The cone dystrophies comprise a heterogeneous group of disorders characterised by visual loss, abnormalities of colour vision, central scotomata, and a variable degree of nystagmus and photophobia. They may be stationary or progressive. The stationary cone dystrophies are better described as cone dysfunction syndromes since a dystrophy often describes a progressive process. These different syndromes encompass a wide range of clinical and psychophysical findings. The aim is to review current knowledge relating to the cone dysfunction syndromes, with discussion of the various phenotypes, the currently mapped genes, and genotype-phenotype relations. The cone dysfunction syndromes that will be discussed are complete and incomplete achromatopsia, oligocone trichromacy, cone monochromatism, blue cone monochromatism, and Bornholm eye disease. Disorders with a progressive cone dystrophy phenotype will not be discussed.

The cone dystrophies are characterised by bilateral visual loss, colour vision abnormalities, central scotomata, variable degrees of nystagmus and photophobia, together with electrophysiological or psychophysical evidence of abnormal cone function. There is considerable clinical and genetic heterogeneity; cone dystrophies showing autosomal dominant, autosomal recessive, and X linked recessive inheritance have all been reported. These disorders may be stationary or progressive. The stationary subtypes are congenital with normal rod function, whereas in progressive cone dystrophies, onset is usually in childhood or early adult life and patients often develop rod photoreceptor dysfunction in later life. The stationary disorders are better described as cone dysfunction syndromes. In this review we will describe the various phenotypes and disease causing genes that have been recently identified in this group of disorders (table 1). We will not consider the various forms of colour vision deficiency; the molecular genetic basis of these disorders has now been well characterised and several reviews have been published on the subject.1 2

**COMPLETE ACHROMATOPSIA**

Complete achromatopsia, typical achromatopsia or rod monochromatism, is a stationary disorder in which there is an absence of functioning cone photoreceptors in the retina.3 4 It is uncommon with an incidence of about 1 in 30 000.1 4 Affected individuals usually present in infancy with pendular nystagmus, poor visual acuity, and photophobia. A hypermetropic refractive error is common and it is often found that the nystagmus wanes with time.5 Fundal examination is usually normal; however, infrequently, central or mid-peripheral retinal pigment epithelial abnormalities are present. Electroretinography (ERG) reveals absent cone responses and normal rod responses.6 Affected individuals usually achieve a visual acuity of 6/60, have absent colour vision, and have normal rod function but absent cone function on psychophysical testing.7

Achromatopsia is recessively inherited and genetically heterogeneous. To date, three achromatopsia genes have been identified, CNGA3, CNGB3, and GNAT2; all three genes will be described in detail in the discussion that follows. The first molecular genetic report of achromatopsia was a cytogenetic analysis of a 20 year old woman with achromatopsia and multiple developmental abnormalities.8 Maternal isodisomy of chromosome 14 was demonstrated (both copies of chromosome 14 were of maternal origin). However, there has been no subsequent confirmation of a locus on chromosome 14. In 1997 a genome-wide search for linkage was performed in a consanguineous Jewish kindred, establishing linkage to a 14 cM region on 2q11.9 This disease interval was further refined to a 3 cM region in 1998 in a study of eight families of different ethnic and racial origins, and CNGA3 was identified as a candidate gene within this interval.10 CNGA3 encodes the α-subunit of the cGMP gated (CNG) cation channel in human cone photoreceptors, the final critical effector in the phototransduction cascade. In the dark, cGMP levels are high in cone photoreceptors, therefore enabling cGMP to bind to the α and β-subunits of CNG channels, resulting in them adopting an open conformation and permitting an influx of cations, with consequent cone depolarisation. However, in light conditions, activated photopigment initiates a cascade culminating in increased cGMP phosphodiesterase activity, thereby lowering the concentration of cGMP in the photoreceptor which results in closure of CNG cation channels and consequent cone hyperpolarisation.11

Missense mutations in highly conserved residues of CNGA3 were initially described in five families with complete achromatopsia from Germany, Norway, and the United States.12 Since then more recent studies have revealed more than 50 disease causing mutations in CNGA3.13 14 Mutations have been identified throughout the CNGA3 protein, including the five transmembrane domains, the pore region,
### Summary of the cone dysfunction syndromes

<table>
<thead>
<tr>
<th>Gene dysfunction syndrome</th>
<th>Mutated gene(s) or chromosome locus</th>
<th>Mode of inheritance</th>
<th>Visual acuity</th>
<th>Refractive error</th>
<th>Nystagmus</th>
<th>Colour vision</th>
<th>Fundi</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cone dysfunction syndrome</td>
<td>CNAG3</td>
<td>Autosomal recessive</td>
<td>6/24–6/60</td>
<td>Often hypermetropia</td>
<td>Usually absent</td>
<td>Usually normal</td>
<td>Normal</td>
</tr>
<tr>
<td>Cone monochromatism</td>
<td>CNAG3</td>
<td>X linked</td>
<td>6/6</td>
<td>Normal</td>
<td>Absent or blue cone monochromatism</td>
<td>X linked</td>
<td></td>
</tr>
<tr>
<td>Blue cone monochromatism</td>
<td>CNAG3</td>
<td>X linked</td>
<td>6/9–6/18</td>
<td>Moderate to high myopia</td>
<td>Absent</td>
<td>Normal</td>
<td></td>
</tr>
</tbody>
</table>

*Note:* The second gene identified in patients with achromatopsia is the null phenotype of CNAG3, which is the null phenotype of CNAG3. A similar study by Kohl et al identified six mutations in CNAG3; three were novel—Arg203stop, Glu366stop, and a putative splice site defect. Rojas et al have since identified Asp149fs in a consanguineous Chilean family. The most frequent mutation of CNAG3 identified to date is the 1 base pair frameshift deletion, 1148delC (Thr383fs), which accounts for up to 84% of CNAG3 mutant disease chromosomes.

Currently there is far greater allelic heterogeneity of CNAG3 mutants (over 50 mutations described) when compared to CNAG3 (~7). It is known that CNAG3 subunits can form functional homomeric channels when expressed alone, whereas CNAG3 subunits alone do not appear to form functional channels. In our opinion it is therefore plausible that some CNAG3 null mutations are not detected since sufficient channel function is possible solely with normal CNAG3 subunits, leading to a relatively normal phenotype.

These studies of CNAG3 and CNAG3 have demonstrated that both the α and β-subunits of the CNAG cation channel are essential for phototransduction in all three classes of cones. The majority of CNAG3 mutations identified to date are missense mutations, indicating that there is little tolerance for substitutions with respect to functional and structural integrity of the channel polypeptide. This notion is supported by the high degree of evolutionary conservation among CNAG channel α-subunits. In contrast, the majority of CNAG3 alterations are nonsense mutations. It is currently proposed that approximately 25% of achromatopsia results from mutations of CNAG3 and 40–50% from mutations of CNAG3. Therefore, while mutations in the cone channel subunit genes, CNAG3 and CNAG3, account for the majority of achromatopsia, there is a significant proportion of patients for whom neither CNAG3 nor CNAG3 mutations can be found (~30%). The phenotype associated with mutations in these two channel protein genes appears to be in keeping with previous clinical descriptions of achromatopsia.

It is of interest that missense mutations in CNAG3 have also been reported in two individuals with cone-rod dystrophy and in a single individual with a progressive cone dystrophy phenotype. Possible reasons for a progressive phenotype in these individuals may include the combination of missense mutations present in these three subjects; some amino acid substitutions may be more deleterious to channel function than others.
phenotypic influences include the presence of other modifier
genes or environmental effects.

Cone degeneration (cd) is an autosomal recessive canine
disease that occurs naturally in the Alaskan Malamute and
German shorthaired pointer breeds and is phenotypically
similar to human achromatopsia. 40–42 Canine CNGB3 mutations
have recently been identified in both of these breeds, thereby
establishing these cd affected dogs as the only naturally
occurring large animal model of human achromatopsia, and
therefore providing a valuable system for exploring disease
mechanisms and evaluating potential genetic therapeutic
intervention in human achromatopsia. 40–42 GNAT2, located at 1p13, is the third gene to be implicated
in achromatopsia. 40–42 GNAT2 codes for the α-subunit of cone
specific transducin. In cone cells, light activated photopig-
ment interacts with transducin, a three subunit guanine
nucleotide binding protein, stimulating the exchange of
bound GDP for GTP. The cone α-transducin subunit, which is
bound to GTP, is then released from its β and γ-subunits and activates
CGMP phosphodiesterase by removing the inhibi-
tory γ-subunits from the active site of this enzyme. CGMP
phosphodiesterase lowers the concentration of CGMP in the
photoreceptor which results in closure of CGMP gated cation
channels. 40–42 All the GNAT2 mutations identified to date result
in premature translation termination and in protein truncation
at the carboxy terminus. 40–42 However, mutations in this
gene are thought to be responsible for less than 2% of
patients affected with this disorder, 40–42 suggesting the presence
of further genetic heterogeneity in achromatopsia.

We have recently undertaken a detailed description of the
phenotype associated with GNAT2 inactivation in a large
consanguineous Pakistani family. 42 The phenotype is char-
acterised by mild photophobia, nystagmus, abnormal colour
vision, and poor visual acuity (6/36 to counting fingers). On
detailed colour vision testing, residual colour discrimination
was detected in three individuals. ERGs revealed absent cone
responses, with normal rod specific ERGs. We were able to
record S cone ERG responses in all patients. In two older
subjects, a worsening of visual acuity with age has been
documented, although we have no definite evidence of
progressive deterioration in retinal function. The residual S
cone function detected in this GNAT2 associated phenotype is
intriguing. The evidence that GNAT2 is expressed in all three
cone types comes from the immunohistochemical demon-
stration that an antibody raised against cone α-transducin
peptides cross reacts with all three classes of cone photo-
receptor in the human retina. 42 This does not however
definitively rule out the possibility that S cones may express
an alternative form of α-transducin, since identical epitopes
may be present on both forms. It may also be significant that
Southern blot analysis of human genomic DNA indicated
that there may be more than one cone α-transducin gene. 42
Therefore, it remains a possibility that GNAT2 is not
expressed in S cones, and that the residual S cone function
detected in our family arises from the use of another distinct
form of α-transducin. The residual tritan colour discrimina-
tion detected may be accounted for by a comparison between
quantum catches in the remaining functional S cones and rod
photoreceptors, in the manner proposed to underlie colour
discrimination detected in blue cone monochromatism. 42–46

The three genes described to date associated with
achromatopsia, CNGA3, CNGB3 and GNAT2, encode proteins
in the cone phototransduction cascade. It is therefore
reasonable to propose that further cone specific intermediates
involved in phototransduction represent good candidates.
These include the genes encoding the cone specific β and γ-
transducin subunits and cone phosphodiesterase. It is of note
that immunological studies of the canine cd affected retina
have demonstrated a specific absence or delocalisation of
β and γ cone specific transducin subunits from the outer
segments of pre-degenerate cone photoreceptors. However,
genomes for both subunit transducin subunits have been excluded as canine
cd genes. 5, 6

INCOMPLETE ACHROMATOPSIS

Previously, before the underlying pathogenesis of blue cone
monochromatism (BCM) had been identified, BCM was
known as X linked incomplete or atypical achromatopsia.
However, the term incomplete/atypical achromatopsia is best
reserved for the description of individuals with autosomal
recessive disease where the phenotype is a variant of
complete achromatopsia. Individuals with incomplete achro-
matopsia (atypical achromatopsia) retain residual colour
vision and have mildly better visual acuity (6/24–6/60) than
those with complete achromatopsia. 5, 6 In all other respects,
the phenotype of these two conditions is indistinguishable.
Three subtypes of incomplete achromatopsia have been
demonstrated via colour matching experiments: 31–35:

- colour matches are governed by rods and M cones
  (incomplete achromatopsia with protan luminosity) 31;
- colour matches are governed by L and M cones;
- colour matches mediated by rods, L cones, and S cones
  (incomplete achromatopsia with deutan luminosity). 31–34

As in the complete form, mutations in CNGA3, the gene
coding the α-subunit of the CGMP gated cation channel in
cones, have been identified in individuals with incomplete
achromatopsia. 34 The psychophysical data provided in this
study 31–34 are inadequate to be able to classify these individuals
into the three colour matching subtypes described above. 31–34
The 19 mutations identified were all missense mutations,
located throughout the channel polypeptide including the
transmembrane domains, ion pore, and CGMP binding
region. However, only three of these missense mutations,
Arg427Cys, Arg563His, and Thr565Met, were found exclu-
sively in patients with incomplete achromatopsia. 31–34 Therefore
in the majority of cases of incomplete achromatopsia, factors
other than the specific causative mutation, such as modifier
genes, or environmental influences, may dictate the pheno-
type. The missense variants identified in incomplete achro-
matopsia must be compatible with residual channel function
since the phenotype is milder than in complete achro-
matopsia. Mutations in CNGB3 or GNAT2 have not been reported in
association with incomplete achromatopsia, despite mutant
CNGB3 alleles being identified twice as common as CNGA3
variants as the cause of complete achromatopsia. However all
GNAT2 mutations to date, and the vast majority of CNGB3
mutations, result in premature termination of translation, and
thereby truncated and most probably non-functional photo-
transduction proteins. Therefore an incomplete achromatop-
sia phenotype is unlikely to be compatible with these
genotypes which are predicted to encode mutant products
lacking any residual function.

OLIGOCONE TRICHRROMACY

Oligocone trichromacy is a rare cone dysfunction syndrome,
which is characterised by reduced visual acuity, mild
photophobia, normal fundi, reduced amplitude of the cone
electroretinogram, but with colour vision within normal
limits. The disorder was first described by Van Lith in 1973. 35 Since then Keunen et al have described a further
four patients, 35 while Neuhann et al, and, more recently, Ehlich
et al have each reported a single case. 36–38 The two cases reported
by Van Lith and Ehlich both had pendular nystagmus.
It has been proposed that these patients might have a
reduced number of normal functioning cones (oligocone


syndrome) with preservation of the three cone types in the normal proportions, thereby permitting trichromacy. Keunen et al tested this hypothesis by screening foveal cone photopigment density. A reduced density difference of the foveal cone photopigment with a normal time constant of photopigment regeneration was found in all patients. Colour matching and increment threshold spectral sensitivity were normal. This provided evidence for the hypothesis of a reduced number of foveal cones (decreased density differences) with otherwise normal functioning residual cones.

We have recently detailed the phenotype of six patients with oligocone trichromacy. All six affected patients had a history of reduced visual acuity from infancy (6/12 to 6/24). They complained of very mild photophobia, but were not aware of any colour vision deficiency. They had no nystagmus and fundi were normal. On examination, all patients were found to have good colour vision. The various colour vision tests either revealed completely normal colour vision or slightly elevated discrimination thresholds. Anomaloscope revealed matching ranges within normal limits, indicating the presence of long and middle wave cones of normal spectral sensitivity at the macula, while the absence of pseudoprotanomaly suggests that photopigment is present at normal optical densities in individual cone photoreceptors. The slightly elevated discrimination thresholds that were detected are compatible with a reduction in cone numbers. The cone ERG findings in our patients were poorly concordant, but could broadly be divided into two classes. In the first group (five individuals) cone responses were absent or markedly reduced. In the second group (one individual), cone b-waves were more markedly reduced than a-waves, implying a predominantly inner retinal abnormality in the cone system. These electrophysiological data suggest that there may be more than one disease mechanism and therefore more than one disease causing gene.

Oligocone trichromacy is likely to be inherited as an autosomal recessive trait. The molecular genetic basis of the disorder is unknown. Genes involved in retinal photoreceptor differentiation, when cone numbers are being determined, may represent good candidate genes.

**CONE MONOCHROMATISM**

Monochromatism is diagnosed by a patient's ability to match any two colours merely by adjusting their radiance when all other cues are absent. In rod monochromatism there is an absence of functioning cone photoreceptors with visual perception depending almost exclusively on rods. The rod monochromat therefore has markedly reduced visual acuity and total colour blindness.

Cone monochromatism is another rare form of congenital colour blindness, in which visual acuity is normal. The incidence of cone monochromatism is estimated at one in 100 million. Unlike rod monochromatism, cone monochromatism has never been noted in more than one family member. The colour vision defect may be incomplete for certain colours and may vary both with the size of the field viewed and the level of luminance. A normal ERG is present in this disorder thereby supporting the notion of abnormal processing central to the retinal photoreceptors and bipolar cells. This notion was first proposed following the demonstration of red and green sensitive pigments at the fovea in cone monochromats and the ability of such patients to use ocular chromatic aberration as a cue for altering accommodation. In addition, Gibson has been able to demonstrate the presence of three mechanism sensitivity curves for the cone monochromat that are similar to those found in normal individuals, representing evidence of colour mediating mechanisms in the cone monochromat, and thereby providing further evidence for a post-receptoral defect in this disorder.

**BLUE CONE MONOCHROMATISM**

Blue cone monochromatism (BCM), previously also known as X linked incomplete achromatopsia, affects fewer than 1 in 100 000 individuals, and is characterised by absence of L and M cone function. Thus, the blue cone monochromat possesses rod vision and a normal short wavelength sensitive cone mechanism.

As in rod monochromacy, BCM typically presents in infancy with reduced visual acuity, pendular nystagmus, photophobia and normal fundi. The nystagmus often wanes with time. Visual acuity is of the order of 6/24 to 6/60. Eccentric fixation may be present and myopia is a common finding. BCM is distinguished from rod monochromatism (RM) via psychophysical and electrophysiological testing. The photopic ERG is profoundly reduced in both, although the S cone ERG is well preserved in BCM. Classification can also be aided by family history, because BCM is inherited as an X linked recessive trait, whereas both subtypes of rod monochromacy show autosomal recessive inheritance.

Rod monochromats cannot make colour judgments, but rather will use brightness cues to differentiate between colours. This contrasts with blue cone monochromats who do have access to colour discrimination, though this does depend upon the luminance of the task: at mesopic levels, they have rudimentary dichromatic colour discrimination based upon a comparison of the quantum catches obtained by the rods and the S cones (blue cones). Colour discrimination is reported to deteriorate with increasing luminance. Therefore blue cone monochromats may be distinguished from rod monochromats by means of colour vision testing: blue cone monochromats are reported to display fewer errors along the vertical axis in the Farnsworth 100 Hue test (fewer tritan errors), and they may also display protan-like ordering patterns on the Farnsworth D-15. In addition, the Berson plates have been claimed to provide a good separation of blue cone monochromats from rod monochromats. Therefore, in order to clinically distinguish RM and BCM one needs to use colour vision tests that probe the tritan axis of colour as well as the deutan and protan, since the presence of residual tritan discrimination suggests BCM.

In order to derive colour vision, the normal human visual system compares the rate of quantum catches in three classes of cones: the longest wavelength sensitive (L) cone; the short (S) wavelength sensitive medium wave (M) cone; and the short wavelength sensitive, and long (L) wavelength sensitive cones are maximally sensitive to light at 430 nm, 535 nm, and 565 nm, respectively. Whereas the L (red) and M (green) pigment genes are located on the X chromosome, the S cone (blue) pigment is encoded by a gene located on chromosome 7. The wild type arrangement of the L and M opsin genes consists of a head to tail tandem array of two or more repeat units of 39 kb on chromosome Xq28 that are 98% identical at the DNA level. The highly homologous L and M opsin genes are as a consequence predisposed to unequal intergenic and intragenic recombination. Transcriptional regulation of the L and M visual pigment genes is controlled by an upstream locus control region (LCR). Mutations in the L and M pigment gene array result in the lack of functional L and M pigments, and thus inactivate the corresponding cones, have been identified in the majority of BCM cases studied.

Mutation analyses introduced by Nathans and collaborators have proved highly efficient at establishing the molecular basis for BCM. The mutations in the L and M pigment gene array causing BCM fall into two classes. In the first class, a normal L and M pigment gene array is inactivated by...
of these multistep pathways produce visual pigment genes. The evidence thus far shows that many monochromat genotypes comprise a heterogeneous group of BCM. The data suggest that 40% of blue cone monochromat BCM where exon 4 of an isolated red pigment gene had been deleted.\(^2\)BCM is generally accepted to be a stationary disorder, although Fleischman and O’Donnell reported one BCM family with macular atrophy and noted a slight deterioration of visual acuity and colour vision during a 12 year follow up period, as well as foveal pigmentary changes.\(^3\) There are two further reports of individual displaying a progressive retinal degeneration.\(^2\)\(^4\) In two of the families that we have studied there has also been progression in the severity of the condition\(^2\)\(^6\) in that visual acuity and residual colour vision have been seen to deteriorate.

Combined results of previous studies\(^5\)\(^7\)\(^8\)\(^9\)\(^10\) provide evidence for the general conclusion, first put forward by Nathans et al, that there are different mutational pathways to BCM. The data suggest that 40% of blue cone monochromat genotypes are a result of a one step mutational pathway that leads to deletion of the LCR. The remaining 60% of blue cone monochromat genotypes comprise a heterogeneous group of multistep pathways. The evidence thus far shows that many of these multistep pathways produce visual pigment genes that carry the inactivating Cys203Arg mutation, which eliminates a highly conserved disulphide bond.\(^11\) This cysteine residue is located in the second extracellular loop of the opsin and, together with a conserved cysteine residue at position 126 in the first extracellular loop, forms a disulphide bond necessary for stabilisation of the tertiary structure of the protein.\(^12\)

These studies have failed to detect the genetic alteration that would explain the BCM phenotype in all assessed individuals.\(^2\)\(^3\)\(^4\)\(^5\)\(^6\)\(^8\)\(^9\)\(^10\) Indeed, Nathans et al\(^13\) found that in nine out of 35 individuals with BCM (25%), the structure of the opsin array did not reveal the genetic mechanism for the disorder. This failure to identify disease causing variants in the opsin array may suggest that there is genetic heterogeneity yet to be identified in BCM.

**BORNHOLM EYE DISEASE**

Myopia can be inherited as an autosomal recessive, autosomal dominant, or as an X linked trait and, in the latter case, it is well known as a component of congenital stationary night blindness\(^14\) and retinitis pigmentosa.\(^15\) X linked myopia has been reported in a large five generation Danish family that had its origins on the Danish island of Bornholm. The syndrome has therefore been named Bornholm eye disease (BED).\(^16\)\(^17\) In that family, the syndrome manifests as moderate to high myopia combined with astigmatism and impaired visual acuity. Additional signs are moderate optic nerve head hypoplasia, thinning of the retinal pigment epithelium in the posterior pole with visible choroidal vasculature, and abnormal photopic ERG flicker function as the most constant finding.\(^18\) Affected members in this family are all deuteranopes, with a stationary natural history. This disorder is therefore best characterised as an X linked cone dysfunction syndrome with myopia and deuteranopia.

Linkage analysis performed in the original BED family has mapped the locus to Xq28, in the same chromosomal region therefore as the L/M opsin gene array.\(^19\) It remains to be seen whether molecular genetic analysis of the opsin array will reveal mutations that account for both the cone dysfunction and the colour vision phenotype. However, it may also be possible that rearrangements within the opsin gene array will be found to account for the colour vision findings, while the cone dysfunction component of the disorder may be ascribed to mutation within an adjacent but separate locus. The cone dystrophy that has been mapped to Xq27 (COD2) however, displays a different phenotype which is progressive.\(^20\)

Nevertheless, it is becoming increasingly common in retinal molecular genetics to find that disparate phenotypes can be caused either by different mutations in the same gene, or even the same mutation in the same gene. In the latter situation it is currently believed that other genetic factors—namely, the “genetic context” within which the primary disease causing mutation is expressed, and/or environmental factors may determine the final phenotype.

**MANAGEMENT**

There is currently no specific treatment for any of the cone dysfunction syndromes. Nevertheless, it is important that the correct diagnosis is made in order to provide accurate information on prognosis and to offer informed genetic counselling. Prenatal diagnosis is possible when the mutation(s) causing disease in the family is known.

Although there is no specific treatment available for this group of disorders, the provision of appropriate spectacle correction, low vision aids, and educational support is very important. Photophobia is often a prominent symptom in the cone dysfunction syndromes and therefore tinted spectacles or contact lenses may be beneficial to patients, in terms of both improved comfort and vision. In achromatopsia a deep red tint is most effective, allowing wavelengths of low luminous efficiency for rod photoreceptors to be transmitted to the retina, while those of a higher luminous efficiency (short wavelength light) are absorbed by the filter.\(^21\) Incomplete achromats are thought to benefit more from reddish brown lenses rather than deep red lenses, which on account of their narrow spectral transmission, would eliminate their residual colour discrimination.\(^22\) In contrast, magenta tints which prevent rod saturation while allowing transmission of blue light are indicated in BCM.\(^23\)

**CONCLUSIONS**

The cone dysfunction syndromes comprise a group of disorders that are both clinically and genetically heterogeneous. Their phenotypes are now well characterised both clinically and psychophysically and many causative genes have been identified. Perhaps not surprisingly these genes mainly encode proteins involved in the cone phototransduction pathway. Other genes remain to be identified before the complete molecular pathology of this interesting group of disorders can be established.

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REFERENCES


etal. Evidence for slow progression, carrier fundus findings, and possible genetic linkage to achromatopsia, Hum Genet 1995;97:266–70.


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