Color Schemes to Represent the Orientation of Anisotropic Tissues From Diffusion Tensor Data: Application to White Matter Fiber Tract Mapping in the Human Brain

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This paper investigates the use of color to represent the directional information contained in the diffusion tensor. Ideally, one wants to take into account both the properties of human color vision and of the given display hardware to produce a representation in which differences in the orientation of anisotropic structures are proportional to the perceived differences in color. It is argued here that such a goal cannot be achieved in general and therefore, empirical or heuristic schemes, which avoid some of the common artifacts of previously proposed approaches, are implemented. Directionally encoded color (DEC) maps of the human brain obtained using these schemes clearly show the main association, projection, and commissural white matter pathways. In the brainstem, motor and sensory pathways are easily identified and can be differentiated from the transverse pontine fibers and the cerebellar peduncles. DEC maps obtained from diffusion tensor imaging data provide a simple and effective way to visualize fiber direction, useful for investigating the structural anatomy of different organs. Magn Reson Med 42:526–540, 1999. © 1999 Wiley-Liss, Inc.

Key words: diffusion; tensor; color; white matter; MRI; anisotropy; brain; human

In recent years, MRI has increasingly been used to study tissue water diffusion in vivo. By acquiring diffusion-weighted images with diffusion gradients oriented in at least six noncollinear directions, it is possible to measure the diffusion tensor \( \mathbf{D} \) in each voxel (1,2). Diffusion tensor data can, in turn, be used to compute quantities that characterize specific features of the diffusion process, such as the trace of the tensor [Trace(\( \mathbf{D} \))], indices of diffusion anisotropy, and the principal directions of diffusion (eigen vectors of \( \mathbf{D} \)) (See Ref. 3 for a review). Trace(\( \mathbf{D} \)), which is proportional to the orientationally averaged water diffusion coefficient, can better delineate ischemic and infarcted brain regions than single-direction apparent diffusion coefficients (4,5). Anisotropy measures yield information about the structure of normal and pathological white matter that cannot be obtained from conventional, relaxation-based MRI (6,7). The principal directions of diffusion provide information on the spatial orientation of anisotropic structures that may be useful to further characterize tissue structural anatomy. For example, in studying cardiac and skeletal muscle fibers, direction mapping might provide insight about the relationship between the structure of the muscle tissue and its function during different physiological and pathological conditions (8–11). Most of what we know in this regard has been derived from invasive procedures on isolated organs. In the brain, fiber direction mapping is useful to identify and differentiate anatomical white matter pathways that have similar structure and composition but different spatial orientation (12,13). Historically, such studies of the brain's structural anatomy have been performed only with histological methods.

While Trace(\( \mathbf{D} \)) and anisotropy measures are scalar quantities, easily displayed using gray scale maps, representing the directional information contained in the diffusion tensor is more problematic. Previously proposed approaches include: gray scale maps of the polar and azimuthal angles of the eigenvector associated with the largest eigenvalue of the diffusion tensor \( V_{\text{max}} \) (14,15); two-dimensional (2D) line fields of the in-plane projection of \( V_{\text{max}} \) (16); display of diffusion ellipses (17) and octahedra (8,9); and three-dimensional (3D) rendering of diffusion ellipsoids (6,18–20) in each voxel. Theoretically, these are rigorous approaches but they face the problem of visualizing 3D objects in two dimensions. For example, cigar-shaped diffusion ellipsoids would be indistinguishable from spheres if the anisotropic structure is pointing at the viewer (i.e., the largest diffusivity is parallel to the viewing axis). Displaying fiber direction using 3D rendering of diffusion ellipsoids or other solids could be substantially improved by viewing the structure from many different angles using computerized animation.Still, 3D rendering is suitable to display only small portions of an image and is cumbersome for global viewing of an entire image or a series of images.

Other methods to visualize fiber direction have used colors (21–26). Some of these methods assign different color components to diffusion-weighted images (DWI) or apparent diffusion coefficient (ADC) maps acquired with gradients applied in perpendicular directions (21,23). As acknowledged by the authors who proposed these schemes, these nontensor-based approaches fail to correctly describe fiber direction when \( \mathbf{D} \) has nonzero off-diagonal elements, a condition often encountered in biological tissues (15,20). It has been suggested that a color RGB representation of the components of \( V_{\text{max}} \), weighted, or filtered, by some measure of diffusion anisotropy, may provide a concise and effective way to display the directions of anisotropic structures in a single image (25,26). Using directionally

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encoded color (DEC) maps is appealing because it facilitates displaying the orientation of fibers that run at an angle to the image plane. Previously proposed tensor-based DEC schemes (25,26) overcome the problems inherent in DWI- or ADC-based schemes, but still have limitations that preclude them from reliably and quantitatively representing fiber direction (see Background section below). Here we propose improved tensor-based DEC mapping schemes that produce a simple, intuitive, and faithful representation of fiber direction. Finally, we use these schemes to identify and display white matter fiber tracts in the normal human brain.

BACKGROUND

We present some objectives that we think should be considered in designing a tensor-based quantitative color representation of fiber direction: (a) Perceived color differences should be proportional to actual differences in direction, characterized either by the Euclidian distance between vectors or by the angle between them. (b) Fiber orientation should be specified in an anatomically based coordinate system, independent from the laboratory reference frame in which the tensor dataset is acquired. (c) Only the directional information contained in the D should be displayed, and the representation should not be affected by parameters related to the magnitude of diffusivity, such as Trace(D) or diffusion coefficients. Some measure of anisotropy, however, could be used as a mask or filter to avoid representing directions in isotropic structures in which a fiber tract direction would be meaningless (25,26). (d) Commonly identifiable color groups (e.g., red, green, blue, yellow, cyan, magenta, etc.) should be approximately associated with a characteristic direction (e.g., direction along x, y, z, directions bisecting the xy, yz, xz planes, etc.).

Previously proposed DEC schemes (25,26) satisfy only a few of these requirements and, in particular, do not address the issues of points (a) and (b) above. To design improved DEC schemes, we investigated three sources of misrepresentation: (i) nonlinear effects of the display device, (ii) properties of human color vision, and (iii) antipodal symmetry of the eigenvectors of D.

Nonlinear Effects of the Display Device (Gamma Correction)

For a particular color to have the same appearance on different devices, one has to use a device-independent color specification [e.g., CIE XYZ+ system (27)]. This is attained through linear transformations of the primary colors for different devices (28). The next step is to account for the nonlinear relationship existing between the true luminosity produced by the device, L, and the digital values stored in the frame buffer, f (29). For each color channel of a cathode ray tube (CRT), this relationship can be written in the simplest form as L = fγ, where γ is the characteristic exponent for a given channel. Once the relationship is determined, one can apply a correction known as the gamma correction. When applying this correction, one has to ensure that the display software or the hardware look-up tables are not already performing it. Typical values for γ are γ ∈ [2.0, 2.4] for almost all CRT-based computer displays except for Macintosh (γ ≈ 1.5) and SGI (γ ≈ 1.3) computer displays. The formula is primarily derived for the CRT device, but is frequently applied to other devices, using different parameter values. With the advent of new display hardware, the nature of the nonlinear relationship can become quite different.

Properties of Human Color Vision

In order to construct a color representation scheme in which perceived color differences are proportional to actual differences in direction, we must consider not only the properties of the display device, but also the properties of our color vision. Throughout the history of color research, attempts have been made to define color ordering schemes based on the perceived difference between colors. Although color spaces have metric properties, the goal to find a uniform color space in which the perceived color difference is proportional to the Euclidean distance in such a space has remained elusive. Moreover, in complex scenes, such as color medical images, color perception involves the perception of contrast, i.e., the perceived color in a given region is influenced by the color of all the surrounding regions. Using the retinex model (30), one can predict the perceived color qualities for any region of a given image. However, constructing color mappings schemes based on the retinex theory having general validity for different images is impossible. If we ignore the effects of contrast, a perceptually uniform color representation can be constructed using approximately uniform color spaces, such as the CIEL*a*b* color space (27). Even a less elaborate approach should not ignore some fundamental relationships in color perception, such as the simple nonlinear relationship between the true luminosity of monochromatic light and its perceived brightness (Steven’s law).

Antipodal Symmetry of the Eigenvectors of D (Discontinuity Artifacts)

The sign of the eigenvectors of D is arbitrary since the parallel and anti-parallel vectors convey the same information (Vmax = −Vmax). In statistics, directional data with such antipodal symmetry are referred to as axial data. To have a unique vector representation of axial data, the vectors have to be mapped to only one hemisphere in a spherical coordinate system. The plane that divides the spherical coordinate system into two hemispheres can be positioned in an arbitrary direction; however, within this plane, fiber orientation will still be represented by two antipodally symmetric vectors Vmax and −Vmax. We call this the plane of discontinuity because a color representation mapped on the surface of a sphere will be discontinuous at the intersection of the sphere with this plane (see Fig. 1). For any structure oriented nearly parallel to the plane of discontinuity a small change in orientation can cause the Vmax to cross the plane of discontinuity and be represented with a very different color. This small change in orientation of the structure in the neighboring voxels can result from the natural curvature of the structure or can be due to noise in measuring Vmax. In general, a continuous

¹CIE stands for Commission Internationale de L’Eclairage (International Color Commission). The three letters following CIE indicate the type of color coordinates used in a given color system.
As in previously proposed schemes (25), the brightness of our DEC maps is weighted by a measure of diffusion anisotropy to avoid representing isotropic or slightly anisotropic structures in which \( V_{\text{max}} \) will have virtually random orientation. The anisotropy measure we used is the lattice index, \( \Lambda \) (20), which is a linear combination of its basic elements \( \Lambda_N \), defined as:

\[
\Lambda_N = \sqrt{3} \frac{D_{\text{ref}} D_N}{8 D_{\text{ref}} D_N} + \frac{3}{4} \frac{D_{\text{ref}} D_N}{D_{\text{ref}} D_{\text{ref}} D_N D_N} \tag{1}
\]

where \( D_{\text{ref}} : D_N \) denotes the tensor dot product between the diffusion tensor of the given voxel \( D_{\text{ref}} \) and that of the neighboring voxel \( D_N \). \( D \) (italic) indicates the anisotropic part the diffusion tensor or deviatoric tensor. The formula of the basic elements of the lattice index is reported here because it contained typographical errors in the original publication (20). A more detailed description of the quantities appearing in the formula and the properties of the lattice anisotropy index are reported in reference (20). We implemented two types of anisotropy filters in our color representations. In the first one, each of the color components contains a weighting factor \( \Lambda_w \), which linearly increases from 0 to 1 within a specified range of lattice index values \( \Lambda \in [\Lambda_{\text{min}}, \Lambda_{\text{max}}] \). In the second, only structures where the lattice index is larger than \( \Lambda_{\text{min}} \) are represented; the others are truncated and set to black (see Appendix A for details).

Color Maps and Symmetry Considerations
To reduce the discontinuity artifacts mentioned in (iii) of the Background section, we allowed different directions to be represented with the same color if some form of symmetry exists between these directions. In this case, we have an orientational ambiguity because the color representation of orientation is not unique. In some situations, however, using such symmetries is justified by existing symmetries in the imaged anatomy. With regard to which type of symmetry is used, we distinguished the following cases:

No Symmetry. This representation is implemented by relating the azimuthal angle of the vector \( \psi \) to color hue and the polar angle \( \theta \) to color saturation. For example, a simple implementation in HSV (Hue, Saturation, Value) color space is \( H = \psi, S = 2\sin(\pi/4), V = \Lambda_w \). This representation is unique because a particular color describes a single direction but suffers from discontinuity artifacts.

Rotational Symmetry (2-Fold). In this color representation any vector and its pair rotated by 180° around the z axis will have the same color representation \( V_{\text{max}}(0, \psi) = V_{\text{max}}(0, \psi + \pi) \), or \( V_{\text{max}}(\psi, 0, \psi) = V_{\text{max}}(-\psi, 0, -\psi) \). This representation is implemented in a fashion similar to the no symmetry approach but with the constraint \( H(\psi) = H(\psi + \pi) \). This scheme removes the discontinuity artifacts but does not provide a unique representation of directions and is, also, perceptually highly nonuniform (see Table 1 for a summary of the properties of different color schemes).

Mirror Symmetry. The mirror symmetric representation implies that any two vectors that are mirror images of each other relative to the yz plane of the anatomical coordinate system for directional data with antipodal symmetry cannot be constructed; hence, the metric properties of the color spaces cannot be used for representation. However, a metric can be constructed if we assign zero distance, and therefore the same color, to all vector pairs that satisfy a 2-fold rotational symmetry around the z axis (i.e., all vector pairs that are images of each other under 180° rotations). In such cases, the antipodally symmetric vector pairs lying within the plane of discontinuity will correctly have the same representation.

**MATERIALS AND METHODS**

**Color Representation of Fiber Orientation**

We assumed that the orientation of fibers is described by the eigenvector associated with the largest eigenvalue, \( V_{\text{max}} \). We established a coordinate system fixed to defined anatomical landmarks (Fig. 1a) satisfying the condition (b) described in the Background section. Proper rotations were used to convert from the laboratory reference frame to the specified anatomical reference frame. The eigenvector \( V_{\text{max}} \) is a unit vector completely determined by its polar (ii) and azimuthal (\( \psi \)) angles that are defined within the anatomical coordinate system as shown in Fig. 1a. When no anisotropy filter is used, a particular color representation can be shown on a unit sphere, which we call the color representation sphere. Figure 1b shows the color representation sphere for the no symmetry scheme described later in this section. As mentioned before, fiber orientation is described by two vectors \( V_{\text{max}} \) and \( -V_{\text{max}} \), or two pairs of angles \( (0, \psi) \) and \( (\pi - 0, \psi + \pi) \), and only the vectors in one of the two hemispheres are used for unique representation. Unless stated otherwise, vectors are expressed in the upper hemisphere \( (z > 0) \) or \( (0 \leq \theta \leq \pi/2) \) and the xy plane is the plane of discontinuity. The lower hemisphere is just an antipodally symmetric copy of the upper hemisphere, as shown in Fig. 1b in which the color representation sphere is split along the plane of discontinuity.

![Figure 1](image-url)
system will have the same color representation \( \mathbf{V}_{\text{max}}(-v_xv_yv_z) = \mathbf{V}_{\text{max}}(v_xv_yv_z) \). Mirror symmetry exists between many structures in the left and the right brain hemispheres.

Absolute Value. Simultaneously assuming the existence of mirror and rotational symmetries together with the inherent antipodal symmetry \( \mathbf{V}_{\text{max}} = -\mathbf{V}_{\text{max}} \), implies that the sign of any vector component is unimportant; only the absolute value is. This yields a simple representation in RGB color space in which RGB components are associated with the absolute value of the components of the vector, as originally proposed in (25), i.e.: 

\[
R = \Lambda_w |v_x|, \quad G = \Lambda_w |v_y|, \quad B = \Lambda_w |v_z|
\]

Preferred Direction. When no symmetry is assumed and the structure of interest lies close to the plane of discontinuity, it is desirable to rotate the coordinates of the color space so that undesirable artifacts are shifted to a different plane. For a given region of interest (ROI), we determine the dominant fiber direction, or we specify a particular preferred direction that we want to explore. This is called the preferred direction and is described by the vector \( \mathbf{v}_p \). Once a preferred direction is selected, the no symmetry representation is implemented in the new coordinate system in which the new z axis coincides with \( \mathbf{v}_p \) (see Appendix A for details) and the discontinuity artifacts are now shifted to a plane perpendicular to \( \mathbf{v}_p \). Very often one is only interested in the structures along the chosen preferred direction and not in those close to the plane of discontinuity. Hence, we choose a single cutoff polar angle \( \theta_C \) (usually between 70° and 80°) to define the structures to be color coded \((\theta \leq \theta_C)\). Structures having \( \theta > \theta_C \), i.e., lying close to the plane where discontinuity artifacts occur, are displayed in black. Alternatively, one can use a cone defined by the preferred direction and a smaller angle \( \theta_C < 70° \) to increase the resolving power of the color representation around the direction of \( \mathbf{v}_p \). Only with the preferred direction approach is it appropriate to use the cutoff angle \( \theta_C \) since in the other schemes previously described, the plane of discontinuity is fixed and the same structures would always be eliminated.

Color Perception Considerations

We used two different approaches to address the properties of color perception. The first was based on the uniform color spaces CIEL*a*b* and CIELUV (27), and the second was an heuristic approach in which desirable properties were obtained empirically.

Uniform Color Space Approach. By using uniform color spaces, we hoped to improve the color representation despite the fundamental limitations described in (ii) of the Background section. Only the no symmetry and absolute value cases can be implemented within uniform color space in a relatively simple way and their implementation is briefly described in Appendix C.

Heuristic Approach. In order to alleviate some of the most pronounced artifacts associated with color display and perception [(i) and (ii) of the Background section], we propose schemes in which a perceptually satisfying representation is obtained interactively by adjusting a small number of parameters. These are called heuristic parameters to emphasize the empirical nature of this approach. These heuristic parameters are listed in Table 2 and a detailed explanation of their implementation is found in Appendix A. We used heuristic parameters \( p_B, p_S, \) and \( L_E \) to achieve approximately uniform brightness for different colors; \( p_B \) to modify the nonlinear relationship between the perceived brightness and the anisotropy measure; and \( p_S \) to modify the dependence between the saturation and the anisotropy and a more uniform color representation. We also treated as heuristic parameters both the gamma correction exponent \( \gamma \) and the Steven's law exponent \( \beta \), which describes the nonlinear relationship between the luminosity and perceived brightness.

Color Circles

Color circles were constructed by projecting the color representation sphere onto the viewing plane using the equal area Lambert projection (Fig. 2b). These color circles are useful to relate a particular color in the DEC map with its corresponding fiber orientation. For axial images the representation sphere is viewed from below (negative z axis of our anatomical coordinate system pointing at the viewer, see Fig. 2b) and for coronal images from the front (positive y axis pointing at the viewer, see Fig. 2b), according to the standard convention for viewing MR images. A grid of parallels and meridians is added to the circle at 15° intervals to indicate the angles \( \theta \) (angle at which the structure is positioned relative to the viewing plane) and colors corresponding to the viewing direction. We modified the scheme for maximum color representation in the structures along the chosen preferred direction and not in those close to the plane of discontinuity.
Table 2
List of Heuristic Parameters*

<table>
<thead>
<tr>
<th>Parameter symbol</th>
<th>Short description of the parameter</th>
<th>Range of suggested (upper field) and allowed (lower field) values</th>
</tr>
</thead>
<tbody>
<tr>
<td>( p_b )</td>
<td>Decreases the saturation of blue hues to achieve better uniformity of perceived brightness for different hues, and more faithful representation of diffusion anisotropy.</td>
<td>([0, 0.3/p_b] )</td>
</tr>
<tr>
<td>( p_c )</td>
<td>Used for equalization of the perceived brightness. For ( p_c = 0 ) RGB components are treated as equivalent; for ( p_c = 1 ) the components are weighted differently to achieve uniform brightness for different hues, similar to CIEL<em>u</em>v*.</td>
<td>([0, 1] )</td>
</tr>
<tr>
<td>( L_E )</td>
<td>Reference brightness to which colors are scaled. For higher values of ( L_E ) the larger values of ( p_b ) should be used and vice versa. For large ( L_E ) the equivalent of ( p_b ) for red should be used.</td>
<td>([0.6, 0.7] ) ( [0, 1] )</td>
</tr>
<tr>
<td>( p_E )</td>
<td>Control parameter. For ( p_E = 0 ) the maximal range of colors is used and uniformity is ignored ( (p_E, p_b, ) and ( L_E ) are disabled); for ( p_E = 1 ), ( p_b ), ( p_E ), and ( L_E ) are fully enabled and a smaller color range is used.</td>
<td>([0, 1] )</td>
</tr>
<tr>
<td>( p_S )</td>
<td>Modifies the relation between the hue and angle ( \theta ) in HSV-based schemes; ( p_S &gt; 0 ). For ( p_S = 0 ) relationship is linear. We use ( p_S = 0.5 ).</td>
<td>([0, 1] )</td>
</tr>
<tr>
<td>( p_i )</td>
<td>Modifies the exponent of the anisotropy index-based weighting factor ((\Lambda_a)). When ( p_i &lt; 1 ), structures with low anisotropy are emphasized.</td>
<td>([0.5, 1] ) ( [0, +\infty) )</td>
</tr>
<tr>
<td>( \beta )</td>
<td>Parameter based on Steven’s law exponent relating true luminosity and perceived brightness.</td>
<td>([0.3, 0.5] ) ( [0, \infty) )</td>
</tr>
<tr>
<td>( \gamma )</td>
<td>Parameter based on gamma correction exponent. For CRT, ( \gamma \in [1.3, 2.4] ); for other devices or if ( \gamma ) is unknown, search the allowed range starting from ( \gamma = 1 ).</td>
<td>([1.3, 2.4] ) ( [0, \infty) )</td>
</tr>
</tbody>
</table>

*Description of the primary role of each heuristic parameter. For some parameters in addition to the range of values that are theoretically allowed, we report the range of suggested values that are likely to produce the best results.

The anatomic coordinate system and the reference system for image display do not necessarily coincide, i.e., axial, coronal, and sagittal images are not lying in the xy, xz, and yz planes of the anatomical coordinate system, respectively. In such case, the color coordinate system appears shifted relative to the color circle grid (e.g., the white point in the circles in Fig. 5c, d, and e would not be exactly in the center of the circle) and the above proposed symmetries in color representation would not be easily seen on the color circle. If the misalignment between the anatomical coordinate system and the reference system for image display is severe, images should be interpolated and redisplayed after proper rotation. However, if the misalignment is small \((<10°)\), as in our case, errors introduced by displaying images in their original form and color circles in the anatomical reference frame are negligible.

MRI

The DEC maps presented in the figures were produced from diffusion data obtained from two 30-year-old female normal volunteers. We used a 1.5-T GE Signa Horizon EchoSpeed spectrometer (GE Medical Systems, Milwau­kee, WI), equipped with a whole-body gradient coil capable of producing gradient pulses up to 22 mT/m with a slew rate of 120 T/m/s. Diffusion images were acquired with an interleaved spin-echo echo-planar imaging sequence with a navigator echo for correction of motion artifacts. A description of the algorithms used for image reconstruction is presented elsewhere (31,32). Imaging acquisition parameters for axial sections were as follow: 32 contiguous slices, 3.5-mm slice thickness, rectangular 165 × 220-mm field of view, 96 × 128 in-plane resolution (6 interleaves, 16 echoes per interleaf), repetition time of greater than 5000 msec, echo time of 78 msec, and cardiac gating (4 acquisitions per heart beat starting with a 200 ms delay after the rise of the sphygmic wave as measured.

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by a peripheral pulse oxymeter). Six diffusion directions were sampled using combinations of the x, y, and z physical gradients as previously described (6). Each physical gradient had a strength of 21 mT/m, yielding an effective strength of 29.7 mT/m in the sampled diffusion direction. The value of the trace of the b-matrix (33), which is equal to the effective b-value in the sampled direction, was 1000 sec/mm². Four images with no diffusion gradients were also acquired for a total of 28 images per slice. The imaging time for the acquisition of the entire diffusion imaging data set was about 30 min. We also acquired conventional fast spin echo T1-, T2-, and proton density-weighted images matching the slices of the diffusion study, having however a higher in-plane resolution (192 × 256). Following image reconstruction, we computed the b-matrix numerically for each diffusion weighted image (33). We calculated D in each voxel according to Basser et al. (1,2), and generated maps of the lattice anisotropy index (20).

RESULTS

Figure 3 shows an axial section of the human brain with a conventional T2-weighted image (Fig. 3a), a map of the lattice anisotropy index (Fig. 3b), and a line field representation of V max visualized using a scheme similar to that presented by Tang et al. (16) (Fig. 3c). With 2D projection representations, and to a lesser extent also with representations based on 3D rendering of solids, such as diffusion ellipsoids, fiber direction is faithfully represented for structures positioned within the plane of the image, such as the corpus callosum. However, structures having a strong directional component along the viewing axis, such as the posterior limb of the internal capsule and the cingulum (arrows), are poorly represented.

Figure 4 shows the same axial section of normal human brain as in Fig. 3 using the previously proposed color representation schemes. Figure 5 shows implementation of the absolute value case in uniform color spaces (Fig. 5a) and several heuristic mapping schemes: absolute value (Fig. 5b), rotational symmetry (Fig. 5c), mirror symmetry (Fig. 5d), no symmetry (Fig. 5e), and the preferred direction (Fig. 5f) cases. The full effect of the heuristic approach, however, cannot be appreciated, since the heuristic parameter p c was set to 0.5 (see Table 2) to yield a larger range of colors and to achieve a better match between the printed and the displayed image.

A comparison between the DEC map suggested by Pierpaoli (25), displayed with no gamma correction (Fig. 4a) and with proper gamma correction (Fig. 4b), demonstrates the importance of properly handling this parameter. The color circle of Fig. 4a has large regions predominantly colored in red, green, and blue, that are separated by narrow boundaries between them. Consequently, fiber directions in the brain image appear artifactually segmented. For example, sharp color transitions are present between the blue and green regions in the posterior limb of the internal capsule (arrows) as well as between the red and green regions in the genu of the corpus callosum (arrows), suggesting abrupt variations in fiber direction. This indeed is an artifact of the color representation because the orientation of these structures varies smoothly, as confirmed by a histogram analysis of the same data, and as can be seen by the smoother transition in Fig. 4b, where gamma correction has been properly implemented. However, the image of Fig. 4b fails to provide an accurate representation of diffusion anisotropy. By comparing Fig. 4b to Fig. 3b, which displays a gray scale map of the lattice anisotropy index for the same section, structures represented in green appear to have a higher anisotropy than they really have, whereas structures represented in blue have a lower apparent anisotropy. This problem is partially corrected in Fig. 5b using the heuristic approach. Figure 4c shows the appearance of the DEC map suggested by Jones et al. (26). In Jones’ approach, voxels having a degree of anisotropy below a certain threshold (here λ min = 0.15) are not considered and are represented in black. This creates ambiguity because anisotropic structures running approximately perpendicular to the plane of the image (such as the internal capsule) are also represented in black. However, the main problem with this approach is that discontinuity artifacts are present. They consist of sharp artifactual boundaries between complementary colors occurring in structures that are approximately oriented within the plane of the image, such as the corpus callosum (arrows in Fig. 4c). Without prior knowledge of brain anatomy, it would be
impossible to discern whether these sharp color transitions are true boundaries between differently oriented structures or just artifacts of the color scheme.

Discontinuity artifacts were also present in the mirror symmetry (Fig. 5d) and no symmetry cases (Fig. 5e). Only the rotational symmetry (Fig. 5d) and absolute value (Fig. 5b) schemes were free of discontinuity artifacts. However, they do not provide a unique representation of directions; additionally, the rotational symmetry scheme produces a perceptually nonuniform representation (sensitivity along $\varphi$ is much greater than along $\theta$). Figure 5f shows that the preferred direction representation can be used to shift the discontinuity artifacts away from the structures of interest. The preferred direction that was used coincided with the average orientation of the fibers contained in the region of the corpus callosum indicated by the arrowhead. The black band in the color circle encompassed directions that were very close to the plane perpendicular to the preferred direction whose representations were eliminated by using the critical angle $\theta_C = 80^\circ$ (see Appendix B). In this way any color transitions, except those between black and saturated hues, reflect true anatomical features and do not originate from discontinuity artifacts. A summary of the three important properties of the color representation: presence of discontinuity artifacts, perceptual uniformity, and presence of orientational ambiguity, is presented in Table 1 for five different representations.

Figure 6a shows a T2-weighted image of an axial section of the pons. Figure 6b and c, show DEC maps of the same section using the absolute value and no symmetry heuristic equations, which are the least unique mapping scheme and the scheme with the worst discontinuity artifacts, respectively (see Table 1). Despite having an in-plane resolution that is higher than that of the DEC maps, the T2-weighted image did not identify any of the white matter pathways running in the pons at this level. On the contrary, the DEC map of Fig. 6b identified the descending motor pathways, ascending sensory pathways, and transverse pontine fibers. The cerebellar peduncles (superior, middle, and inferior) can also be easily recognized in Fig. 6b. In this figure, however, the nonuniqueness of fiber direction as a result of taking the absolute value of each component of $V_{\text{max}}$, can be observed in some regions. For example, it can be seen that the inferior cerebellar peduncle (ICP) and superior cerebellar peduncle (SCP), which have very different orientations, appeared with similar colors. The true orientations of ICP and SCP are indicated on the corresponding color circles. These ambiguities were eliminated using the no symmetry approach (Fig. 6c); however, discontinuity arose at the boundary between the left ICP and adjacent
FIG. 5. Different heuristic color schemes for fiber direction mapping. Maps of the same axial section of the human brain seen in Fig. 3 are shown using the formulas for (b) absolute value, (c) rotational symmetry, (d) mirror symmetry, (e) no symmetry, and (f) preferred direction. The arrowhead in (f) points to the region in the splenium of the corpus callosum that defines the preferred direction used in this case. The absolute value case implemented in CIEL*u*v* uniform color space is also presented (a).
FIG. 6. Axial section of the pons as depicted by (a) T2-weighted imaging, and DEC mapping using the (b) absolute value and (c) no symmetry heuristic equations.

fibers. This ambiguity can be resolved by using the preferred direction representation with an ROI positioned on the left ICP (not shown).

Figure 7 shows DEC maps obtained with the absolute value heuristic equation in five coronal sections of normal human brain. This is the DEC scheme we favored for initial viewing of fiber direction in brain imaging. Most of the main association, projection, and commissural pathways can be clearly identified in these images. It is interesting that neighboring white matter pathways having similar anisotropy (as well as similar signal intensity in conventional MRI) can be discriminated on the basis of their different directions. One interesting example is in the most posterior section (left bottom of the figure) where the optic radiation, running predominantly in an anterior-posterior direction, can be differentiated from the neighboring vertical occipital fibers oriented in a superior-inferior direction.

DISCUSSION

We have addressed the issue of representing fiber direction from diffusion tensor imaging data using colors. Compared to graphic schemes (vector fields, diffusion ellipses, octahedra, ellipsoids, etc.), color representation schemes potentially better represent fibers oriented at an angle with the plane of the image. Nevertheless, one important issue in color representation is to preserve the quantitative nature of the directional information contained in the diffusion tensor. In this paper, we identified and described three main sources of artifacts in previously proposed DEC schemes: (i) nonlinear effects of the display device; (ii) properties of human color perception; and (iii) antipodal symmetry of the eigenvectors of $\mathbf{D}$ (see the Background section). While (i) can be corrected, (ii) and (iii) pose fundamental limitations to achieving the ideal case of having perceived color differences proportional to the actual differences in direction.

It is a natural tendency of the human visual system to interpret separate regions with similar colors as belonging to the same object or structure. Thus, it is insufficient just to assign different colors to different directions and the perceived differences between the colors must also be taken into account. Although perceptually uniform color spaces can be constructed, their validity is limited when applied to complex color images. Additionally, the constraint of uniformity requires omitting a significant range of colors (see Appendix C and Fig. 5a), thus producing an unsatisfactory representation. Therefore, we favor an empirical or heuristic approach to the issues of color perception. For a given display device, we found that a perceptu-
FIG. 7. Set of coronal DEC maps constructed using the absolute value heuristic equation. All the main association, projection, and commissural pathways can be easily identified in these images. The color circle for coronal images has a different appearance from that used for axial images. This is because the DEC mapping scheme is defined in an anatomic coordinate reference system so that each structure maintains its color independently from the viewing angle, while the color circle is constructed to allow identification of directions in the reference frame of the plane in which the image is displayed, as described in Materials and Methods and in Fig. 2.
ally satisfactory representation could be obtained by adjusting the heuristic parameters and judging interactively the effects on the color circle which maps all possible directions. This approach is described in detail in Appendix A and Table 2. Improvements over previously proposed schemes are shown in the Results section. By expressing the colors obtained in a heuristic representation with \( L^*u^*v^* \) components of CIEL\( ^* \) space, the improvements over previously used schemes can be assessed quantitatively. For example, in trying to achieve the goal that perceived brightness indicates the degree of anisotropy, we want regions of constant anisotropy to correspond to regions of constant perceived brightness. When the anisotropy weighting factor \( \lambda_w \) is set to 1, for example, each color is set to its maximal brightness, the range of perceived brightness for different directions/colors is very large when no heuristic corrections are performed. In this case the perceived brightness ranged from \( L^* = 0.4 \) for the blue to \( L^* = 1 \) for white (\( L^* = 0 \) is black). When heuristic corrections were used (\( \rho_c = 1, \rho_B = 0.2, \rho_E = 1, L_E = 0.6 \)), this range was much narrower (\( L^* \in [0.59, 0.63] \)) indicating much better uniformity in the perceived brightness for the constant anisotropy index. Another significant limitation in achieving a quantitative color representation is related to the antipodal symmetry of the eigenvectors of \( D \). The problem stems from the fact that the antipodally symmetric directional data (axial data) cannot be mapped into a continuous metric space. If one tries to construct a representation in which a given color uniquely corresponds to a given direction, a discontinuity in the representation occurs resulting in artificial boundaries or undesirable speckled patterns (Figs. 4c and 5d and e). Schemes that assume rotational symmetry can alleviate this problem but they do not offer a unique color representation of directions (see Table 1). Another approach is to selectively represent structures by using the preferred direction scheme. When using the preferred direction scheme, one can be confident that boundaries between colored (nonblack) regions indicate true differences in direction, provided that \( \alpha_C \) is set sufficiently smaller than 90\(^\circ\). Only the boundaries between the black and saturated hues may be artifacts because black regions may represent either anisotropic structures lying close to the plane of discontinuity or isotropic tissue. This approach is best used for selective viewing within an ROI to visualize the orientation of a particular structure without ambiguity or artifacts.

**THE DEC MAPS PROTOCOL**

From the above, it is evident that using only a single DEC map leads to ambiguities and misrepresentation. We propose using different representation schemes to address each of the problems separately. The protocol that we implemented in our Institution enables one to use any of the mentioned coloring schemes and to choose between truncation or weighting with a rotationally invariant anisotropy index, such as the lattice index. We used the heuristic absolute value (Fig. 5b) and rotational symmetry (Fig. 5b) schemes to view initially brain images. Although these schemes do not provide a unique representation of fiber direction, they are void of discontinuity artifacts. The rotational symmetry scheme has a less ambiguous (more unique) representation of directions than the absolute value scheme because one color represents two directions instead of four. However, its discriminative power for angles in the polar (\( \theta \)) direction is poorer than in the azimuthal (\( \phi \)) direction, a disadvantage not present in the absolute value scheme. For a more detailed representation of fiber direction, images were also viewed with the heuristic no symmetry (Fig. 4c) scheme. Once a particular fiber tract is to be analyzed, we determined its average direction in a small ROI and used the preferred direction (Fig. 4d) scheme to further improve visualization of both the tract and similarly oriented neighboring structures.

Although we chose this protocol to work with brain images, alternative approaches may be more appropriate for other tissues and organs. For example, for images of the heart on sections perpendicular to the long axis, the rotational symmetry scheme may be optimal because of the directional arrangement of cardiac fibers.

**OTHER ISSUES**

In regions where the diffusion ellipsoids are prolate (cigar shaped), it is reasonable to assume that the fiber orientation is determined by the eigenvector associated with the largest eigenvalue, \( \mathbf{V}_{\text{max}} \). However, in regions where diffusion ellipsoids are oblate and axisymmetric (pancake shaped), using \( \mathbf{V}_{\text{max}} \) to indicate fiber direction is meaningless because \( \mathbf{V}_{\text{max}} \) is randomly oriented within the plane perpendicular to the direction of the smallest diffusivity. Some regions of the brain indeed have an architectural arrangement of fibers resulting in a diffusion displacement profile described approximately by oblate ellipsoids \( (6) \). Because these regions correspond to highly negative values of the skewness of the eigenvalues \( (15, 34) \), filtering by the skewness of the eigenvalues could be implemented, just as we used an index of diffusion anisotropy to mask voxels where diffusion is isotropic.

Implementation of our formulas is mainly intended for the 24-bit color graphics CRT devices, which we used in this work. If an 8-bit color display with a maximum of 256 colors has to be used, one needs to resort to sophisticated dithering schemes to achieve satisfactory results. If this is unavailable through display software, we suggest implementing a less accurate scheme that is a simple discretization of the above-mentioned representations. In it, the number of the discrete cells for vector representation has to be equal or smaller than the number of available colors, otherwise, colors will be approximated with the closest available ones and this often produces significant distortion. We viewed images only with 24-bit graphics cards. When printing images, we used the printer driver to perform device-independent color correction, including correction for the difference between the gamma exponents. Even high-quality color printers, however, cannot produce printouts of images that faithfully reproduce what one sees on the computer screen. It should be considered that our description of the figures is based on their appearance on the screen and their reproduction in the final print may not be accurate.

**CONCLUSIONS**

DEC maps can be used to represent the orientational information contained in the diffusion tensor. We identi-
fied several desirable properties that DEC mapping schemes should possess in order to provide a satisfactory representation of fiber direction. We also identified sources of errors that arose in representing this information using color. These are related to the discontinuity artifacts resulting from the anti-podal symmetry of eigenvector data as well as to the display and perception of colors. The goal of having a representation in which perceived color distances are proportional to the actual differences in orientation is, generally, not achievable. Based on our investigation and experience in working with DEC maps, we recommend that perceptual properties of human vision and the non-linear properties of the display devices be handled through an heuristic approach. A set of heuristic parameters is calibrated (tuned) by using color circles that are equal area projections of the color representation space, and by searching for a perceptually satisfactory representation on a given device. Discontinuity artifacts can be mitigated by taking advantage of symmetries encountered in the tissue, or by selecting a preferred direction so that the structure of interest is oriented perpendicular to the plane of discontinuity. The main goal of this work was not to suggest a particular implementation, but rather to address general issues involved in color representation of fiber direction that are relevant when constructing the appropriate DEC protocol for a particular imaging situation. Finally, by using our color approach to map white matter fiber tracts in the human brain, we have clearly identified the location and orientation of several white matter pathways which were not previously mapped in vivo using a noninvasive technique.

APPENDIX A: IMPLEMENTATION OF THE HEURISTIC APPROACHES

Color Maps of $V_{\text{max}}$

Here, we map the vector $V_{\text{max}}(v_x, v_y, v_z)$, or $V_{\text{max}}(\theta, \phi)$, into a RGB color space. Let $R_i$, $G_i$, and $B_i$ be the RGB components of the color representation of $V_{\text{max}}$, which are defined in the interval [0, 1]. In most schemes, $R_i$, $G_i$, and $B_i$ are obtained from HSV (Hue, Saturation, Value) components using the standard HSV to RGB conversion (28). The implementation of the color maps based on the four symmetries described in Materials and Methods are as follows:

No Symmetry

$$H = (\phi - \phi_R + 2\pi) \text{mod} 2\pi$$

$$S = \frac{\sin(p_S \theta)}{\sin(p_S \pi/2)}$$

$$V = 1$$  \hspace{1cm} [A1]

in which $\phi_R$ is the $\phi$ angle of the vector lying in the $xy$ plane that will be represented in a pure red color, and $p_S \in [0, 1]$ is an heuristic parameter (e.g., $p_S = 0.5$ in Fig. 5.c). When $p_S = 0$, $S$ is approximately a linear function of $\theta$. For the next two representations the $S$ and $V$ are the same as in [A1] and $H$ is as follows:

Rotational Symmetry

$$H = 2(\phi - \phi_R + 2\pi) \text{mod} 2\pi$$  \hspace{1cm} [A2]

Mirror Symmetry

$$\varphi$$ is azimuthal angle of $V_{\text{max}}(v_x, v_y, v_z)$

$$H = 2(\varphi - \phi_R + \pi) \text{mod} \pi$$  \hspace{1cm} [A3]

The formulas for the $H$, $S$, and $V$ components in the preferred direction approach are described separately in Appendix B. In the absolute value approach $R_i$, $G_i$, and $B_i$ are obtained directly as:

Absolute value

$$R_i = |v_x|$$

$$G_i = |v_y|$$

$$B_i = |v_z|$$  \hspace{1cm} [A4]

Anisotropy Filter

We allowed the brightness of a given color to be modulated by the degree of anisotropy. We implemented two types of anisotropy filters in our color representations. In the first one, each of the color components contains a weighting factor $\Lambda_w$ of the form $\Lambda_w = \max [0, \min(1, \Lambda - \Lambda_{\text{min}})/(\Lambda_{\text{max}} - \Lambda_{\text{min}})]$, where $\Lambda_{\text{min}}$ and $\Lambda_{\text{max}}$ delimit the range of values of the lattice index within which the weighting factor linearly increases from 0 to 1. In the second, only structures where the lattice index was larger than $\Lambda_{\text{min}}$ are represented, the others are truncated and set to black. In schemes based on perceptually uniform color spaces, the weighting by lattice index is included within the representation model, e.g., luminosity $L^* \sim \Lambda_w$. This is the only correct way to preserve the perceptual uniformity after the filtering by $\Lambda_w$. In the heuristic schemes, we added an heuristic correction to the simple multiplication by $\Lambda_w$, which yielded acceptable results (see the following subsection).

Heuristic Corrections

In this section, the $R_i$, $G_i$, and $B_i$ are further processed to produce a representation in which heuristic parameters $p_E$, $p_S$, $L_E$, $p_B$, $p_S$, and $p_C$ are applied to improve the color representation. We also treated the gamma correction exponent ($\gamma \equiv 2.2$) and the Steven’s law exponent ($\beta \equiv 0.4$) as heuristic parameters, i.e., we allowed them to be empirically adjusted. In our approach we tried to preserve the intuitive character of color representation and to remove some artifacts in a simple way using only a few free parameters. Regarding the nonuniformity in perceived distances, the most significant factor was improper implementation of the gamma correction. Parameters $\gamma$ and $p_S$ were specifically used for modification of color uniformity, and together with other heuristic parameters, produced a satisfactory level of uniformity. Any more elaborate scheme would require abandoning the goal of obtaining an intuitive association between typical color groups and directions [see (d) in Background]. This goal, for example, is not
achieved with schemes based on uniform color spaces. Also significant is the difference in perceived brightness between different hues. Perceived brightness for maximum value of pure red, green, and blue components, according to CIEL*a*b* color space for our computer monitor, are approximately 0.4, 0.6, and 0.8, respectively; and for yellow, cyan, and magenta, L* = 0.95, 0.87, and 0.7 (L* = 0 is black, L* = 1 is white). We found that the perceived brightness can be approximated according to L* = (0.3R + 0.59G + 0.11B)^0.4, where L*, R, G, and B are all mapped into the interval [0,1]. This approximation is valid for a wide range of colors for which L* > 0.1 and is derived from the CIE specifications for our computer screen. Knowing this, we derived a scaling factor Fl which, when applied to the CIE specifications for our computer screen. Knowing this, we derived a scaling factor Fl which, when applied to RGB components, brought all color compositions to the same brightness Le. The choice of Le had to be high enough to include a wide range of colors. We chose a moderately high value for Le in the range Le ∈ (0.6, 0.8). Since some color compositions cannot achieve this brightness, we shifted the blue component, and also for Le > 0.7 the red component, toward the white point. The higher the value of Le the larger the shift that was required. Uniform brightness is important when one wants to assign a normal to the plane of vectors for directional information and the degree of anisotropy simultaneously. In other situations, when the degree of anisotropy is less important one may use a larger range of colors. This is governed by the parameter pC. In the first step of implementation we shifted “blue” toward the white point according to:

\[
\begin{align*}
R_S &= \frac{F_m}{B_1 + (1-C_0)B_1} \\
G_S &= \frac{C_0}{B_1 + (1-C_0)G_1} \\
B_S &= B_1
\end{align*}
\]

In the same manner, we modified R_S, G_S, and B_S by shifting the red component but with a smaller shift (pR/4). From the R_S, G_S, and B_S values we obtained the digital values f_r, f_g, f_b to be stored in the frame buffer of the display device according to:

\[
\begin{align*}
f_r &= F_m \left( \frac{A_w}{L_f} \right)^{1/y} \\
f_g &= F_m \left( \frac{A_w}{L_f} \right)^{1/y} \\
f_b &= F_m \left( \frac{A_w}{L_f} \right)^{1/y}
\end{align*}
\]

where F_m is the largest integer that can be stored in each of the color channels (for 24-bit display, the channels have 8-bit registers and thus F_m = 255); A_w is the weighting factor based on the lattice index; and L_f is the scaling factor obtained by using the heuristic parameters L_E, pC and pE as follows:

\[
\begin{align*}
L_f &= p_c F_l + (1-p_c) L_m \\
F_l &= \min \left[ \frac{c_1 R_S + c_2 G_S + c_3 B_S}{L_f^{1/y}}, 1 \right] \\
L_m &= \max [R_g, G_b, B_b] \\
c_1 &= \frac{1}{3} - \frac{p_e}{2} \\
c_2 &= \frac{1}{3} + \frac{p_e}{4} \\
c_3 &= 1 - c_1 - c_2
\end{align*}
\]

where \( \beta = 0.4 \) can be treated as an heuristic parameter. In addition, equations for constants c_1, c_2, and c_3 can be varied if a non-CRT based display device is used. The form of Eq. [A6] is based on gamma correction and also includes the anisotropy filtering by A_w. If truncation is preferred instead, complete exclusion of voxels having anisotropy below a certain threshold A_w \_min with no weighting for the remaining voxels is obtained by setting the factor A_w \_min = 1 if A > A_w \_min, otherwise A_w \_min = 0.

**APPENDIX B: IMPLEMENTATION OF THE PREFERRED DIRECTION SCHEME**

Let us assume that the preferred direction is described by a vector \( \mathbf{v}_p = [x_p, y_p, z_p] \). We want to express the original polar and azimuthal angles \( \theta \) and \( \phi \) as angles \( \theta_p \) and \( \phi_p \), which are corresponding angles in the new coordinate system in which the z axis coincides with \( \mathbf{v}_p \). For an arbitrary vector \( \mathbf{v} \) we derive \( \theta_p \) and \( \phi_p \) according to:

\[
\begin{align*}
\theta_p &= \arccos \left( \frac{\mathbf{v} \cdot \mathbf{v}_p}{| \mathbf{v} |} \right) \\
\phi_p &= \begin{cases} 
(\arccos \left( \frac{\mathbf{v} \times \mathbf{v}_p}{| \mathbf{v} \times \mathbf{v}_p |} \right), & \mathbf{v} \cdot \mathbf{v}_p > 0, \mathbf{v}_p \neq \mathbf{v} \\
2\pi - \arccos \left( \frac{\mathbf{v} \times \mathbf{v}_p}{| \mathbf{v} \times \mathbf{v}_p |} \right), & \mathbf{v} \cdot \mathbf{v}_p < 0, \mathbf{v}_p \neq \mathbf{v} \\
0, & \mathbf{v} = \mathbf{v}_p
\end{cases}
\end{align*}
\]

where \( \mathbf{v}_p \) is the unit vector defining a normal to the plane which contains vector \( \mathbf{v}_p \) and defines a plane of vectors for which \( \phi_p = 0 \), or \( \phi_p = \pi \). We choose such a plane to be
Color Maps of Fiber Direction

defined by \( \mathbf{v}_p \) and the y axis, i.e.:

\[
\mathbf{n}_s = \begin{cases} 
\mathbf{v}_p \times \hat{\mathbf{y}} & \mathbf{v}_p \neq \hat{\mathbf{y}} \\
\frac{\mathbf{v}_p \times \hat{\mathbf{y}}}{\| \mathbf{v}_p \times \hat{\mathbf{y}} \|} & \mathbf{v}_p = \hat{\mathbf{y}} 
\end{cases}
\]

where \( \mathbf{x} \) and \( \mathbf{y} \) are the unit vectors defining the x and y axes. The polar and azimuthal angles in the rotated coordinate system, \( \theta_p \) and \( \phi_p \), are then used with HSV-based color as an heuristic approach with no symmetry, with the additional parameter \( \theta_c \) used to mask the discontinuity artifacts present in the image. For \( \theta_p \leq \theta_c \):

\[
H = (\phi_p - \phi_r + 2\pi) \bmod 2\pi
\]

\[
S = \frac{\sin (p_S p_t \theta_c)}{\sin (p_S \pi/2)}
\]

\[
V = 1
\]

\[
S_N = \frac{\pi}{2\theta_c}
\]

and for \( \theta_p > \theta_c \) we set \( V = 0 \). For \( \theta_p > \theta_c \) we also allow that both \( V \) and \( S \) are smoothly decreased according to \( S,V = [1-(\theta_p - \theta_c)/(\pi/2 - \theta_c)]D \) \( D > 2 \) controls the smoothness of transition. The latter case offers a visually more pleasing representation; however, if multiplicative scaling with a lattice index is used there is ambiguity between the structures with low anisotropy and those outside the cone defined by \( \theta_c \).

APPENDIX C: IMPLEMENTATION OF THE UNIFORM COLOR SPACE APPROACH

We omitted the many details of our implementations because of the unsatisfactory results that this approach produces. Therefore, we did not describe the transformations that map between RGB space of a given display device and CIEL\(*u\,*v\,*\) uniform color space (27). These require precise CIE specifications for a given display device to be known. Also, we found that all color parameters of the final color gamut of the uniform color spaces without significantly reducing the range of colors used. We have omitted the many details of our implementations because of the unsatisfactory results that this approach produces. The results for this approach require that all color parameters of the final printing device be known. Since these were unavailable, Fig. 5a is not truly a uniform color space implementation.

REFERENCES