

S1. Generation of Heatmaps for Model Explainability

Input (prior, subsequent, time_interval, time_start)

- 1 **Reference score** → survival score on the original images
 - 2 **ROI** → cube of 64x64x64 in the top left back corner
 - 3 **Occluded images** → set intensities within the **ROI** to zero in both prior and subsequent scan
 - 4 **Occluded score** → Compute the survival score on the occluded_images
 - 5 **ROI importance** → $| \text{occluded_score} - \text{reference_score} |$
 - 6 **Prognostic map[ROI]** → $\text{maximum}(\text{prognostic_map}[\text{ROI}], \text{roi_importance})$ [A]
 - 7 Move the **ROI** 8 voxels along one of the axis
 - 8 If **ROI** has not scrolled through the whole image yet, go to **Step 3**
 - 9 **Deformation map** → anatomical changes between prior and subsequent, as returned by the registration
 - 10 **Return** $\text{gaussian_filter}(\text{deformation_map} \times \text{prognostic_map})$ [B]
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[A] Since the ROI are overlapping, we chose to use the maximum function, which prevents erroneous overriding of previous estimation. [B] We fuse the prognostic map and the deformation map together (S3-4.0) to refine the hotspots to regions of changes, and therefore help along the radiologist in the visual interpretation.