## **Supplementary Notes 1: Details of Data Augmentation**

Following up on the Data Augmentation sub-section in Methods, this section contains a tabular illustration of *some* of the data augmentation techniques integrated into GaNDLF. The variety showcased here highlights the built-in flexibility of the entire framework.

Туре	Augmentation	Description of Specific Application
Spatial	Affine	Random affine transformations
	Elastic	Dense random elastic deformations
	Flipping	Reversal of the order of elements in an image along the given axes
	Rotation	Rigid rotations of 90 or 180 degrees across the specified axes
	Anisotropic	Down-sample and up-sample images along the provided axes
Intensity	Blur	Blurring using a random-sized Gaussian filter
	Noise	Gaussian noise with random parameters
	Gamma	Random change of contrast by raising values to the power of $\gamma$
MRI Space	Bias field	Random MRI bias field artifact
	MRI motion	Random motion artifact
	Ghosting	Random ghosting artifact
	Spike	Random spike artifacts

Supplementary Table 1: All available data augmentations provided in GaNDLF.

## **Supplementary Methods: Network Architectures**

GaNDLF seeks to provide both well-established and state-of-the-art network architectures showing promise in the field of healthcare. Since the literature on novel network architectures is continuously being expanded, at the time of publication the following list/table of architectures are offered by GaNDLF, the topologies of which are shown in the Methods section. Detailed description of each architecture is provided in the Supplementary Material.

- **UNet**: The UNet with (**ResUNet**) and without residual connections<sup>1–5</sup> (Supplementary Figure 1) is one of the most well known architectures of Convolutional Neural Networks (CNN) used for 2D and 3D segmentation. The UNet consists of an encoder, comprising convolutional layers and downsampling layers, and a decoder offering upsampling layers (applying transpose convolution layers) and convolutional layers. The encoder-decoder structure contributed in automatically capturing information at multiple scales/resolutions. The UNet further includes skip connections, which consist of concatenated feature maps paired across the encoder and the decoder layer, to improve context and feature re-usability.
- Fully Convolutional Network (FCN): The FCN architecture<sup>6</sup> (Supplementary Figure 2) introduced in 2017, utilizes hierarchical feature extraction with an encoder recognizing both imaging patterns and spatial information of each input image, with varying receptive fields. FCN has smaller computational requirements compared to UNet, due to the absence of the decoding module, incorporating convolution and transpose convolution operations. FCN simply upsamples the encoded features to the required output segmentations to generate masks. It hence provides faster, yet coarser, segmentations for various domains<sup>7</sup>.
- **Inception UNet (UInc)**: The Inception module<sup>8,9</sup> can be used to substitute the standard convolutional block (which is a simple series of convolutional layers) of the UNet to create the UInc architecture (Supplementary Figure 3). This module describes parallel pathways of convolutional layers of different kernel sizes, to improve the representation of multi-scale features. UInc has been applied towards semantic segmentation workloads<sup>10</sup>.
- **Spatial Decomposition Network (SDNet)**: The SDNet<sup>11</sup> (Supplementary Figure 4) is a well-known content-style disentanglement model for medical image segmentation. SDNet uses two different encoders to separate anatomy from appearance; a UNet encodes the anatomical information into a spatial representation and a variational autoencoder encodes the appearance into a vector one. The encoded anatomical information is represented as multi-channel binary maps of the same resolution as the input. A segmentation module is applied on the anatomy latent space to learn to predict the segmentation masks. A decoder is responsible for reconstructing the input by combining the two latent variables at multiple levels of granularity using AdaIN layers<sup>12</sup>. The original architecture uses FiLM layers<sup>13</sup> to combine these variables and a mask discriminator to support semi-supervised learning. GaNDLF currently supports the fully supervised training scheme.
- **TransUNet**: The TransUNet<sup>14</sup> (Supplementary Figure 5) architecture is a variant of UNet that uses a CNN-transformer hybrid encoder rather than just a CNN (UNet) or transformer (UNetR) encoder. This allows for the transformer portion of the encoder to capture long-range dependencies and global context using self-attention, while the leveraging the

resolution of the CNN feature maps. GaNDLF supports TransUNet of variable depth and will scale the number of transformer layers accordingly. GaNDLF supports the use of TransUNet on both 2D and 3D images.

- **UNetR**: The UNetR<sup>15</sup> (Supplementary Figure 6) architecture is a variant of UNet that uses a transformer encoder rather than the traditional CNN-based encoder. The transformer allows for capturing of long-range dependencies and global context, and it consists of a multi-head self-attention module that maps between a query and the value and key representations followed by a multilayer perceptron. GaNDLF supports UNetR of variable depth and will scale the number of transformer layers based on the size of the input image. GaNDLF supports the use of UNetR on both 2D and 3D images.
- VGG: The VGG<sup>16,17</sup> (Supplementary Figure 7) is a well-known network for performing classification and regression workloads. The original VGG has 16 convolutional layers and 3 dense layers. We have modified the final classifier layers to include a global average pooling layer followed by a single dense layer, which allows greater flexibility<sup>18</sup> for different types of workloads and reduce the effect of overfitting due to dense layers<sup>19</sup>. It is well known for its performance on the ImageNet classification challenge<sup>20</sup>. VGG reinforced the idea that networks should be simple and deep. VGG uses 3 × 3 convolution filters and 2 × 2 max-pooling layers with a stride of 2 throughout the architecture. The original architecture uses ReLU activation function<sup>21</sup> and categorical cross-entropy loss function. The initial layers of the VGG perform feature extraction and the last softmax layers act as the classifier. GaNDLF supports multiple variants of the VGG, namely, VGG11, VGG13, VGG16, VGG19, with and without batch normalization for both 2D and 3D datasets to maximize flexibility.
- **DenseNet**: The DenseNet<sup>22</sup> (Supplementary Figure 8) architecture is a type of convolutional neural networks that consist of dense blocks, where each layer in the block is densely connected, which is a mechanism to address the vanishing-gradient problem<sup>23</sup>. A unique property of the dense connections in DenseNet is that the previous layer's output and the current layer's output are concatenated instead of getting added. After the concatenation, there is a pooling layer, batch normalization, and non-linear activation layer. The DenseNet architecture can be customized based on the number of layers, and GaNDLF currently supports the DenseNet-121, DenseNet-160, DenseNet-201 and DenseNet-264 variants.
- **ResNet**: The ResNet<sup>3</sup> (Supplementary Figure 9) module uses shortcut connections to allow for learning of deeper architectures while avoiding the degradation of training accuracy. When the dimensions of the input and output to the block are the same, identity mapping is used, which does not introduce new parameters or increase computation complexity. If the dimensions change, linear projections are applied by the shortcut connection to match dimensions. GaNDLF supports variants of ResNet, including ResNet-18, ResNet-34, ResNet-50, ResNet-101, and ResNet-152. ResNet-18 and ResNet-34 use a pair of  $3 \times 3$  convolutions for each block, while ResNet-50, ResNet-101, and ResNet-152 use a bottleneck of  $1 \times 1$ ,  $3 \times 3$ , and  $1 \times 1$  convolutions to reduce the number of parameters.
- EfficientNet: The EfficientNet<sup>24</sup> (Supplementary Figure 10) module uses a compound scaling method to uniformly network scale width, depth, and resolution using a set of fixed constants, which allows for networks to be scaled while achieving improved accuracy without sacrificing efficiency. The base architecture uses mobile inverted bottleneck convolutions (MBConv) with squeeze-and-excitation optimization, which increases efficiency by the use of a narrow to wide to narrow approach rather than the wide to narrow to wide approach of residual blocks. GaNDLF supports multiple variants of EfficientNetB0 through EfficientNetB7.
- **ImageNet-trained 2D models**: GaNDLF also provides functionality of transfer learning based on popular architectures pre-trained on the ImageNet data<sup>20</sup>. Every architecture's first and last layers are modified to be able to process input images of any size, and only output the relevant number of classes for each problem, respectively. The rest of the layers retain weights from the ImageNet data. This allows for more efficient training, with a potential for better convergence result<sup>25</sup>.





























Supplementary Figure 8: DenseNet, a well-known architecture for tackling regression and classification tasks.



Supplementary Figure 9: ResNet, a well-known state-of-the-art architecture for tackling regression and classification tasks.



Supplementary Figure 10: EfficientNet, a well-known state-of-the-art architecture for tackling regression and classification tasks.

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