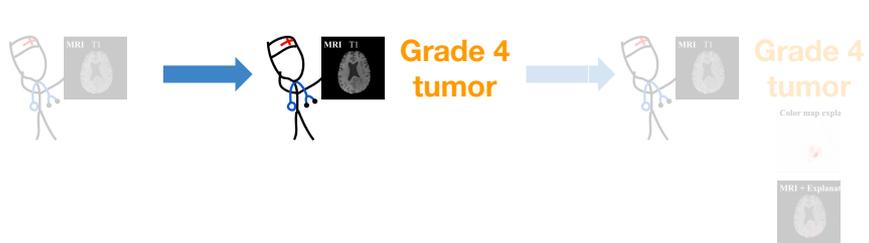
What grade of glioma would you predict this MRI to be?

Grade 2 or Grade 3 glioma

Grade 4 glioblastoma

## Study design

- National online **survey**, 25 MRI, 30-40 min.  
Participants gave judgment at three conditions:



1. DR

Doctor only

2. DR+AI

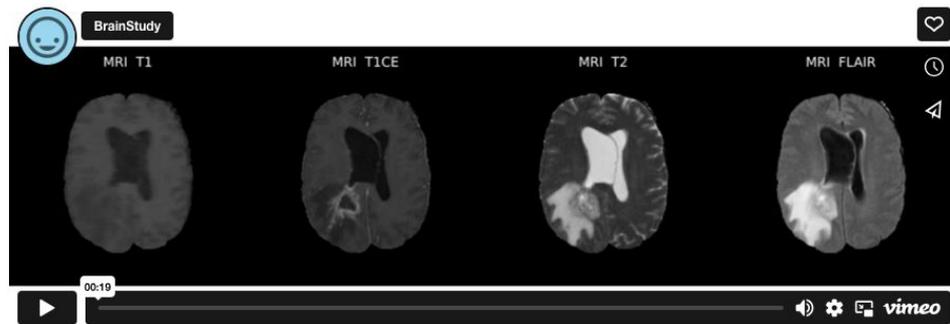
Doctor assisted  
by AI **prediction**

3. DR+XAI

Doctor assisted  
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\* 9. Your prediction is **Grade 4 glioblastoma**. AI's prediction is **Grade 4 glioblastoma**.

After viewing AI's suggestion, what is your current judgment on the tumor grade?

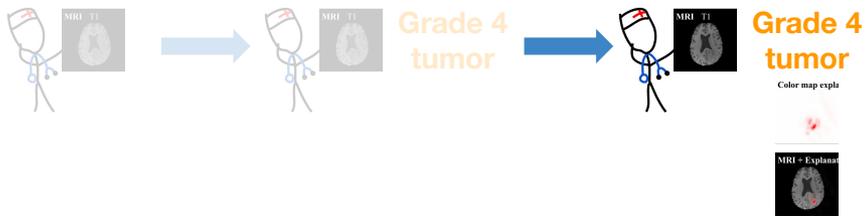


Grade 2/3 glioma

Grade 4 glioblastoma

## Study design

- National online **survey**, 25 MRI, 30-40 min.  
Participants gave judgment at three conditions:



### 1. DR

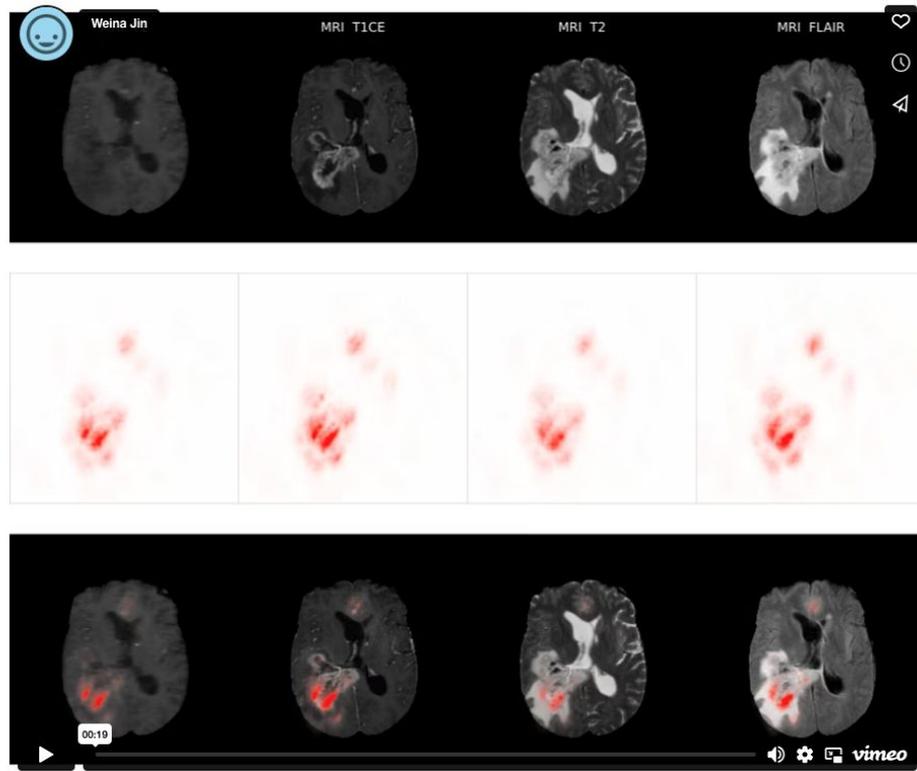
Doctor only

### 2. DR+AI

Doctor assisted  
by AI **prediction**

### 3. DR+XAI

Doctor assisted  
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**explanation**



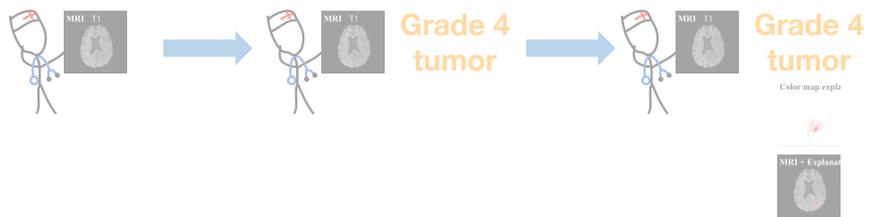
After viewing AI's explanation, what is your final judgment on the tumor grade?

Grade 2/3 glioma

Grade 4 glioblastoma

## Study design

- National online **survey**, 25 MRI, 30-40 min.  
Participants gave judgment at three conditions:



### 1. DR

Doctor only

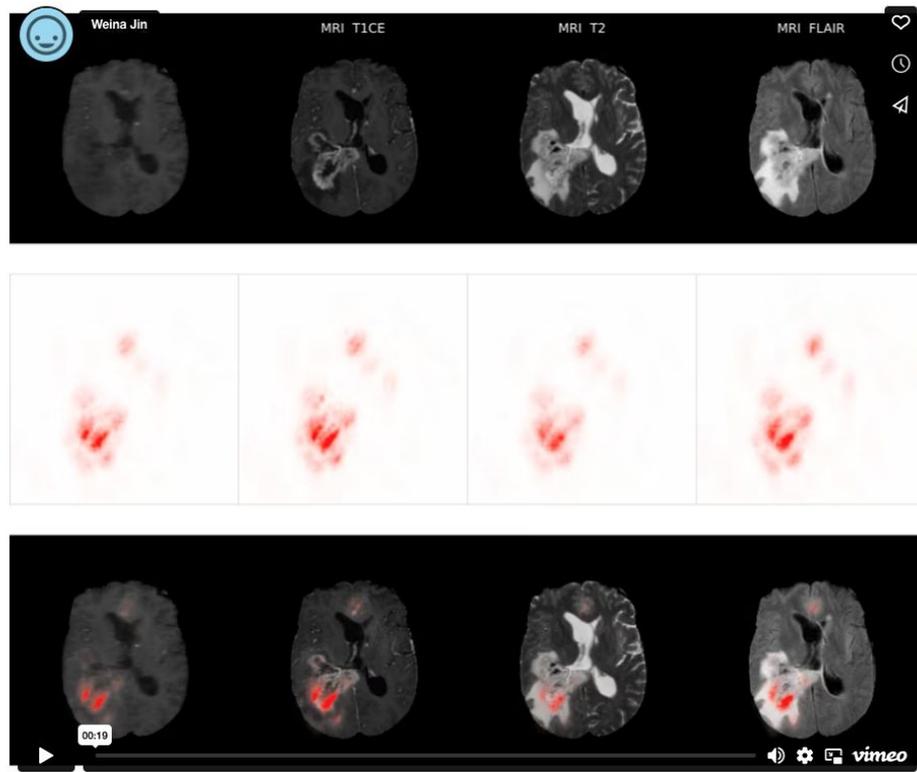
### 2. DR+AI

Doctor assisted  
by AI **prediction**

### 3. DR+XAI

Doctor assisted  
by AI prediction &  
**explanation**

- Post-survey, one-to-one **interview**, 20-30 min



After viewing AI's explanation, what is your final judgment on the tumor grade?

Grade 2/3 glioma

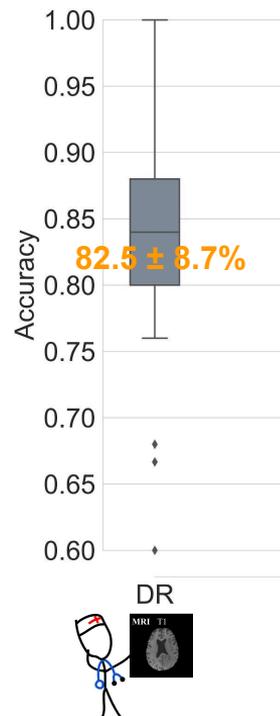
Grade 4 glioblastoma

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## Result

Is doctor+AI better than doctor only?



# Result

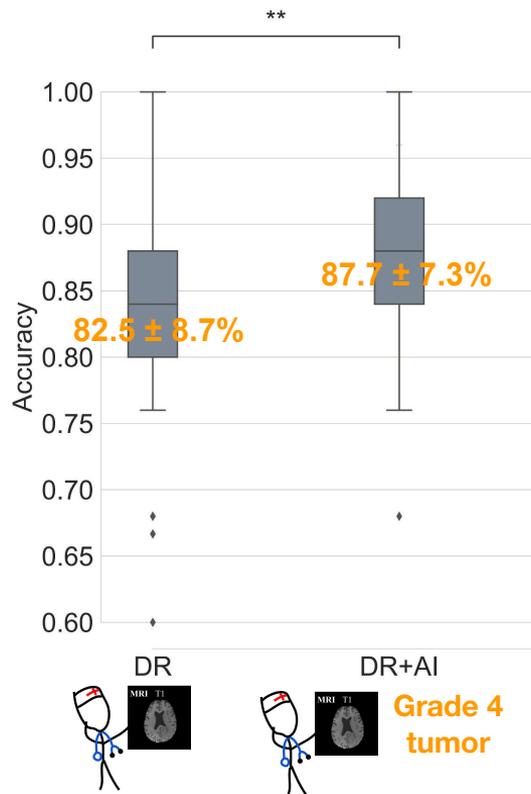
Is doctor+AI better than doctor only?

**Doctor + AI > Doctor** ✓

On improving doctors' task performance:

- AI prediction (**DR+AI**) is helpful

Wilcoxon signed-rank tests with Bonferroni correction



# Result

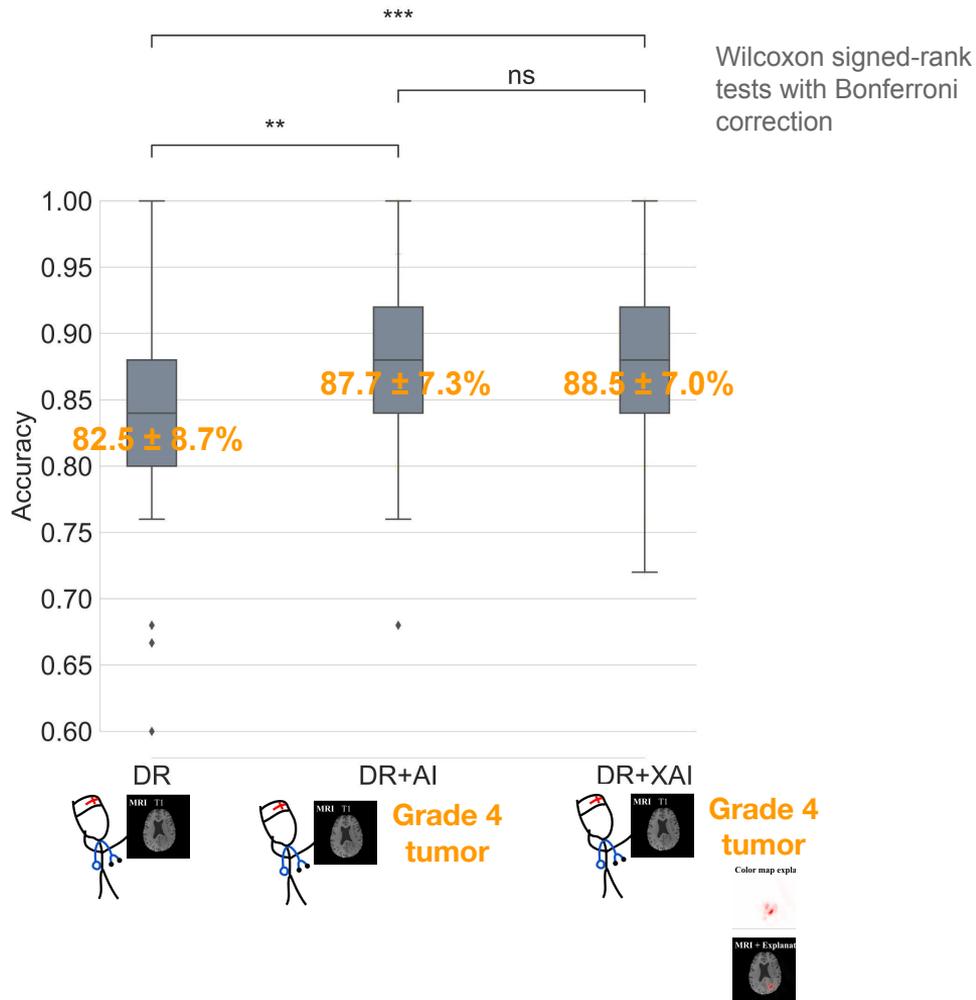
Is doctor+AI better than doctor only?

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Suggestion      Explanation

On improving doctors' task performance:

- AI prediction (**DR+AI**) is helpful
- AI explanation (**DR+XAI**) not show additional value



# Result

Is doctor+AI better than doctor only?

**Doctor + AI > Doctor** ✓

Suggestion Explanation

On improving doctors' task performance:

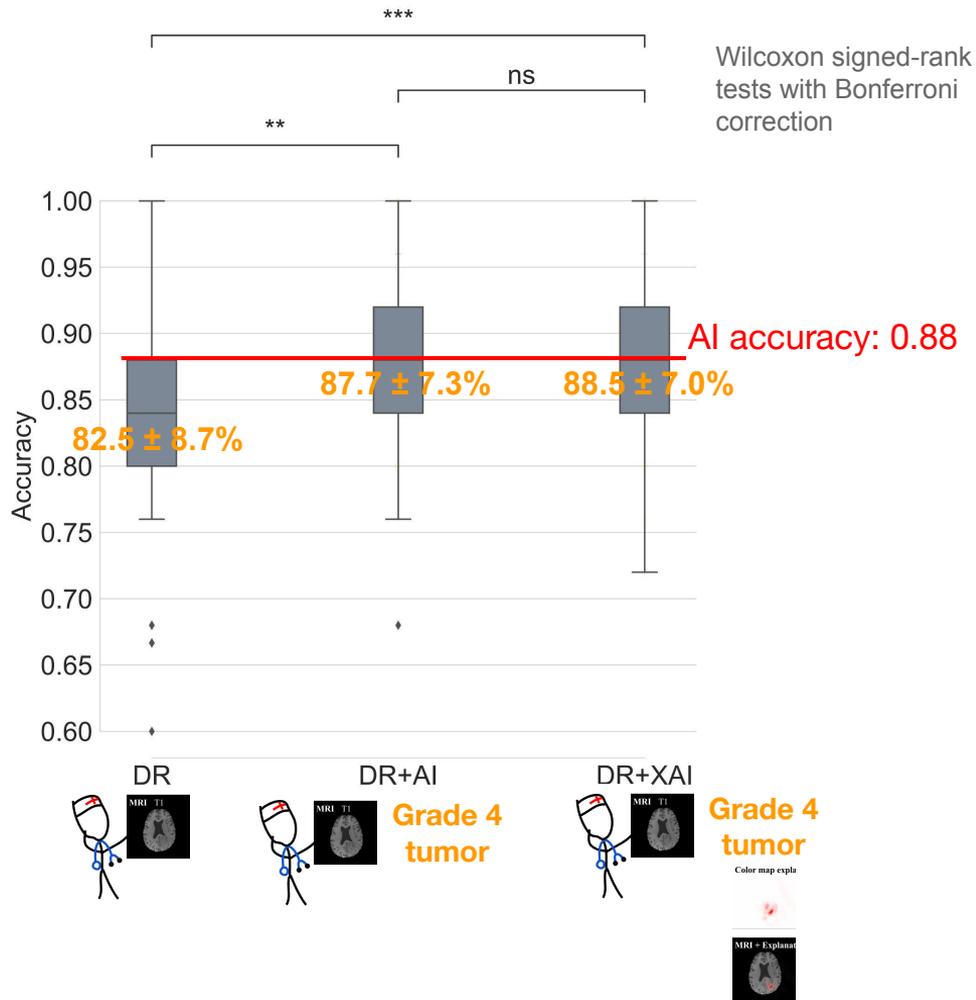
- AI prediction (**DR+AI**) is helpful
- AI explanation (**DR+XAI**) not show additional value

If so,

**Doctor + AI > max(Doctor, AI)** ✗

Suggestion Explanation

Not achieve complementary doctor-AI performance in **DR+AI** or **DR+XAI**

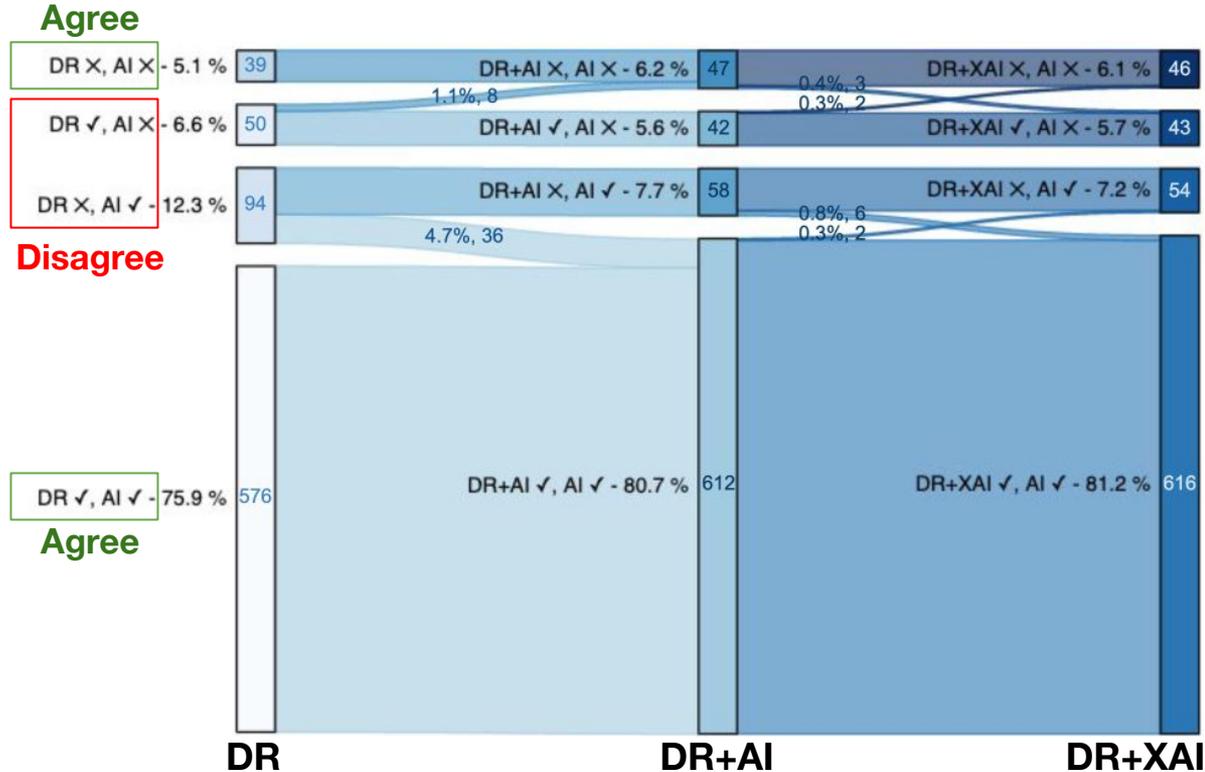




# Result Why did not AI explanation help?

Because physicians had both **changed decisions correctly and incorrectly** with basically the same amount.

I.e.: explanation cannot help doctors to discern potentially questionable decisions of AI.



““

**What does that (color map region) mean?** Like hey, which part of my car gets my car moving? It should say press the accelerator. But yours would just show a dashboard of the car, and show that this button had some red, that button had some red, but it's not an explanation. Let's go to an example, and you'll see, what about the red areas under MRI T1CE (modality)? **Was it central necrosis?** But it couldn't be the central necrosis, because there's more central necrosis in the temporal lobe, and that area didn't get highlighted. So anyway, I don't know, it's just confusing.

...These color maps were totally useless **without text, without any context or explanation**, like those details. The color maps were just pretty, but they didn't explain anything.

– Neurosurgeon #3

““

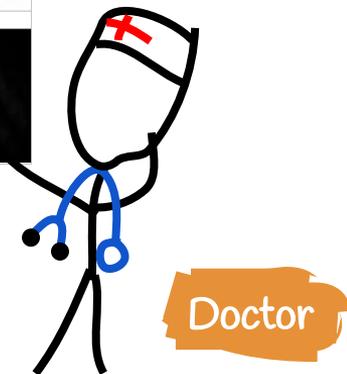
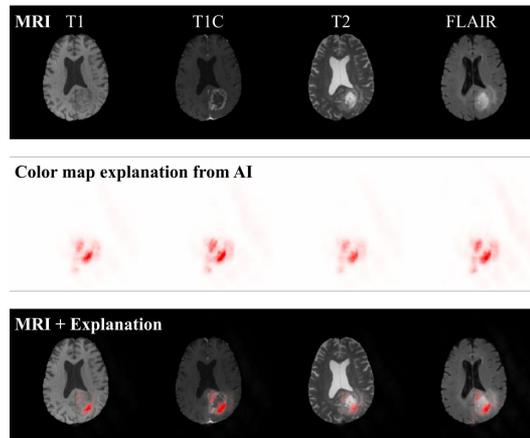
Though the color map is drawing your eyes to many different spots, but I feel like I didn't understand why my eyes were being driven to those spots, like **why were these very specific components important?** And I think that's where all my confusion was.

– Neurosurgeon #2

## Qualitative results

### Why AI explanation did not help?

Because the existing AI explanations do not explain in a clinically relevant way



# Existing AI explanations do not speak clinical language to explain



What (explanation) we get currently, when a radiologist read it, they **point out the significant features**, and then they **integrate those knowledge**, and say, to my best guess, this is a glioblastoma. And I have the same expectations of AI (explanation).

– Neurosurgeon #3

*Physicians' clinical image interpretation process:*

Medical image



Human-interpretable feature

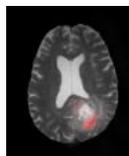
Human-interpretable reasoning based on the features

Clinical decision

Tumor grade 4

*Physicians' interpretation process of AI explanation:*

Heatmap explanation



Contrast-enhanced region of the tumor

Contrast-enhanced region is an indicator of higher grade tumor

Clinical decision

Tumor grade 4

“Context of the important features”

## Takeaway A phase 2 clinical user study of AI on glioma grading

- A national clinical study with 35 neurosurgeons on glioma grading task using MRI
- With **AI prediction** assistance, doctors significantly improved task performance
  - Due to overreliance on AI suggestions
- Additional **AI explanation** did not change doctors' performance
  - Due to lack of clinically relevant information in AI explanation

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*Thank you!*