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**Citation:** Huntjens, B., Asaria, T. S., Dhanani, S., Konstantakopoulou, E. and Ctori, I. (2014). Macular pigment spatial profiles in South Asian and white subjects. *Investigative Ophthalmology & Visual Science*, 55(3), pp. 1440-1446. doi: 10.1167/iovs.13-13204

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# Macular Pigment Spatial Profiles in South Asian and White Subjects

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Submitted: September 5, 2013  
Accepted: January 17, 2014

Citation: Huntjens B, Asaria TS, Dhanani S, Konstantakopoulou E, Ctori I. Macular pigment spatial profiles in South Asian and white subjects. *Invest Ophthalmol Vis Sci.* 2014;55:1440-1446. DOI:10.1167/iovs.13-13204

**PURPOSE.** Variability in central macular pigment optical density (MPOD) has been reported among healthy individuals. These variations seem to be related to risk factors of AMD, such as female sex, smoking, and ethnicity. This study investigates variations in the spatial profiles of MPOD among ethnicities.

**METHODS.** Using heterochromatic flicker photometry (HFP), MPOD was measured at seven retinal locations in 54 healthy, young South Asian and 19 white subjects of similar age. Macular pigment spatial profiles were classified as either typical exponential, atypical ring-like, or atypical central dip.

**RESULTS.** Central MPOD was significantly greater in South Asian ( $0.56 \pm 0.17$ ) compared with white subjects ( $0.45 \pm 0.18$ ;  $P = 0.015$ ). Integrated MPOD up to  $1.8^\circ$  (i.e., average MPOD [MPOD<sub>av(0-1.8)</sub>]) was also significantly increased in South Asian ( $0.34 \pm 0.09$ ) compared to white subjects ( $0.27 \pm 0.10$ ;  $P = 0.003$ ). Average MPOD(0-1.8) was significantly increased in all subjects presenting a ring-like profile ( $0.35 \pm 0.08$ ) or central dip profile ( $0.39 \pm 0.09$ ), compared with typical exponential profiles ( $0.28 \pm 0.09$ ;  $P < 0.0005$ ). We found a statistically significant association between ethnicity and spatial profile type ( $P = 0.008$ ), whereby an exponential profile was present in 79% of white compared with 41% of the South Asian subjects.

**CONCLUSIONS.** Central MPOD, MPOD<sub>av(0-1.8)</sub>, and the prevalence of atypical spatial profiles were significantly increased in South Asian compared with white subjects. Atypical profiles resulted in increased integrated MPOD up to  $1.8^\circ$ , and may therefore offer enhanced macular protection from harmful blue light.

**Keywords:** macular pigment optical density, ethnicity, heterochromatic flicker photometry, macular pigment spatial profiles

The spatial profile of macular pigment (MP) optical density has been shown to vary considerably among subjects. The optical density of MP, measured in log units, typically peaks centrally and declines sharply with eccentricity away from the foveola.<sup>1-3</sup> Central MP optical density (MPOD) has been reported to be lower with age,<sup>4</sup> smoking,<sup>5</sup> in the presence of inflammation promoting conditions (e.g., diabetes),<sup>6</sup> in females,<sup>7</sup> and in the presence of light iris color.<sup>8,9</sup> Previous studies described MP spatial profiles with either a single peak decaying exponentially,<sup>2,10,11</sup> a central dip (i.e., without a central peak),<sup>10,11</sup> or exhibiting a secondary peak up to  $2^\circ$  eccentricity, also referred to as a subpeak, shoulder, bimodal, or ring-like structure.<sup>2,10</sup> Using psychophysical heterochromatic flicker photometry (HFP), Hammond et al.<sup>2</sup> found that the MP distribution of 32 Caucasian subjects was best described by an exponential fit. However, the authors also discovered that approximately 40% of subjects presented secondary subpeaks (defined as increments greater than 0.05 optical density units from the exponential fit) at  $1^\circ$  and  $2^\circ$ . More recent studies have shown similar bimodal MP spatial profiles in a significant proportion of subjects.<sup>10,12-15</sup> The presence of a parafoveal ring was also shown in 20% to 50% of subjects when using objective autofluorescence imaging (AFI) techniques.<sup>10,15-17</sup> Moreover,

using AFI, the frequency of ring-like profiles was found to be significantly greater in females and in nonsmokers,<sup>15,16</sup> and in healthy subjects (43%) compared with patients with age-related maculopathy (23%).<sup>15</sup> Similar findings have also been demonstrated in ethnicities with a low prevalence of AMD, whereby 86% of African subjects presented with secondary peaks versus 68% non-Hispanic, white subjects.<sup>17</sup> However, it was also suggested that the lack of a central peak could possibly have an adverse effect on the protective role of MP in AMD, as the prevalence of a central dip has been found to increase with age and smoking in Caucasian subjects.<sup>11</sup>

Several studies have investigated ethnic differences in central MPOD.<sup>14,17-21</sup> White subjects presented significantly lower mean central MPOD compared to South Asian,<sup>18</sup> African,<sup>17,19</sup> and non-white subjects, including Asian, black, and Hispanic ethnicities.<sup>14</sup> However, the central MPOD of white subjects did not differ greatly compared with Chinese subjects.<sup>21</sup> Additionally, in a study where darker iris color was linked to increased average MPOD over the central  $1^\circ$  area, the results implied that central MPOD was not related to ethnicity. However, possible differences in MP density due to race were minimized as only a small percentage of non-Caucasian (Asian and African American) subjects were included.<sup>9</sup> Published data

on MPOD variations between South Asian (from India, Pakistan, and Bangladesh) and white subjects is limited.<sup>2,9,14,18</sup> Using the HFP technique, Howells et al.<sup>18</sup> reported a significantly increased mean central MPOD in South Asian ( $0.43 \pm 0.14$  log units) versus white subjects ( $0.33 \pm 0.13$  log units;  $P < 0.0005$ ), with increased MPOD in the Asian males compared with Asian females ( $P < 0.01$ ). This was not true for the white subjects; while the males presented with lower central MPOD, this was not statistically significant ( $P = 0.39$ ). Less is known about the ethnic differences in the distribution of MP away from the fovea. A study by Hammond et al.<sup>2</sup> found that MPOD distribution was not related to ethnicity.<sup>9</sup> Nolan et al.<sup>14</sup> also reported no association between the prevalence of a ring-like profile and ethnicity. However, both studies included limited numbers of non-White subjects (including South Asian) in comparison with the white group. To our knowledge, this is the first comparison study to investigate the prevalence of MP spatial profiles among South Asian and white subjects.

## METHODS

### Macular Pigment Measurements

Macular pigment optical density was assessed using a visual display unit based Macular Assessment Profile (MAP) test.<sup>22</sup> The MAP test uses HFP to measure MPOD at the center of the fovea ( $0^\circ$ ) and at six other retinal locations ( $0.8^\circ$ ,  $1.8^\circ$ ,  $2.8^\circ$ ,  $3.8^\circ$ ,  $6.8^\circ$ , and  $7.8^\circ$  eccentricity from the fovea). Like other tests employing HFP techniques, the MAP test is based on the spectrally selective properties of MP. Two beams of light are produced optically by the phosphors of the MAP test display unit. The test beam is composed of short wavelength (SW) blue light, peaking at approximately 450 nm, which is maximally absorbed in the central retina by MP. The reference beam is of a longer wavelength (LW) light that is not absorbed by the MP.<sup>23</sup> A notch filter is used in front of the test eye to increase the separation between the test and the reference beam. When the luminance of these wavelengths is not equal, a counter phased sinusoidal pattern is produced and the stimulus appears to flicker.<sup>1,24</sup> A larger difference in luminance yields a stronger sensation of flicker.

The center stimulus is a disc of  $0.36^\circ$  diameter. The peripheral stimuli are sectors of an annulus, which are presented concentric to the fovea. Both the angular subtense and the width of the peripheral stimuli increase with eccentricity<sup>22</sup> to ensure greater flicker sensitivity in the peripheral retina. Although the test supports any selected meridian, all the measurements reported in this study were performed with the stimulus centered along the horizontal meridian. In addition, a static mirror symmetric stimulus was presented at the corresponding location in the visual field to minimize the subject's tendency to saccade to the flickering peripheral target.

During the MAP test, the luminance of the test beam is altered until the perception of flicker is canceled or minimized. In order to ascertain the range of luminance for which the perception of flicker is absent, the MAP test calculates a low and a high threshold using a double reversal technique. The average of the low and high values is computed to give the luminance of the test beam required to cancel the reference beam (the flicker null point). The test is repeated in a random order eight times (four high and four low thresholds) at each eccentricity and the average is calculated to give the mean luminance of the SW test beam required to achieve the flicker null point. Macular pigment optical density is calculated by comparing the mean luminance adjustment of this SW light in

the central retina with a reference point in the peripheral retina using the equation

$$\text{MPOD} = \log_{10}(L_i/L_o), \quad (1)$$

where  $L_i$  is the mean luminance of the SW test beam at location  $i$  and  $L_o$  is the average of the test beam luminance of the  $6.8^\circ$  and  $7.8^\circ$  peripheral locations (where MP levels are thought to be negligible<sup>10</sup>).

### Study Protocol

The study took place at the Division of Optometry and Visual Science at City University London. Study data was collected from 54 South Asian and 19 white participants between May 2008 and November 2010. The average age of the South Asian participants was not statistically different from the average age of the white participants ( $P = 0.068$ ). Ethnicity was self-reported as white or South Asian (born in India, Pakistan, or Bangladesh, or born in the United Kingdom (UK) from Indian, Pakistani, or Bangladeshi parents; hereafter referred to as Asian). All participants had LogMAR visual acuity greater than 0.3 log units in the eye being tested. Exclusion criteria were ocular pathology, including inflammation, AMD or cataract, (self-reported) pregnancy, current use of carotenoid supplementation, and/or medication that may affect retinal function. Participants completed a lifestyle and health questionnaire, providing information about general and ocular health, use of medication, nutritional supplementation, and smoking history. Prior to using the MAP test, each participant was given a practice run of the  $0^\circ$ ,  $1.8^\circ$ , and  $2.8^\circ$  spatial locations. This provided a uniform introduction to the test and ensured complete dark adaptation.

### Classification of MP Spatial Profiles

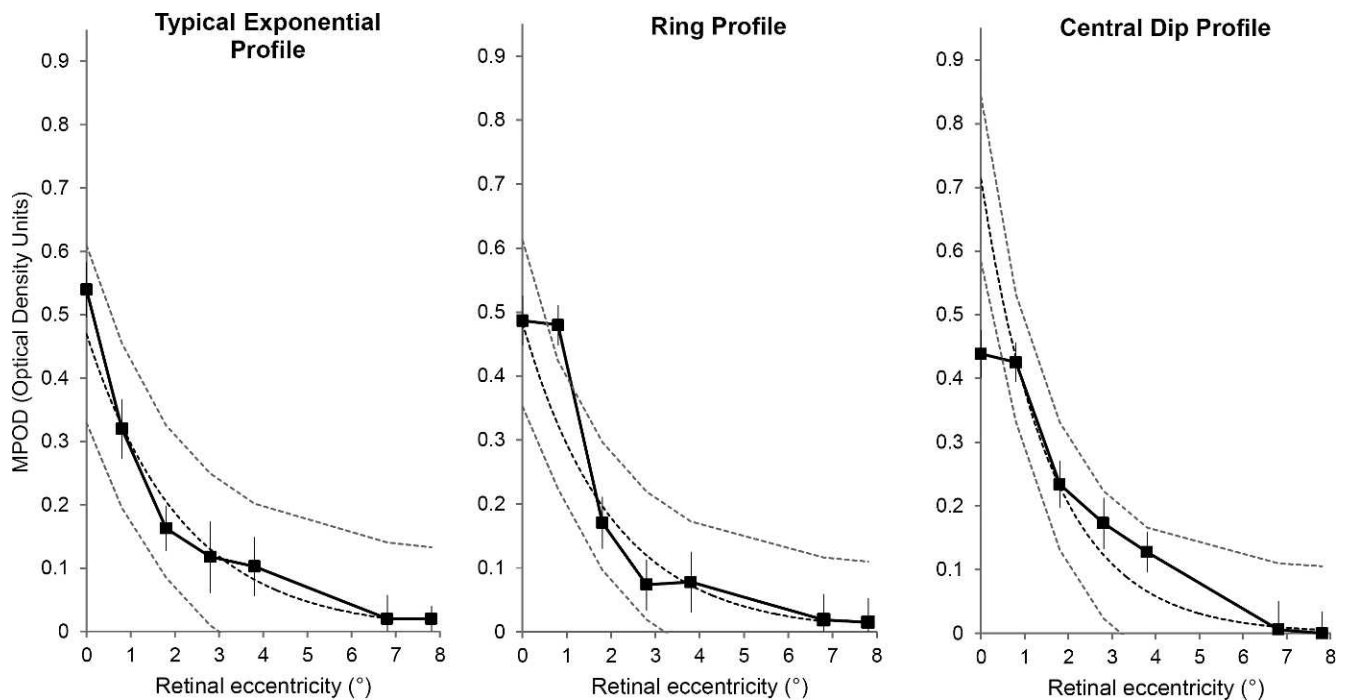
For each study participant, an exponential curve was fitted to the average absolute MPOD measurements at all retinal locations. The MP spatial profile presentation of each study participant was classified into typical exponential or atypical (nonexponential). The coefficient of repeatability (CoR; i.e., the average within-subject SD) was calculated from the eight repeated MPOD measurements at each eccentricity for both ethnicities. The exponential profile was classified by MPOD at  $0^\circ$ ,  $0.8^\circ$ , and  $1.8^\circ$  being within one CoR of the value predicted by the exponential curve. All others were assumed atypical. We subclassified our atypical group into ring-like and central dip profiles. Using the method described by Hammond et al.,<sup>2</sup> a positive deviation greater than the MAP test CoR from the exponential curve at  $0.8^\circ$  and/or  $1.8^\circ$  was classified as a ring-like profile. A negative deviation from the exponential profile greater than the MAP test CoR from the exponential curve at  $0^\circ$  was considered to be a central dip profile (Fig. 1).<sup>10</sup>

### Average Blue Light Transmittance ( $T_{av}$ ) and Average MPOD ( $\text{MPOD}_{av}$ )

At each eccentricity measured by the MAP test, the transmittance ( $T_i$ ) is a measure of the SW blue light-filtering capacity of the MP at location  $i$  and is given by

$$T_i = 10^{-\text{MPOD}_i}. \quad (2)$$

The value of  $T_i$  was plotted against retinal eccentricity, and the trapezium rule was used to calculate the area under the curve ( $T_{av}$ ), representing the integrated transmittance of the MP between eccentricities. Average blue light transmittance between  $0^\circ$  and  $1.8^\circ$  corresponding to a  $3.6^\circ$  diameter circular



**FIGURE 1.** Macular pigment optical density as a function of eccentricity for three participants: examples of exponential, ring, and central dip profiles. All three graphs include the mean absolute MPOD values  $\pm$  SD of eight measurements at each eccentricity. The *black dotted line* represents the exponential curve fitting to the mean absolute MPOD values. The *gray dashed lines* represent the MAP test measurement error according to the subject's ethnicity at each eccentricity from the exponential curve. Note the MPOD at  $0.8^\circ$  in the ring-like profile presents more than one coefficient of repeatability (CoR) above the expected exponential curve at  $0.8^\circ$ . The MPOD at  $0^\circ$  in the central dip profile shows more than one CoR below exponential curve.

aperture was calculated using the formula

$$T_{av(0-1.8)} = \frac{0.5(T_0 + T_{0.8})(\pi 0.8^2 - 0) + 0.5(T_{0.8} + T_{1.8})(\pi 1.8^2 - \pi 0.8^2)}{\pi 1.8^2}, \quad (3)$$

where  $T_0 = 10^{-\text{MPOD}}$  at  $0^\circ$ ,  $T_{0.8} = 10^{-\text{MPOD}}$  at  $0.8^\circ$ , and  $T_{1.8} = 10^{-\text{MPOD}}$  at  $1.8^\circ$ . The value of  $T_{av(0-1.8)}$  was used to calculate an average integrated MPOD between  $0^\circ$  and  $1.8^\circ$ :

$$\text{MPOD}_{av(0-1.8)} = -\log_{10} T_{av(0-1.8)}. \quad (4)$$

### Ethical Approval and Consent

Ethical approval was obtained from the Optometry Research and Ethics Committee at City University London, and written informed consent was obtained from all subjects, conforming to the tenets of the Declaration of Helsinki.

### Statistical Analysis

All statistical analyses were performed using SPSS version 19.0 for Windows (SPSS, Inc., Chicago, IL). Values in the text and tables are presented as the mean  $\pm$  SD. Kolmogorov-Smirnov tests revealed no significant deviation from a normal distribution for MPOD at different spatial locations. Independent Student's *t*-tests and one-way, between-groups ANOVA analyzed the differences between the ethnic groups, sex, and smoking status. The Pearson  $\chi^2$  test and Mann-Whitney *U* test were used to assess any difference between categories and groups that showed an abnormal distribution. Analysis of the variance was used to investigate any differences between the three different

distribution profiles of MP. Statistical significance was accepted at the 95% confidence level ( $P < 0.05$ ). Power statistics revealed that a sample size of 38, 19 subjects per group, was needed to detect a standardized difference of 0.91, using 80% power at 5% significance level.<sup>25</sup> This calculation was based on an estimated significant mean difference in MPOD of 0.1 with group SDs of 0.11 (based on the average MAP test coefficient of repeatability; Huntjens B, Asaria TS, Dhanani S, unpublished data, 2010).

### RESULTS

Demographics between the ethnic groups, and mean MPOD measured at each eccentricity are summarized in Table 1. There was a significant difference between the two ethnic groups: the Asian group included fewer current smokers compared with the white group ( $P = 0.039$ ). Age was not significantly correlated with central MPOD or any of the other spatial locations ( $r = -0.110$ ;  $P = 0.35$ ). Mean MPOD for individual eccentricities up to  $2^\circ$  showed a significant difference between the groups (Table 1). Average MPOD(0–1.8) (corresponding to integrated MPOD over the central  $3.6^\circ$  area) was significantly increased in Asian versus white subjects ( $t[71] = 3.07$ ;  $P = 0.003$ ). The significant difference in MPOD<sub>av</sub> up to  $1.8^\circ$  between ethnicities was maintained with smoking as a covariant ( $F[1,70] = 7.43$ ;  $P = 0.008$ ).

### Sex

When the group was considered as a whole ( $n = 73$ ), females had higher central MPOD values ( $0.55 \pm 0.19$ ) compared with males ( $0.50 \pm 0.16$ ); however, this difference was not statistically significant ( $t[71] = 1.25$ ;  $P = 0.22$ ). A one-way, between-groups analysis was conducted to explore the impact

TABLE 1. Demographics and MPOD Results for All Subjects and Separate Ethnic Backgrounds

	All	Asian	White	P Value
Number	73	54	19	
Age, y				
Mean $\pm$ SD	21.3 $\pm$ 3.2	20.9 $\pm$ 3.2	22.4 $\pm$ 2.8	0.068
Range	16–34	18–34	16–28	
Sex				
Male	24 (33%)	14 (26%)	10 (53%)	0.065
Female	49 (67%)	40 (74%)	9 (47%)	
Current smoker?				
Yes	8 (12%)	3 (6%)	5 (26%)	0.039*
No	65 (88%)	51 (94%)	14 (74%)	
Mean $\pm$ SD MPOD, log units				
MPOD 0°	0.53 $\pm$ 0.18	0.56 $\pm$ 0.17	0.45 $\pm$ 0.18	0.015*
MPOD 0.8°	0.44 $\pm$ 0.14	0.46 $\pm$ 0.13	0.37 $\pm$ 0.14	0.010*
MPOD 1.8°	0.19 $\pm$ 0.08	0.20 $\pm$ 0.09	0.14 $\pm$ 0.07	0.007*
MPODav(0–1.8)	0.32 $\pm$ 0.10	0.34 $\pm$ 0.09	0.27 $\pm$ 0.10	0.003*

Independent *t*-tests and  $\chi^2$  tests were conducted to determine statistically significant differences in MP measurements between Asian and white participants.

\*Statistical significance at the 0.05 level.

of sex on MPODav(0–1.8) between the ethnicities. Average MPOD(0–1.8) did not show a statistically significant difference between Asian males, Asian females, white males, and white females ( $F[3,69] = 2.25$ ;  $P = 0.06$ ).

### Smoking Status

Among all participants, central MPOD was increased in nonsmokers ( $0.54 \pm 0.18$ ) when compared with current smokers ( $0.47 \pm 0.17$ ); however, this difference was not statistically significant ( $t[71] = 1.01$ ;  $P = 0.32$ ). Additionally, a one-way, between-groups analysis did not show a significant difference in MPODav(0–1.8) between smoking and nonsmoking Asian and white subjects ( $F[3,69] = 2.69$ ;  $P = 0.053$ ).

### Spatial Profiles

When the group was considered as a whole, a typical exponential profile was seen in half of the group ( $n = 37$ ), while 36 participants showed a nonexponential (i.e., atypical) profile. Pearson's  $\chi^2$  test using the appropriate continuity correction indicated a statistically significant association between ethnicity and spatial profile type ( $\chi^2 [1, n = 73] = 6.75$ ,  $P = 0.009$ , Cramer's  $V = 0.335$ ). The results show that within ethnicities, 79% of white subjects presented an exponential profile in comparison to 41% of the Asian subjects (Fig. 2). In showing an atypical profile, 98% of participants were of Asian phenotype. We also observed an interesting relationship between the ethnicities and the three spatial profiles of MP as described in the Methods. When the group was considered as a whole, an exponential profile occurred in half the group, a ring in 30% of the group and the central dip profile was present in 19% of the subjects. Furthermore, 82% of subjects showing a ring and 100% of subjects showing a central dip profile were of Asian descent (Fig. 2). The Pearson's  $\chi^2$  test indicated a statistically significant association between ethnicity and spatial profile type ( $\chi^2 [2, n = 73] = 9.68$ ,  $P = 0.008$ , Cramer's  $V = 0.364$ ).

We explored the relationship between spatial profile type and MPOD at individual spatial locations up to 2° and MPODav(0–1.8) (Table 2). Average MPOD(0–1.8) was significantly increased in participants that showed an atypical when compared with an exponential spatial profile ( $t[71] = -4.56$ ;  $P < 0.0005$ ). This was also true for MPOD at 0.8° and MPOD at 1.8°, but not for central MPOD ( $t[67] = -1.35$ ;  $P = 0.19$ ). When the same analysis was conducted for each ethnicity, identical statistically significant results were found for the Asian subjects

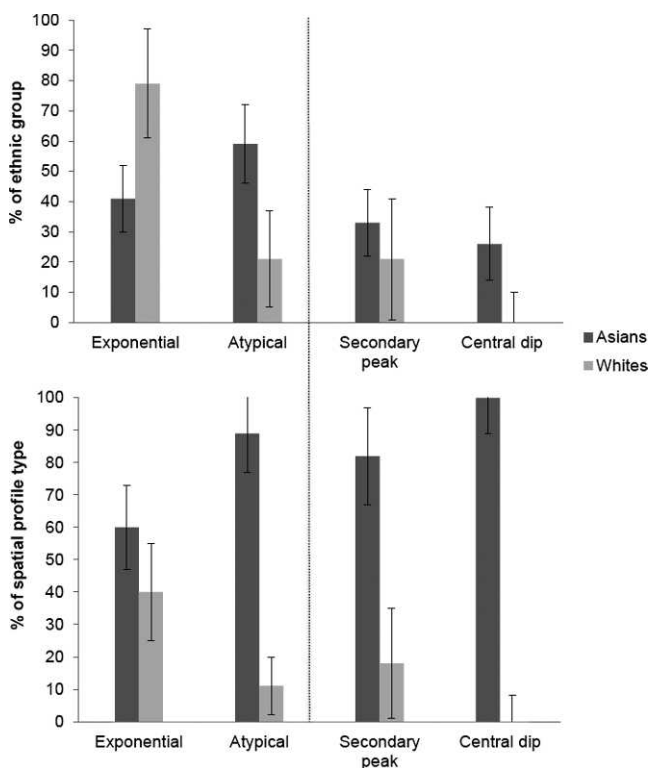


FIGURE 2. The frequency of spatial profile types. The upper graph shows typical exponential versus atypical MP spatial profiles as a percentage of each ethnic group. The lower graph shows the prevalence of ethnicity within each of the spatial profile groups. On the right side, the prevalence of individual atypical profiles (ring and central dip) is shown for both ethnic groups. Error bars represent the 95% confidence interval for proportions.

TABLE 2. Summary of MPOD Values Per Spatial Profile Type for All Participants

	Mean $\pm$ SD MPOD, Log Units				P Value
	Typical Exponential, <i>n</i> = 37	Atypical, <i>n</i> = 36	MP Ring, <i>n</i> = 22	Central Dip, <i>n</i> = 14	
MPOD 0°	0.51 $\pm$ 0.20	0.56 $\pm$ 0.15			0.19
MPOD 0.8°	0.36 $\pm$ 0.13	0.52 $\pm$ 0.11			<0.0005*
MPOD 1.8°	0.16 $\pm$ 0.06	0.22 $\pm$ 0.09			0.003*
MPODav(0-1.8)	0.28 $\pm$ 0.09	0.37 $\pm$ 0.08			<0.0005*
MPOD 0°	0.51 $\pm$ 0.20		0.57 $\pm$ 0.16	0.55 $\pm$ 0.14	0.43
MPOD 0.8°	0.36 $\pm$ 0.13†‡		0.52 $\pm$ 0.11	0.51 $\pm$ 0.11	<0.0005*
MPOD 1.8°	0.16 $\pm$ 0.06‡		0.19 $\pm$ 0.08‡	0.27 $\pm$ 0.10	<0.0005*
MPODav(0-1.8)	0.28 $\pm$ 0.09†‡		0.35 $\pm$ 0.08	0.39 $\pm$ 0.09	<0.0005*

\* Indicates statistical significance at the 0.05 level.

† Statistically significantly different from ring-like profile.

‡ Statistically significantly different from central dip profile.

but not for white subjects. Analysis of the variance showed statistically significant differences for all MPOD values (Table 2) when all three spatial profiles (exponential, ring, and central dip) were considered, with the exception of central MPOD ( $P = 0.43$ ). Post hoc analysis using the Tukey honest significant difference test indicated that the mean MPODav(0-1.8) for the exponential profile group ( $0.28 \pm 0.09$ ) was significantly decreased compared with the MP ring group ( $0.35 \pm 0.08$ ) and the central dip group ( $0.39 \pm 0.09$ ), but not between the two atypical profile groups. This was also true for MPOD at  $0.8^\circ$ . Interestingly, mean MPOD at  $1.8^\circ$  for the exponential group ( $0.16 \pm 0.06$ ) was not significantly different from the ring group ( $0.19 \pm 0.08$ ), but they were both significantly decreased from the subjects in the central dip group ( $0.27 \pm 0.10$ ;  $P < 0.0005$ ).

## DISCUSSION

Consistent with previous studies,<sup>18,26</sup> we found increased central MPOD in Asian ( $0.56 \pm 0.17$ ) versus white subjects ( $0.45 \pm 0.18$ ;  $t[71] = 2.50$ ;  $P = 0.015$ ). This is in agreement with the work of Howells et al.<sup>18</sup> where an average of  $0.43 \pm 0.14$  in 117 Asian and  $0.33 \pm 0.13$  in 52 white subjects was reported. Overall, their slightly lower average MPOD values compared with the present study are possibly due to the different HFP instruments used. However, the difference in central MPOD values between the ethnicities is similar between the studies. In contrast, Raman et al.<sup>26</sup> reported a mean central MPOD (at  $0.25^\circ$  retinal eccentricity) of  $0.63 \pm 0.16$  in 60 Asian subjects aged 20 to 29 years, and  $0.72 \pm 0.22$  in 60 Asian subjects aged 30 to 39 years. These values are higher when compared with our results, which again may be due to the different HFP instruments. Furthermore, the Asian subjects were of South Indian origin living in India (Mumbai); however, similar to Howell's study,<sup>19</sup> the Asian subjects included in our study were of Indian, Pakistani, and Bangladeshi descent, the majority born and living in the UK (78%; 42 out of 54 Asian subjects). The country of origin and residence may be significant because of differences in diet. The traditional south Asian diet typically consisting of a diet rich in carotenoids may be altered after migration, particularly in the young or second generation Asians<sup>27</sup>; this may contribute to the lower MPOD levels found in our group.

The integrated transmittance of the MP between eccentricities was used to calculate the average MPOD up to  $1.8^\circ$ . Similar to central MPOD, mean MPODav(0-1.8) was significantly increased in Asian ( $0.34 \pm 0.09$ ) compared with white subjects

( $0.27 \pm 0.10$ ;  $t[71] = 3.07$ ;  $P = 0.003$ ). Lower central MPOD has been associated with factors that may increase the risk of AMD, such as female sex<sup>4,7,20,21,28,29</sup> and smoking.<sup>5,28</sup> The relationship between spatial profiles and ethnicities, including covariates such as sex and smoking status, were difficult to establish in the present study due to the small sample size of each subgroup. Nonetheless, we did not find a sex association with MPOD, with central MPOD values of  $0.55 \pm 0.19$  for the females compared with  $0.50 \pm 0.16$  for the males ( $P = 0.22$ ).

When the groups were analyzed by ethnicity, a similar trend was found for both Asian and white participants. Previous studies of Asian subjects with a similar age range to our study have reported that males have higher mean MPOD than females.<sup>18,26</sup> One study found this to be statistically significant.<sup>18</sup> The difference between MPODav(0-1.8) in nonsmokers ( $0.33 \pm 0.09$ ) compared with smokers ( $0.27 \pm 0.11$ ) did not reach statistical significance ( $P = 0.15$ ). We note that the lack of a difference may be due to the small sample of smoking subjects (8 out of 73 subjects) and the short smoking history.

Our data suggest that atypical profiles (i.e., ring and central dip) occur more frequently in Asian compared with white subjects ( $P = 0.009$ ). The average integrated MPOD up to  $1.8^\circ$  was significantly increased in Asian subjects presenting with atypical ( $0.38 \pm 0.08$ ) versus exponential profiles ( $0.29 \pm 0.10$ ;  $t[52] = -3.86$ ;  $P < 0.0005$ ). In white subjects, this finding was not significant ( $0.30 \pm 0.07$  and  $0.26 \pm 0.10$ , respectively;  $t[17] = -0.85$ ;  $P = 0.41$ ). Therefore, it seems that an atypical spatial profile is a representative characteristic of the Asian group, and indeed may be considered typical in this ethnic group. Since there was no significant difference between central MPOD in Asian ( $t[35] = -0.71$ ;  $P = 0.48$ ) or in white subjects presenting with an atypical profile compared with an exponential profile ( $t[17] = 0.26$ ;  $P = 0.80$ ), our results suggest that, compared with an individual MPOD measurement at a single retinal spatial location or an average of MPOD measurements at several retinal spatial locations, MPODav(0-1.8) provides a better representation of the amount of MP present. Although some of the subjects show a sizable decrease in MPOD at the fovea, many others do not. In spite of large variability in MPOD caused by averaging MPOD over the area of the stimulus and the variability in fixation accuracy during the HFP test, the results using a small central target (i.e.,  $0.36^\circ$  diameter) suggest that a ring-like profile is possible. However, the main conclusion of the study based on the measured differences in short wavelength transmittance over the centre  $3.6^\circ$  has become more significant by analyzing the results in terms of area weighted central transmittance.

This is the first comparative study to investigate MP spatial profiles in Asian and white subjects. Several studies have reported on the different spatial distributions of MP; however, there is little consensus on the definition of an atypical profile. Additionally, there are various methodologies used to measure MP density and results are consequently not always interchangeable. The spatial profile of MP is normally described as following an exponential decline, although 20% to 50% of the population in studies where MP is measured by HFP and objective imaging techniques have shown a deviation from the exponential curve at 0° or at a location away from the central fovea.<sup>10,15,16</sup> The lack of spatial resolution in the measurement of central MPOD can be largely attributed to the size of the central target, as well as the subject's ability to maintain steady fixation. In comparison with other HFP techniques, the MAP test aims to minimize this effect by employing a very small central (0.36°) and static peripheral stimuli. A nonexponential spatial profile was found in 21% (4 out of 19) of white subjects and 59% (32 out of 54) of Asian subjects. Atypical profiles have been previously defined as those not exhibiting a typical exponential profile, but showing either a annulus of higher MP or ring, where the central peak is surrounded by a ring of increased density,<sup>15</sup> or a central dip (i.e., MPOD at 0.25° not visually exceeding MPOD at 0.5°,<sup>13</sup> or MPOD at 0.25° not exceeding MPOD at 0.5° by more than 0.04 optical density units<sup>30</sup>). The presence of a MP ring has been found significantly increased in ethnicities with low AMD prevalence,<sup>17</sup> suggesting it may enhance the MP's protective role. Wolf-Schnurrbusch et al.<sup>17</sup> showed significantly increased frequency of a parafoveal ring ( $P < 0.0001$ ) and central MPOD ( $P < 0.0001$ ) in African subjects, when compared with non-Hispanic white subjects. In contrast, since increased prevalence of a central dip was found to be associated with increased age and smoking, it was proposed that a central dip decreased the protective role of MP.<sup>11</sup>

Interestingly, when we considered the atypical spatial profiles in all participants, we found that MPOD values at 0.8° and 1.8° and MPOD<sub>av(0-1.8)</sub> were increased in the profiles showing a ring or central dip, compared with the exponential profile. Table 2 shows that this was statistically significant, with the exception of central MPOD. There was no difference in central MPOD between the exponential, ring and surprisingly, the central dip profile groups. Unexpectedly, the mean MPOD at 1.8° for the group presenting a ring was not significantly different from the exponential group, but was significantly lower than for the central dip group ( $P < 0.0005$ ). These results show that the central dip profile has more MPOD at or close to the location where the MP ring profile shows its additional peak. It seems that a central dip has not lost its peak, but possibly broadened its lateral distribution. We, therefore, propose that the presence of a central dip profile may actually offer increased integrated MPOD up to 1.8°, and therefore increased macular protection from harmful blue light. Moreover, our data suggest that there may be a disparity in the occurrence of MP spatial profiles amongst ethnicities. Not only were atypical spatial profiles more frequently present in Asian subjects ( $P = 0.008$ ), but also the central dip was entirely absent in white subjects. This implies that there may be need for subclassification of MP spatial profiles other than typical (i.e., exponential) versus atypical, as previously suggested by Berendschot and van Norren.<sup>10</sup> Additionally, we propose using exponential versus nonexponential profile types, since atypical profiles for some ethnicities may represent typical characteristics for that group.

Considering previous reports of dietary differences between ethnicities,<sup>31,32</sup> our data support the hypothesis that the central dip could be the result of a high conversion of lutein to meso-zeaxanthin<sup>33,34</sup> resulting in an increased MPOD at the

0.8° and 1.8° locations. Additionally, there is supporting evidence that lutein and zeaxanthin supplementation increases MPOD in the human foveal and parafoveal areas.<sup>35-37</sup> The distribution of zeaxanthin (centrally) and lutein (more peripherally) within the macula may suggest that an exponential or atypical ring profile represent a relative enrichment of zeaxanthin, while an atypical central dip profile represents a relative enrichment of lutein. However, Zeimer et al.<sup>38</sup> suggested that lutein and zeaxanthin supplementation in AMD and control subjects might amplify, not create, atypical MP spatial profiles. A limitation of our study was that we did not measure lutein and zeaxanthin dietary intake. Neither could we relate these differences in spatial profiles to the iris color, or family history of AMD, since we did not collect this data. While not controlled for in our study, iris color and dietary intake of carotenoids may be the largest source of variation between our two groups. Nonetheless, our results have shown an uneven distribution of MP spatial profile types between white and Asian subjects, which confirms the need for wider-scale studies, including other ethnic phenotypes, iris color, and dietary intake of carotenoids.

## CONCLUSIONS

This is the first study to investigate the prevalence of different MP spatial distributions for Asian and white subjects. Our results show that central MPOD was significantly increased in our 54 Asian subjects, compared with 19 white subjects of similar age. We classified spatial distributions of macular pigment into typical exponential and atypical (nonexponential) profiles. Atypical profiles were significantly more prevalent in Asian compared with white subjects. Additionally, we noted that ring and central dip spatial profiles varied between the ethnicities, whereby the prevalence of central dip was significantly increased in Asian group. Additionally, integrated MPOD up to 1.8° was significantly increased in a central dip compared with an exponential profile. This suggests that, similar to a MP ring, a central dip represents enhanced retinal protection from harmful blue light.

## Acknowledgments

The authors thank John Barbur for use of MAP test, and greatly appreciate the helpful comments from John Lawrenson and John Barbur during the writing of this manuscript.

Disclosure: **B. Huntjens**, None; **T.S. Asaria**, None; **S. Dhanani**, None; **E. Konstantakopoulou**, None; **I. Ctori**, None

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