Supplementary Material.

Cumulative order-preserving image transforming network

In the original paper, the Google DeepMind demonstrated their spatial transformer network with three different popular transformations: affine, perspective and thin-plate spline transformation.¹⁶ It is clear that affine or perspective transformation will not be suitable for modeling changes of structures in the anterior chamber. While we suspect thin plate spline transformation to be capable of modeling anterior chamber changes, we have decided to develop a new image transforming network, which is computationally simpler and easier to implement.²⁷ The only condition imposed on the new transformation is preservation of the original order of *x* and *y* coordinates, because anatomic structures are not likely to change their order; e.g., the corneosclera or the iris is not likely to be twisted.

Following the original definition, the output pixels are defined to lie on a regular grid $G = \{G_{i,j}\}$ of pixels $G_{i,j} = (x_{i,j}^t, y_{i,j}^t)$. As AS-OCT images have been aligned so that horizontal midline of an image lies approximately at the center of the *x*axis and the top of the cornea lies at almost the same position on the yaxis, we define *G* with two $m \times n$ matrices X^t and Y^t corresponding to *x* and ycoordinates as follows:

$$X^{t} = \begin{bmatrix} x_{1,1}^{t} & \cdots & x_{1,p}^{t} & x_{1,p+1}^{t} & x_{1,p+2}^{t} & \cdots & x_{1,n}^{t} \\ x_{2,1}^{t} & \cdots & x_{2,p}^{t} & x_{2,p+1}^{t} & x_{2,p+2}^{t} & \cdots & x_{2,n}^{t} \\ \vdots & \ddots & \vdots & \vdots & \vdots & \ddots & \vdots \\ x_{m,1}^{t} & \cdots & x_{m,p}^{t} & x_{m,p+1}^{t} & x_{m,p+2}^{t} & \cdots & x_{m,n}^{t} \end{bmatrix} = \begin{bmatrix} -p & \cdots & -1 & 0 & 1 & \cdots & p \\ -p & \cdots & -1 & 0 & 1 & \cdots & p \\ \vdots & \ddots & \vdots & \vdots & \vdots & \ddots & \vdots \\ -p & \cdots & -1 & 0 & 1 & \cdots & p \end{bmatrix}$$

and
$$Y^{t} = \begin{bmatrix} y_{1,1}^{t} & y_{1,2}^{t} & \cdots & y_{1,n}^{t} \\ y_{2,1}^{t} & y_{2,2}^{t} & \cdots & y_{2,n}^{t} \\ \vdots & \vdots & \ddots & \vdots \\ y_{m,1}^{t} & y_{m,2}^{t} & \cdots & y_{m,n}^{t} \end{bmatrix} = \begin{bmatrix} 0 & 0 & \cdots & 0 \\ 1 & 1 & \cdots & 1 \\ \vdots & \vdots & \ddots & \vdots \\ m - 1 & m - 1 & \cdots & m - 1 \end{bmatrix}$$
(3)

where $p = \frac{n-1}{2}$ for odd number *n*. For even values of *n*, we modify X^t in the equation (3) to make it centered:

$$X^{t} = \begin{bmatrix} x_{1,1}^{t} & \cdots & x_{1,p-1}^{t} & x_{1,p+1}^{t} & x_{1,p+2}^{t} & \cdots & x_{1,n}^{t} \\ x_{2,1}^{t} & \cdots & x_{2,p-1}^{t} & x_{2,p}^{t} & x_{2,p+1}^{t} & x_{2,p+2}^{t} & \cdots & x_{2,n}^{t} \\ \vdots & \ddots & \vdots & \vdots & \vdots & \vdots & \ddots & \vdots \\ x_{m,1}^{t} & \cdots & x_{m,p-1}^{t} & x_{m,p}^{t} & x_{m,p+1}^{t} & x_{m,p+2}^{t} & \cdots & x_{m,n}^{t} \end{bmatrix}$$
$$= \begin{bmatrix} -(p-0.5) & \cdots & -1.5 & -0.5 & 0.5 & 1.5 & \cdots & p-0.5 \\ -(p-0.5) & \cdots & -1.5 & -0.5 & 0.5 & 1.5 & \cdots & p-0.5 \\ \vdots & \ddots & \vdots & \vdots & \vdots & \vdots & \ddots & \vdots \\ -(p-0.5) & \cdots & -1.5 & -0.5 & 0.5 & 1.5 & \cdots & p-0.5 \end{bmatrix}$$
(4)

where and $p = \frac{n}{2}$.

For even number *n*, we define two $m \times n$ matrices *R* and *S*, which corresponds to the intervals between coordinates of the regular grid X^t and Y^t :

$$R = \begin{bmatrix} -1 & \cdots & -1 & -0.5 & 0.5 & 1 & \cdots & 1 \\ -1 & \cdots & -1 & -0.5 & 0.5 & 1 & \cdots & 1 \\ \vdots & \ddots & \vdots & \vdots & \vdots & \vdots & \ddots & \vdots \\ -1 & \cdots & -1 & -0.5 & 0.5 & 1 & \cdots & 1 \end{bmatrix} \text{ and } S = \begin{bmatrix} 0 & 0 & \cdots & 0 \\ 1 & 1 & \cdots & 1 \\ \vdots & \vdots & \ddots & \vdots \\ 1 & 1 & \cdots & 1 \end{bmatrix}$$
(5)

Then the regular grid can also be defined as the cumulative sum of intervals from the origin:

$$X^{t} = \left[R_{l} \cdot L_{p} \mid R_{r} \cdot U_{p}\right] \text{and } Y^{t} = \left((S^{T}) \cdot U_{m}\right)^{T}$$

$$\tag{6}$$

were R_l and R_r are the left and right half of the matrix R, L is a square lower triangular matrix of ones and U is a square upper triangular matrix of ones.

The central idea behind COPIT is to adjust the distances between the coordinates such that the order of the *x* and *y* coordinates are preserved. A COPIT parametrized by θ is defined with two components as $(x_{ij}^s, y_{ij}^s) = T_{\theta}(G_{i,j}) = M_{\theta}(G_{i,j}) + A_{\theta}(G_{i,j})$, where M_{θ} is a multiplicative component of T_{θ} and A_{θ} is an additive component of T_{θ} .

A multiplicative component is calculated according to the formula (6) but with modified intervals:

$$M_{\theta}(X^{t}) = M_{x}(W_{\theta}) = \left[\phi(W_{l}) \circ R_{l} \cdot L_{p} | \phi(W_{r}) \circ R_{r} \cdot U_{p}\right] \text{and}$$
$$M_{\theta}(Y^{t}) = M_{y}(V_{\theta}) = \{(\phi(V_{\theta}) \circ S)^{T} \cdot U_{m}\}^{T}$$
(7)

where $\phi(z) = \begin{cases} \frac{1}{1-z} & z < 0, \\ z+1 & z \ge 0 \end{cases}$, ois the Hadamard product, W_l and W_r are the left and right half of the $m \times n$ matrix, and W_{θ} , V_{θ} is a matrix of size $m \times n$. If the elements of W_{θ} and V_{θ} are real numbers, the intervals are multiplied by positive real numbers, then the elements of $\phi(W_l), \phi(W_r)$ and $\phi(V_{\theta})$ are positive real numbers. Hence, the order of x and y coordinates are preserved after transformation M_{θ} . Also, M_{θ} is an identity transformation if all elements of W_{θ} and V_{θ} equal zero. The transformation M_{θ} does not affect the origin of the coordinates (0,0). Hence, we define an additive component of the transformation A_{θ} , which performs a translation to the origin:

$$A_{\theta}(X^{t}) = A_{x}(E_{\theta}) = E \cdot I_{n}^{T} \text{ and}$$
$$A_{\theta}(Y^{t}) = A_{y}(F_{\theta}) = I_{m} \cdot F^{T}$$
(8)

where E_{θ} and F_{θ} are column matrices of real numbers parametrized by θ with size $m \times 1$ and $n \times 1$, and I_m and I_n are a column matrix of ones of size $m \times 1$ and $n \times 1$.

Any number of transformations can be combined without violating the order-preserving properties if we define a new operator \boxplus as follows:

$$T_{\alpha\beta}(G) = T_{\alpha}(G) \boxplus T_{\beta}(G) = T_{\beta}(G) \boxplus T_{\alpha}(G) = \left(M_{x}(W_{\alpha} \circ W_{\beta}) + A_{x}(G_{\alpha} + G_{\beta}), M_{y}(V_{\alpha} \circ V_{\beta}) + A_{y}(H_{\alpha} + H_{\beta})\right)$$
(9)