# The Good, The Bad and The Orals

COPF# 92501-PH

Walter O. Whitley, OD, MBA, FAAO **Director of Optometric Services** Virginia Eye Consultants Regional Medical Director - Mid-Atlantic Eyecare Partners, LLC

Disclosures - Walter O. Whitley, OD, MBA, FAAO has received consulting fees, honorarium or research funding from:

- Apellis Pharmaceuticals: Consultant, Speaker Bausch and Lomb: Advisory Board, Speaker
- Bruder Healthcare: Advisory Board CSI Dry Eye Inc.: Consultant
- Dry Eye Coach Co-Medical Editor Epion Consultant
- Glaukos Consultant Harrow: Consultant
- 12-Vision, Inc: Consultant Iveric Bio: Consultant Mediprint Pharma: Consultant

- Alcon: Advisory Board, Consultant, Speaker
  Aldeyra Therapeutics: Consultant
  Allergan: Advisory Board, Consultant, Speaker

  Nordic Pharma: Consultant
  OD Immersion: Consultant

  - Regener-Eyes: Consultant Review of Optometry Contributing Editor Science Based Health: Advisory Board Sight Sciences: Consultant

  - Sun Pharmaceuticals: Advisory Board Tarsus Pharmaceuticals: Advisory Board
  - Thea Pharmaceuticals: Consultant Trukera Consultant
  - Visus Pharmaceuticals: Advisory Board
     Viatris Consultant

1

# Which Orals Do you Commonly Prescribe?

# **Prescribing Considerations**

- Indications
- Brand vs. generics
- · Does the insurance cover prescriptions?
- · Costs of medications
- Compliance

3

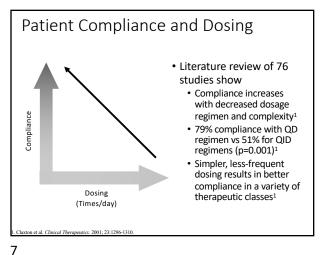
6

# Patient Assistance Programs

- · www.myalcon.com
- · www.allergan.com
- · www.bausch.com
- www.tarsusrx.com
- www.rxhope.com
- www.RxOutreach.org
- · www.NeedyMeds.org
- www.pparx.org
- www.goodrx.com

# Barriers to Adherence

- Cost
- Polypharmacy
- · Side effects
- Forgetfulness
- · Ability to take medications
- Providers role





# **Prescription Considerations**

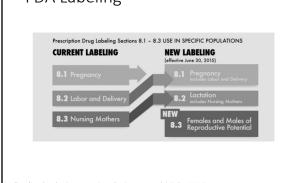
- · Review medical history
  - · Renal function
  - Liver function
- Review current medications
- Side effect vs. true allergies
- · Pregnant or nursing
- Rx for children

Prescribing for Women

- Certain medications are OK in pregnancy
- · Breast feeding
- · Consult OB-GYN if necessary

9

FDA Labeling



10

# So What Can Be Used During Pregnancy?

- Antibiotics
  - Amoxicillin
  - · Amoxicillin/clavulanate
  - Azithromycin
  - Erythromycin
- Antivirals
  - Acyclovir
  - Valacyclovir
- Anti-inflammatory
  - Prednisone

- Analgesics
  - Acetaminophen
  - Ibuprofen
  - Tylenol #3
  - Vicodin
- Allergy
  - Diphenhydramine
  - Loratadine

# What About Topical Medications **During Pregnancy?**

- Category B
  - Antibiotics tobramycin
  - Allergy alcaftadine
  - Glaucoma brimonidine
- · Category C
  - Allergy olopatadine
  - Anti-inflammatory steroids, cyclosporine
  - Anti-viral ganciclovir, trifluridine

# Prescribing Considerations for Kids

- 1. Know the age
- 2. Know the weight >88lbs
- 3. Look up the dosage
  - mg/kg/day
- 4. Be good at math
  - Or call the pharmacist
- 5. Avoid
  - Tetracyclines
  - Fluoroquinolones

13 14

#### Math is Fun!

- Augmentin 20-40 mg/kg/day po in 3 doses
- Convert pounds to kg
   50 lbs / 2.2 lbs = 23kg
- Multiply kg by dosage to get DAILY dose mg
   23kg \* 20 mg/kg = 460 mg
- Daily dose divided by number of doses per day
  460 mg / 3 = 150 mg po tid

Clinical References/Apps

**Epocrates** 

GoodRx

Eye Handbook

15 16

#### Case #1

- 34 YOWF presents with pain and swelling OD lid and right side of face. Started about 4 days ago suddenly and not getting better. Started amoxicillin/clavulanate 1 PO BID 2 days prior. She is 20 wks pregnant and can only take Tylenol for pain. Had a similar episode a couple years prior which
- Oc Hx: Epiphora OD
- Med Hx: 20 weeks pregnant / unremarkable

• Allergies: NKMA

#### Examination

- BCVA:
  - OD 20/25
  - OS 20/20
- Neg APD
- CVF/EOMs Full OU

# What is Our Likely Diagnosis?

- 1) Chalazion
- 2) Dacryocystitis
- 3) Dacryoadenitis
- 4) Preseptal cellulitis
- 5) Does it really matter???

# Which Antibiotic Would You Choose?

- 1) Change to ciprofloxacin 500mg BID PO
- 2) Erythromycin ointment BID OD
- 3) Continue amoxicillin/clavulanate 875 mg BID po
- 4) Change to azithromycin as directed
- 5) Does it really matter???

So what did we do??? We lanced it and kept on amoxicillin/clavulanate and Scheduled DCR

19

# Amoxicillin / clavulanate

- Penicillinase-resistant penicillin
- Disrupts the synthesis of peptidoglycan in bacterial cell walls
- Bactericidal
- $\bullet$  500 mg TID or 875 mg, 1000mg BID for one week
- Can be used in kids and pregnancy category B
- Side Effects/Contraindications:
  - Cannot be taken if penicillin allergy
  - GI upset
  - · Liver injury

Fluconazole

20

- Effective against candidiasis, cryptococcal meningitis
- Interferes with fungal cytochrome P450 activity (lanosterol14- $\alpha$ -demethylase) decreasing ergosterol synthesis
- Pregnancy Category C
- 150 mg as a single oral dose
- Side effect
  - Headache

21 22

# Eyelid / Conjunctival Cultures

- Eyelid
  - Moisten swab, rub along the lid margins
- Conjunctiva
  - Inferior palpebral conjuntiva
- · Inoculate solid media plates
- Culture
  - Calcium alginate swab
  - Cotton-tipped applicator
  - · Transport medium

**Example of Culture Report** 

- Hold for:
  - Bacteria 1 week
  - Viral 2 weeks
  - Fungal 1 month
- Test for all sensitivities

#### Treatments for MRSA

- 100% to vancomycin1
- 97.7% to sulfisoxazole1
- 95% to Polytrim<sup>2</sup>
- 93.2% were sensitive to tetracycline<sup>1</sup>
- 63.6% were sensitive to bacitracin<sup>1</sup>
- 14.8% of MRSA isolates were sensitive to ciprofloxacin and erythromycin1
- · Besifloxacin has been reported to be effective

Freidlin J. Acharya N, Lietman TM, et al. Spectrum of eye disease caused by methicillin-resistant staphylococcus aureus. Am J Ophthalmol. 2007 Aug;144(2):313-5 Asbell PA Colby KA Deng S, et al. Ocular TRUST: nationwide antimicrobial susceptibility patterns in ocular isolates. Am J Ophthalmol. 2008 Jun;145(6):951-58.

# Trimethoprim/sulfamethoxazole

- Trimethoprim: Inhibits dihydrofolate reductase, thereby blocking production of tetrahydrofolic acid from dihydrofolic acid
- Sulfamethoxazole: Inhibits bacterial synthesis of dihydrofolic acid by competing with para-aminobenzoic acid
- 80mg/400mg or 160mg/800mg
- Pregnancy Category C
- Effective against MRSA
- · Side Effects
  - Nausea, vomiting, loss of appetite
  - · Mild itching or rash
  - Neurological abnormalities / hyperkalemia / renal complications
     Steven Johnson's Syndrome

26 25

# You've Got to be Kidding Me!

- 27yowm presents with red, painful, blurry VA OS. Started 10 days ago after returning from a trip to Italy. Taking 500mg Naprosyn for HA.
- Health Unremarkable

• Vasx: OD 20/20-3

OS 20/25-3 with NI

• IOP: 9 / 10

• SLE:

- OD Mild limbal flush / 1+ Cells
- OS 2+ Inj / 2+ Cells

#### What is Your Treatment?

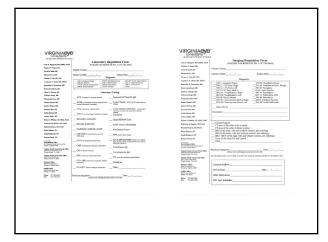
- Prednisolone acetate 1% vs. difluprednate 0.05% vs. loteprednol etabonate .5%
- Homatropine 5% vs. Scopolamine 0.25% vs. Atropine 1%
- · Would you consider lab testing?
- · Would you prescribe an oral medication?

27 28

- · Acute, bilateral non-granulomatous, anterior uveitis OU
- Cause???
- Treatment
  - Difluprednate qid OD, q2h OS
  - Cyclopentolate 2% TID OU

**Pulse Therapy** 

- QID to Q 1 Hour for 7 to 10 Days
- Zero Tolerance for AC Cells
- Avoids Surface Toxicity
- · Quick & Dirty
- Hit It Hard and Fast: Aggressive



# Screening Tests for Syphilis

- · Venereal Disease Research Lab (VDRL)
  - VDRL may become non-reactive in latent syphilis or after successful treatment
  - False positives may occur in:
    - Pregnancy
    - Infectious mononucleosis
    - · Systemic lupus erythematosis
- Rapid Plasma Reagin (RPR)
  - Alternative to VDRL

31

32

# Fluorescent Treponemal Antibody Absorption (FTA-ABS)

- Detects specific antibodies against T pallidum
- · Confirms diagnosis of syphilis
  - More specific than VDRL
  - · More sensitive in primary syphilis
- Test may remain positive for life
- Reactive:
  - Primary syphilis 95%
  - Secondary 100%
  - Late latent 100%
  - Tertiary 96%
  - False positives may occur in pregnancy and SLE

**Syphilis** 

- STD caused by T pallidum / great imitator / any tissue and organ
- Sexually active / multiple partners
- Systemic Sx Depends on stage primary painless ulcer / secondary skin rash palms, soles, trunk / tertiary neurosyphilis
- All types of ocular inflammation
- Labs
  - VDRL / RPR
  - FTA ABS
  - ESR elevated
- Tx penicillin therapy
- Good prognosis if treated early

33

34

36

# So He Has an Allergy to PCN?

- Augenbraun M, Workowski K. Ceftriaxone therapy for syphilis: report from the emerging infections network. Clin Infect Dis. 1999 Nov. 29(5):1337-8
  - Tetracycline, erythromycin, and ceftriaxone have shown antitreponemal activity in clinical trial

Doxycycline

- Inhibits bacterial protein synthesis
- Cannot be used for kids <8 and pregnancy/nursing</li>
   Category D
- Anti-infective dose: 100 mg BID for 10 days
- Anti-inflammatory dose: 50 mg BID for one month then qd 1-3 months
- Side effects/Contraindications:
  - GI upset: caution patient to take this with food
  - Photosensitivity
  - Pseudotumor cerebri

### Azithromycin

- Inhibit bacterial protein synthesis
- Z pak: 500 mg Day 1, 250 mg Day 2-5 or 1g dose
- Pregnancy Category B
- Side Effects/Contraindications:
  - Gl upset
  - Headache
  - Rash

37

- May worsen myasthenia gravis symptoms
- Kidney or liver dysfunction
- · Consider risk of fatal heart rhythms

# MGD: Azithromycin vs. Doxy

- Purpose: To assess the efficacy and safety of oral azithromycin compared with oral doxycycline in patients with meibomian gland dysfunction (MGD) who had failed to respond to prior conservative management.
- Conclusion: Although both oral azithromycin and doxycycline improved the symptoms of MGD, 5-day oral azithromycin is recommended for its better effect on improving the signs, better overall clinical response and shorter duration of treatment

Kashkouli et. Al. Oral azithromycin versus doxycycline in meibomian gland dysfunction: a randomised double masked open-label clinical trial. Br J Ophthalmol. 2015 Feb;99(2):199-204.

# MGD: Azithromycin vs. Doxy

- Satitpitakul V, Ratanawongphaibul K, Kasetsuwan N, Reinprayoon U. Efficacy of azithromycin 1.5% eyedrops vs oral doxycycline in meibomian gland dysfunction: a randomized trial. Graefes Arch Clin Exp Ophthalmol. 2019 Jun;257(6):1289-1294.
- Upaphong P, Tangmonkongvoragul C, Phinyo P. Pulsed Oral Azithromycin vs 6-Week Oral Doxycycline for Moderate to Severe Meibomian Gland Dysfunction: A Randomized Clinical Trial. JAMA Ophthalmol. 2023 May 1;141(5):423-429

# Ocular Uses for Antibiotics

Bacterial keratitis

Lyme disease

Canaliculitis

38

40

• Eyelid lacerations

Chalazion/Hordeolum

• Ocular surface disease

• Chlamydia

 Orbital blow-out fracture

DacryocystitisDacryoadenitis

Preseptal cellulitis

39

#### **Oral Antibiotics**

- Is an oral antibiotic truly needed?
- Coverage
- Check allergy information
- Consider generics
- Take full course of therapy

# **Blowout Fracture**

- Check VA
- Base and medial walls of orbit are very thin
- Does not need to be a major trauma
- · Look for trapped EOMs
- Sunken eye
- · Infraorbital hypoesthesia
- Diplopia
- Pain on eye movement
- Nausea

# Repair?

- · Within 2 weeks
  - Symptomatic diplopia within 30º of primary gaze
  - Muscle entrapment (prevent ischemia and necrosis)
  - Fracture greater than 50% of orbit floor
  - · Displaced orbital rim fracture
  - > 2mm of enophthalmos
  - · Significant hypo-ophthalmos
- Monitor
  - Diplopia outside central 30º
  - · Modest isolated fractures
  - Improvement over first 2 weeks

43

# Cephalexin

- First generation cephalosporin
- · Caution in those with PCN allergy
- Disrupts the synthesis of peptidoglycan in bacterial cell walls
- Bactericidal but less susceptible to penicillinase
- Pregnancy Category B
- 500 mg BID for one week
- 5-10% cross-sensitivity with PCN
- Side Effects/Contraindications:
  - Renal dysfunction
  - GI disease

#### Orbital Trauma in Children

- Trap door orbital floor fractures are very common
  - · More elastic orbits
  - More common to get muscle entrapment

Abx Cross-sensitivity Reactions

when using first-generation cephalosporins or

• For penicillin-allergic patients, the use of third- or

cephalosporins with dissimilar side chains than the

offending penicillin carries a negligible risk of cross

Campagna, J.D. et. Al. The use of cephalosporins in penicillin-allergic patients: a literature review. J Emerg Med. 2012

cephalosporins with similar R1 side chains

fourth-generation cephalosporins or

• The overall cross-reactivity rate is approximately 1%

• Evaluation for repair typically in 5-7 days

<u>—</u> 45

46

44

#### Levofloxacin

- Broad spectrum antibiotic
- Can be used in patients with PCN allergy
- Not to be used in children, pregnancy, nursing
- Dosage: 500 mg qd for one week
- Warnings:
  - Tendinitis and tendon rupture
  - Increase risk of RD?
  - · Peripheral neuropathy

# Spider Bite

allergy

- 63YOWM Referred by PCP for sudden decrease VA OD and swelling of eyelids OD>OS for 1 week
  - Pressure from forehead to cheek
  - · Worse in evenings
  - · Mild seasonal allergies
  - Some tearing and redness OD
- Bitten 3 weeks ago on top of the head while working in the yard which become swollen that evening
- Went to PCP and given oral ABX which finished yesterday

47

#### Examination

Non-healing scab on R forehead

 Conjunctiva: 2+ injection OD
 Cornea: 2+SPK, 2+ MCE, 1+ KPs, No dendrites OD

• AC: 2+ Cells OD

• Lens: 2+ NS OD / 1+NS OS

• IOP: 31/13

# Diagnosis???

- Considerations
  - PCP told him he had an infection not shingles
  - Episode started 3 weeks prior
- Treatment
  - Valacyclovir 1000mg TID po
  - Difluprednate QID OD
  - Timolol 0.5% QAM OD
  - F/u 1 week

49

50

# Herpes Varicella-Zoster Virus

- Primary infections: Chicken pox
  - Remains latent in dorsal root or other sensory ganglia after primary infection
  - May lie dormant for years to decades
- Later infections: Shingles
  - Virus specific cell-mediated immune responses decline
  - Localized cutaneous rash erupting in a single dermatome
  - HZO accounts for 10-25% of all cases of shingles

# Herpes Zoster Ophthalmicus

- 90% of U.S. population infected with VZV by adolescence
- 100% of U.S. population by 60 years of age
- 1.5-3.4 cases per 1,000 individuals

http://emedicine.medscape.com/article/783223-overview#aw2aab6b4

51

52

# Herpes Zoster Ophthalmicus

- Conjunctivitis
- Scleritis
- Pseudodendrites
- Keratic precipitates
- Iritis
- Synechiae
- Neurotrophic keratitis
- Elevated IOP
- Potential vascular occlusion
- Nerve palsies
- Glaucoma (longer-term)

# HZO: Signs and Symptoms

- Prodromal phase: fatigue, malaise, low-grade fever
- Unilateral rash over the forehead, upper eyelid, and nose
  - 60% of patient have dermatomal pain prior to rash
  - Erythematous macules to papules to vesicles to pustules to crusts
  - Other symptoms: eye pain, conjunctivitis, tearing, decrease VA, eyelid rash
  - Hutchinson's sign
- Post-herpetic neuralgia: >12 months for 50%

53

#### HZO: Treatment

- Local wound care
- Analgesia
- Antivirals
  - Valtrex 1g TID
- Antibiotics??
- Oral corticosteroids
- Post-herpetic neuralgia
  - Tricyclic antidepressants
  - Topical capsaicin ung
  - Gabapentin

### **Oral Antivirals**

- Inhibit viral DNA polymerase without inhibiting normal cellular activity
- Works best if treatment initiated within 72 hours
- Pregnancy category B
- · Caution in patients with renal disease

Antiviral Drug	HSV	HZO
Acyclovir	400 mg 5x/day for 1 week	800 mg 5x/day for 1 week
Valacyclovir	500 mg TID for 1 week	1000 mg TID for 1 week
Famciclovir	250 mg TID for 1 week	500 mg TID for 1 week

55

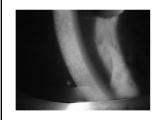
# Vaccines for HZO - Shingrix (GSK)

- Vaccine indicated for prevention of herpes zoster (shingles) in adults aged 50 years and older
- 2 IM doses (0.5 mL each) at 0 and 2 to 6 months
- Reduced risk of developing HZ from 85 – 97%
- Efficacy against PHN 85.5%

Data available from Shingrix PI

57

# Orals for Simplex???





58

56

# Herpetic Eye Disease Study I

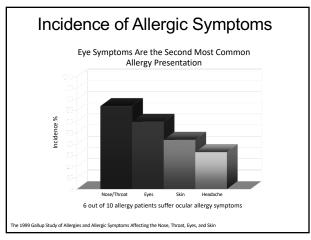
- · Herpes Stromal Keratitis, Not on Steroid Trial
  - Pred Phosphate faster resolution and fewer treatment failures
  - Delaying treatment did not affect outcome
- Herpes Stromal Keratitis, on Steroid Treatment
  - No apparent benefit in the addition of oral acyclovir to the treatment of topical corticosteroid and topical antiviral
- HSV Iridocyclitis, Receiving Topical Steroids
  - Trend in the results suggests benefit in adding oral acyclovir

# Herpetic Eye Disease Study II

- HSV Epithelial Keratitis Trial
  - No benefit from oral ACV with topical trifluridine in preventing the development of stromal keratitis / iritis
- Acyclovir Prevention Trial
  - Reduced by 41% the probability of recurrence
  - 50% reduction in the rate of return of the more severe form
- Ocular HSV Recurrence Factor Study
  - No results available

# Herpes and Bell's Palsy

- HSV or HZV has been shown to cause Bell's Facial Nerve Palsy
- Main concern is dry eye secondary to poor lid function



62

61

#### Oral Antihistamines

- · Central acting antihistamines
  - Benadryl (diphenhydramine)
  - Chlortrimeton
- · Peripherally acting antihistamines
  - Selective peripheral histamine H1 receptor blockade
  - Less CNS and anticholinergic effects
  - · Less sedating but also less effective
  - With / without a decongestant

# Common OTC Allergy Meds

- Inhibits action of histamine by blocking H1 receptors preventing symptoms of allergy
  - Cetirizine (Zyrtec) 5 or 10mg qd
  - Desloratadine (Clarinex) 5mg qd
  - Fexofenadine (Allegra) 60mg bid; 180mg qd
  - Loratidine (Claritin) 10 mg qd
- Pregnancy category C
- Side Effects/Contraindications:
  - Hypersensitivity reactions
  - Dry mouth
  - Headache

Nervousness

63

65

64

# Diphenhydramine

- Temporarily relieves these symptoms due to hay fever or other upper respiratory allergies
  - Runny nose
  - Sneezing
  - Itchy, watery eyes
  - Itching of the nose or throat
- Pregnancy Category B
- 25-50 mg q6-8 hours
- Side effects
  - Blurred VA, Diplopia
  - CNS depression / somnolence

#### Montelukast sodium

- Leukotriene receptor antagonist
- Indications:
  - Prophylaxis and chronic treatment for asthma
  - Acute prevention of exercise-induced brochoconstriction
  - · Relief of symptoms of allergic rhinitis
- 10 mg tablet qd
- Side effects
  - Behavior or mood changes, URI, fever, headache, sore throat, cough, stomach pain, diarrhea, ear ache or ear infection, flu, runny nose, and sinus infection

Triaging Questions

Risk Pactor Analysis

Diagnostic Tests

Diagnostic Tests

Copyright © 2017 Elsewier Inc. James and Conditions.

68

70

67

# Oral Medications for Dry Eye

- Nutritional supplements
  - 1,000 mg BID of Omega-3 Fish Oil
- Lovaza (Rx fish oils)
  - 4g per day po
  - Indicated as an adjunct to diet to reduce triglyceride levels in adult patients with severe hypertriglyceridemia
- Oral pilocarpine
  - Salagen\*: 5 mg qid for dry mouth
  - Evoxac\*: 30 mg tid for dry mouth

69

Effect of Oral Re-Esterified Omega-3 Nutritional Supplementation on Dry-Eye Disease: Double-Masked Randomized Placebo-Controlled Study

- 105 patients with dry eye disease
  - Four capsules (2 gm) once a day containing 1680mg EPA and 560mg DHA (PRN Dry Eye Omega Benefits) for 3 months or four capsules of placebo.
  - All patients underwent a screening, baseline, 6 week and 12 weeks visit.
  - On each visit patients were tested for tear osmolarity, MMP-9, fluorescein corneal staining, Schirmer's testing, and OSDI. On the screening exam and week 12 evaluation patients had their omega index tested.
- This study demonstrated that oral consumption of reesterified omega-3 fatty acids (1680 mg EPA and 560 mg DHA once daily for 12 weeks) is an effective treatment of dry eye disease and results in a statistically significant improvement in tear osmolarity, OSDI, tear break up time and omega index levels.

Donnenfeld ED, Holland, EJ, Bucci FA, et.

District Americal of Numerical page 1 of 10

O'The America 200

Lipid structure does not modify incorporation of EPA and DHA into blood lipids in healthy adults: a randomised-controlled trial

Anneete L. West<sup>1</sup>, Graham C. Burdge<sup>1</sup> and Philip C. Cilder<sup>12</sup>

\*Anderset list of Homes Excellegament and Joseph (Secretary) of suchampane, Suchampane 3016 GHz, IX:

\*Natural Intensity in Intensity in Intensity in Suchampane and Joseph (Secretary) of suchampane, Suchampane 3016 GHz, IX:

\*Natural Intensity in Franciscum True, Electrony of Suchampane, Suchampane 3016 GHz, IX:

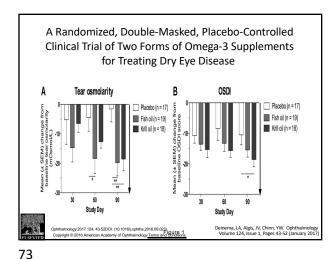
\*Natural Intensity in Franciscum True, Electrony of Suchampane, Suchampane 3016 GHz, IX:

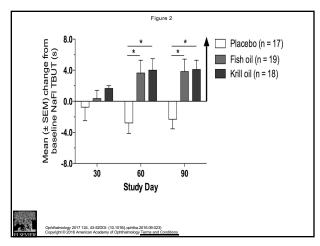
\*Columnal Intels Secretary True American Natural Intensity of Suchampane, Suchampane 3016 GHz, IX:

\*District Such 2016 - Franciscum True, Electrony of Suchampane, Suchampane 3016 GHz, IX:

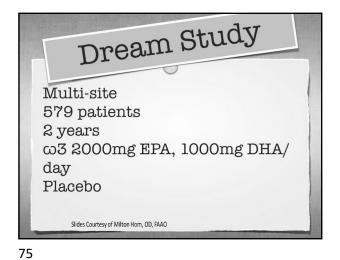
\*Columnal Intels Secretary Suchampane Natural Intels Secretary of Suchampane, Suchampane 3016 GHz, IX:

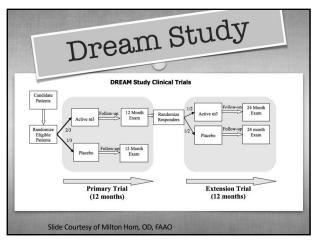
\*Part 11 Suchampane 1 Suchampane 1





74





76

#### Results

- No significant difference between fish / olive oil groups
- However, both groups improved significantly in primary endpoint of symptoms (plus secondary endpoints of corneal / conj staining, TBUT)
- Olive oil may not have been best choice for study
- There's been much confusion over findings

#### What does this mean to clinicians?

- We should all make effort to educate ourselves on research

   headlines often get it wrong
- Fish oil may be beneficial, but may not be the only option we should consider
- Other omegas (e.g. GLA), and nutrients have clinical evidence in OSD, weren't examined in DREAM

Can daily supplementation with marine  $\omega$ -3 fatty acids prevent the development of dry eye disease (DED)?

- Findings In this randomized clinical trial of 23,523 US adults who at study entry were free of a previous diagnosis of DED and were not experiencing severe dry eye symptoms, daily supplementation with 1 g of marine ω-3 fatty acids for a median (range) 5.3 (3.8-6.1) years had no significant effect on the incidence of diagnosed DED or reported DED symptoms.
- Meaning The results do not support recommending marine  $\omega$ -3 fatty acid supplementation to reduce the incidence of DED.

# Eyelids and Pain

- Typically inflammation induced
- Many capsaicin receptors

79 80

# Eyelids

- Pain is often inflammation and swelling based
- Decrease swelling = decrease pain
  - · Cold vs. Hot compress
  - Steroid Dosepak
  - Loteprednol ung

# It's Red Again

- 38 year old, African American, Female presents with red, painful, and photophobic OS
- Started 3 weeks ago / similar episode 10 years ago
- Tried dexamethasone 0.1% but no relief
- BCVA OD 20/25 OS 20/20
- IOP: 17 mmHg

81 82

# Differentials

- Conjunctivitis
- Episcleritis
- Scleritis
- Uveitis

#### When Should Lab Tests Be Ordered?

- · Bilateral cases
- Hyperacute cases
- · Atypical age group
- · Worsens with tapering
- Recurrent uveitis
- VA worsens
- Scleritis
- Immunosuppressed
- Recalcitrant cases

#### Treatment for Scleritis

- NSAIDS
- Systemic steroids
- Immunosuppressive therapy
- Topical steroids???

# Ibuprofen

- Analgesic, antipyretic, anti-inflammatory properties
- Suppresses inflammatory cascade by inhibiting COX pathway
- Pregnancy Category
  - C Prior to 30 weeks gestation
  - D After 30 weeks gestation
- OTC 200 mg tablets/capsules
- Analgesic dosage 1,200 mg / day
- Anti-inflammatory dosage 3,200 mg / day
- · Generics available

85 86

# Oral Anti-inflammatory Indications

- · Allergic reaction
- Uveitis
- Scleritis
- AION
- Optic neuritis
- Orbital pseudotumor
- Herpes zoster
- Contact dermatitis
- Episcleritis

# 82 yowf Sudden Loss of VA OD

- VAcc:
  - OD: LP
  - OS: 20/50 +2
- CVF
  - OD: constricted inferior 180
  - OS: Full to finger counting

- Pupils
  - OD: 1+ APD
  - OS: round and reactive
- IOP: 18mmHg/18mmHg by Goldmann
- EOM
  - Full OU

88

Ocular history:

87

- Primary open angle glaucoma OU
- Epithelial basement membrane dystrophy OU
- Pseudophakia OU
- Early Dry ARMD OU
- Medical history:
  - Arthritis
  - Hypertension
  - High Cholesterol
  - Peripheral NeuropathyRestless leg Syndrome
  - GERD

- Ocular Medication
  - Timolol/brimonidine BID OS
  - Travaprost QHS OU
- Systemic Medication
  - Crestor 5mg
  - Amlodipine-Benazepril 5/10mg
  - Pramipexole 0.125mg
  - Tramadol HCL
  - Nexium 40mg
  - Lidoderm patch
  - Gabapentin 300mg
  - Celebrex 200mg
  - Iron supplement
  - Krill oil supplement

#### **PW-GCA**

- Assessment
  - Ischemic Optic Neuropathy OD
    - Pt denied any jaw pain, headaches, shoulder or hip pain, change in weight and malaise
- Plan
  - · Labs ordered: ESR, CRP, CBC w/diff
  - Medication: Prednisone 20mg 3 PO
  - Meds are not to be started before having blood drawn
  - Follow up in 1 week pending lab results

#### PW - GCA

- Lab Results:
  - ESR: 95 (High)
  - CRP: 7.09 (High)
  - Platelet: 465 (High)
- Temporal artery biopsy scheduled in 2 weeks

91

92

#### PW-GCA

- Temporal Artery Biopsy Result
  - Active arteritis with rare giant cells, consistent with temporal arteritis
  - Mild arteriosclerosis
  - · Disruption and focal loss of internal elastic lamina
- Informed the patient that her PCP will monitor her labs from now on and adjust her oral prednisone dose accordingly. She is to continue on the 60mg/day dosing for right now until he instructs her otherwise
- Follow up in 1 month

Prednisone

- Suppresses inflammatory cascade and immune response
- Optic neuritis
  - Methylprednisolone 1g/day i.v. for 3 days
  - 60-100mg qd p.o. for 11 days
  - Only after initial IV steroid treatment per ONTT to decrease risk of recurrence
- AION: 60-100mg qd
- Scleritis/Uveitis
  - Not responding to topical treatment
  - 40-80 mg as an initial dose with taper

93

94

#### Prednisone

- Side Effects/Contraindications:
  - Increased IOP
  - Cataract formation
  - Fluid retention (moon face, buffalo hump)
  - Increase blood sugar levels in diabetics
  - Gastric ulcers
  - · Not to be used if pregnant
  - Mood changes
- Advantages:
  - Widely available
  - Inexpensive

Oral Corticosteroid Considerations

- Accurate diagnosis is essential
- Indicated for acute inflammatory eye, orbital and eyelid conditions
- Pregnancy category C
- Dosepaks available
  - $\bullet\,$  24 mg, 30 mg, 60 mg with taper
- Best taken with meals
- Short term rarely has ocular side effects

95

# **Oral Analgesics**

- Hydrocodone/acetaminophen is the most frequently prescribed oral medication in the U.S.
- · Indicated for:
  - Corneal abrasions
  - · Recurrent corneal erosions
  - · Severe keratitis
  - Severe iritis
  - · Refractive surgery

# **Aspirin**

- Pain, inflammation, fever, anti-platelet
- Pregnancy Category D
- OTC 325-650mg every 4-6 hours
- Avoid aspirin 1-2 weeks prior to surgery
- Consider in patients with CRVO, retinal emboli
- Side effects
  - Hypersensitivity
  - Rhinitis
  - Bleeding disorders
  - · Reye's syndrome
  - Pregnancy

97

98

# Acetaminophen

- Analgesics and antipyretic
- Indications:
  - Pain relief associated with corneal abrasions, chemical burns, headaches associated with eye pain, scleritis
- Pregnancy Category B
- Side Effects/Contraindications:
  - Rash, Hives
  - Itching
  - Difficulty swallowing/breathing
  - Overdose may damage liver
  - Do not take with alcohol

# Oral Narcotic Analgesics

- · Centrally acting opioid receptor blockers
- Safe and effective for acute, short-term pain
- Clinically used in combination with acetaminophen
- Generally prescribed as one tablet po q4-6hours prn
- Onset 20 minutes, peak 1 hour, duration 4-6 hours

99

100

# Hydrocodone/acetaminophen

- Dosage:
  - • Vicodin contains 5mg hydrocodone with  $\underline{\textbf{300 mg}}$  APAP
  - Vicodin ES contains 7.5mg hydrocodone with 300 mg APAP
  - Vicoprofen contains 7.5mg hydrocodone with 200 mg ibuprofen
  - Pregnancy Category C
- 1 tablet po q4-6 hours
- Indicate how many in writing
- · Generics available

Controlled Drug Act

- Schedule I drugs with a high abuse risk. These drugs have NO safe, accepted medical use in the United States. Some examples are heroin, marijuana, LSD, PCP, and crack cocaine.
- Schedule II drugs with a high abuse risk, but also have safe and accepted medical uses in the United States. These drugs can cause severe psychological or physical dependence. Schedule II drugs include certain narcotic, stimulant, and depressant drugs.
- Schedule III, IV, V drugs with an abuse risk less than Schedule II. These drugs also have safe and accepted medical uses in the United States. Schedule III, IV, or V drugs include those containing smaller amounts of certain narcotic and nonnarcotic drugs, anti-anxiety drugs, tranquilizers, sedatives, stimulants, and non-narcotic analgesics.

Accessed from http://www.deadiversion.usdoj.gov/schedules/index.html on 8/20/1:

101

# Tramadol hydrochloride

- Moderate to severe pain
- Non-narcotic opioid receptor agonist
- Pregnancy Category C
- 50-100mg q4-6 hours
- Side effects
  - Hallucinations
  - Fever
  - · Nausea and vomiting
  - Seizure
  - Skin rash
  - Shallow breathing, weak pulse

#### Side Effects of Pain Meds

- Constipation
- Nausea and vomiting
- Sedation
- Dizziness
- Itching
- Respiratory depression
- Addiction

103 104

#### Promethazine

- Used to treat allergy symptoms
- Prevents motion sickness
- Treats nausea and vomiting or pain after surgery
- · Sedative or sleep aid
- 25 mg QID po

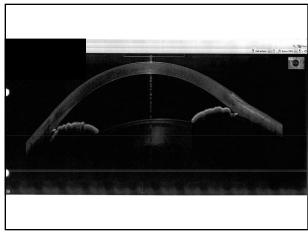
# Angle Closure Glaucoma

- Fewer than 10% of US glaucoma cases
- Anatomically narrow angle
- Sex
  - 3 X higher in caucasian women
  - In blacks, men + women equally affected
- Incidence increases with age

105 106

# **ACG Treatment Options**

- Surgical Care
  - · Laser iridotomy
  - · Laser gonioplasty
- Medication
  - Alpha-adrenergic agonist
  - Beta-blockers
  - Miotic agents
  - Prostaglandins
  - · Carbonic anhydrase inhibitors



107 108

ancet. 2016 Oct 1;388(10052):1389-1397. doi: 10.1016/S0140-6736(16)30956-4.

Effectiveness of early lens extraction for the treatment of primary angle-closure glaucoma (EAGLE): a randomised controlled trial.

zuara-Blanco A<sup>1</sup>, Burr J<sup>2</sup>, Ramsay, C<sup>3</sup>, Cooper D<sup>3</sup>, Foster PJ<sup>4</sup>, Friedman DS<sup>5</sup>, Scotland G<sup>6</sup>, Javanbakht M<sup>7</sup>, Cochrane C<sup>3</sup>, Norrie J<sup>6</sup>; EAGLE study group.

- Randomized trial: PAC and PACG patients > 50 yo without cataract
  - · Clear Lens Extraction
  - LPI with topical meds

30% reduction in IOP and 53% decrease in number of medications

ACUTE ACG - 71% IOP reduction and almost no need for post-phaco medications

110

109

#### Acetazolamide

- · Carbonic Anhydrase Inhibitor
- Diuretic, interfering with conversion of bicarbonate and reducing aqueous formation
- Contraindicated in renal, hepatic, and respiratory disease
- Pregnancy Category C
- Decreases IOP by 40-60%
- 125 mg, 250 mg, 500 mg sequels

#### Acetazolamide

- · Angle closure
  - 250mg X 2 plus topical glaucoma meds
  - Urgent LPI
- Chronic open angle glaucoma
- Diamox sequels 500mg BID
- Pseudotumor cerebri
  - Must have labs performed prior to treatment

Randomized Controlled Trial > Lancet. 2019 Apr 20;393(10181):1609-1618 doi: 10.1016/S0140-6736(18)32607-2. Epub 2019 Mar 14.

Laser peripheral iridotomy for the prevention of angle closure: a single-centre, randomised controlled trial

Interpretation: Incidence of angle-closure disease was very low among individuals classified as primary angle closure suspects identified through community-based screening. Laser peripheral iridotomy had a modest, albeit significant, prophylactic effect. In view of the low incidence rate of outcomes that have no immediate threat to vision, the benefit of prophylactic laser peripheral iridotomy is limited; therefore, widespread prophylactic laser peripheral iridotomy for primary angle-closure suspects is not recommended.

- MRI
- Spinal tap
- Diamox Sequels 500 mg BID

111 112

#### Acetazolamide

- Side Effects/Contraindications:
  - · Not for those with sulfa allergy
  - · Kidney dysfunction
  - · Metallic taste
  - Tingling and burning in hands/feet
  - Aplastic anemia
  - Diabetics susceptible to ketoacidosis

Ocular Side Effects of Systemic Medications

# 10 Most Commonly Prescribed Drugs

- 1) Hydrocodone (combined with acetaminophen)
- 2) Generic Zocor (simvastatin), a cholesterol-lowering statin drug
- 3) Lisinopril (brand names include Prinivil and Zestril), a blood pressure
- 4) Generic Synthroid (levothyroxine sodium), synthetic thyroid hormone
- Generic Norvasc (amlodipine besylate), an angina/blood pressure drug
- 6) Generic Prilosec (omeprazole), an antacid drug
- 7) Azithromycin (brand names include Z-Pak and Zithromax), an antibiotic
- 8) Amoxicillin (various brand names), an antibiotic
- 9) Generic Glucophage (metformin), a diabetes drug
- 10) Hydrochlorothiazide (various brand names), a water pill used to lower blood pressure

http://www.webmd.com/news/20110420/the-10-most-prescribed-drugs

115

Drug	Use	Ocular Effect
Alendronate	Osteoporosis	Episcleritis/scleritis/uveitis
Amiodorone	Ventricular arrhythmia	Optic Neuropahty
Antihistamines	Allergies	Decreased tear secretion
Chloroquine	Arthritis / Lupus	Bull's Eye Maculopathy
Contraceptives	You know	Decrease lacrimation
Corticosteroids	Anti-inflammatory	Increased IOP / PSC
Coumadin	Anticoagulant	Retinal Heme / SCH
Digoxin	Congestive Heart Failure	Yellow vision
Isotrentinoin	Recalcitrant acne	Dry eye / MGD
Gold Salts	Arthritis	Deposits on K / Lens
Sildenafil citrate	ED	NAION
Tamoxifen	Anti-cancer	Crystalline retinopathy
Tamsulosin	BPH	IFIS
Tetracycline	Antibacterial	Pseudotumor
Thioridazine	Anti-psychotic	Pigmentary retinopathy
Thorazine	Anti-psychotic	Stellate ASC
Topiramate	Migraines	Acute myopica / ACG

116

118

#### Sudden Decrease in VA

- 24 YOWF, decrease VA 5 minutes ago, OD temp VF
- Bitten by a spider 3 days prior
- Ohx: LASIK 3 months prior
- Mhx: Unremarkable
- Shx: PT smoker
- VA ODsc: 20/25 OSsc: 20/20
- (-) APD • GAT: 7/10

Baseline VF 06/01/11



117

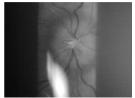
# OD Visual Field 06/03/11 OD 06/08/11 OD

# **Oral Contraceptives**

- You know what it's for
- Ocular Side Effects
  - Optic neuritis
  - Pseudotumor
  - Dry eye
- · Increase risk with smoking
  - Heart attack
  - Blood clots
  - Stroke
  - HTN
  - Migraines

119 120

### Pseudotumor Cerebri





121

# Hydroxychloroquine sulfate

- Indicated for the treatment of discoid and systemic lupus erythematosus, rheumatoid arthritis, and malaria
- Dosage: 200mg to 400mg per day
- · Primary risk factors
  - Duration > 5 years
  - Cumulative dose >1000g

  - Systemic High BMI, liver, kidney dysfunction
  - Ocular retina or macular changes

123

# Fingolimod

• FDA-approved oral disease-modifying therapy for the treatment of <u>multiple sclerosis</u> (MS). It acts by modulating the sphingosine-1-phosphate (S1P) receptor, preventing the release of lymphocytes into the systemic circulation and therefore reducing autoimmune demyelination in the central nervous system Macular edema (ME) has been reported as a significant ocular adverse event associated with the use of fingolimod for the treatment of MS and is termed fingolimod-associated macular edema (FAME).

What Medications Exacerbate Dry Eyes?

- Anti-hypertensive agents
- Antihistamine / Decongestants
- · Hormonal Replacement Therapy
- Antidepressants
- Pain Relievers
- GI medications
- Chemotherapy
- Antipsychotics

122



American Academy of Ophthalmology Statement

#### **Recommendations on Screening for** Chloroquine and Hydroxychloroquine Retinopathy (2016 Revision)

The Refringancy is not reversible, and there is no present therapy, Recognition at an early stage (before lices) is important to prevent central visual loss. However, questionable lest results should be repeated or a visu

124

#### Fluoxetine

- Antidepressant
- · Ocular side effects
  - Dry eye
  - Blurred VA
  - Mydriasis
  - Photophobia
  - Diplopia
  - Conjunctivitis
  - Ptosis

125 126

# Phenothiazines

- Indicated for depression and anxiety
- Ocular side effects
  - Night blindness
  - Anterior subcapsular cataracts
  - Salt and pepper fundus

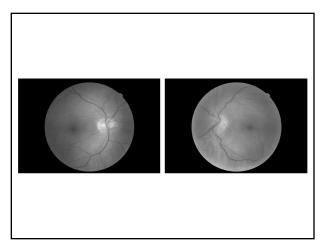
# Levothyroxine

- Indicated for hypothyroidism
- Ocular side effects
  - Diplopia
  - Ptosis
  - EOM paralysis
- Graves Diseases
  - 5X W>M
- NOSPECS

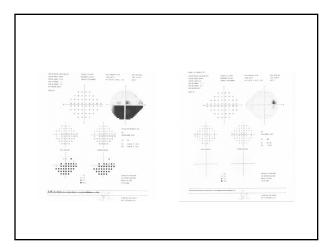
127 128

# Ethambutol

- Indicated for tuberculosis
- Used in combination:
  - Rifampin
  - Isoniazid
  - Pyrizinamide
- Ocular side effects
  - Optic neuropathy
  - SE dose dependent



129 130



# Sildenafil citrate

- Selective inhibitor of phosphodiesterase type 5
- Impairment of color discrimination (B/G)
- Non-arteritic ischemic optic neuropathy

131 132

# Can Anything be Done?

- Hayreh SS, Zimmerman MB. Non-arteritic anterior ischemic optic neuropathy: Role of systemic corticosteroid therapy. Graefes Archives of Clinical Exp Ophthalmology 2008; 246:1029-1046.
- Initial VA 20/70 or worse, treated within 2 weeks of onset of symptoms
  - Visual outcome at 6 mo
    - · Treated eyes 70% improved
  - Untreated eyes 41%.
  - Visual Fields
    - Treated 40.1% improvement
    - Untreated 24.5% improvement

#### Tamsulosin

- Indication for the treatment of benign prostatic hyperplasia
- Alpha-1 blocker
- Intraoperative floppy iris syndrome
- Importance to communicate prior to cataract surgery



133

134

#### Amiodorone

- Indicated for the treatment of life-threatening recurrent ventricular arrhythmia
- Side Effects
  - Halos
  - Photosensitivity
  - Optic neuropathy
  - Optic neuritis
  - Disc swelling



Differentials for Vortex Keratopathy

- · Drug induced
  - Amiodorone
  - Chloroquine
  - Tamoxifen
  - Ibuprofen
  - Indomethacin
- Stem cell deficiency
- Fabry's disease

135

136

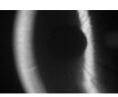
# Fabry Disease

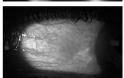
- X-linked disorder due to a deficiency of alphagalactosidase resulting in the buildup of globotriaosylceramide
- Signs and symptoms include:
  - Severe pain in the extremities
  - Exercise intolerance
  - Renal involvement
  - Skin lesions angiokeratoma corporis discusum consists of clusters of superficial cutaneous dark-red angiokeratomas
  - Tortuosity of conjunctival and retinal vessels

#### Alendronate

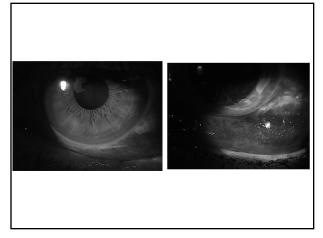
- Osteoporosis in postmenopausal women, Paget's disease
- Inhibitor of osteoclaticmediated bone resorption
- Side Effects
  - Episcleritis
  - Scleritis
  - Uveitis

138



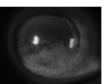


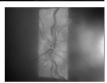
137



# Isotrentinoin

- Used in the treatment of recalcitrant nodular acne
- Dry eyes
- Keratitis
- Conjuctivitis
- Decreased night VA
- Cataracts
- Optic Neuritis
- Pseudotumor cerebri





139 140

# Topiramate

- Indicated for the prophylaxis of migraine headaches
- Choroidal effusions
- Acute myopia
- Acute angle closure
- Treatment cessation of drug, cycloplegics, and topical hypotensives

Phentermine and Topiramate

- FDA approved for weight loss in overweight adults who are also diagnosed with hypertension, type 2 diabetes mellitus, or hyperlipedimia
- Ocular Side Effects
  - Acute myopia
  - Bilateral Angle closure glaucoma
  - Choroidal effusion

141 142

# Tamoxifen

- Anti-estrogen used as adjunctive therapy for the treatment or prophylactics of breast cancer
- Maculopathy with crystalline deposits and macula edema

Home Remedies, Herbal Supplements and Whatever MOM Told Me to Take

# My Top 10 Oral Meds

- Preseptal Cellulitis Keflex 500mg BID
- MGD Azithromycin as directed
- HSV/HZO Valacyclovir 500/1000mg TID
- Allergies Singulair 50mg QD
- DED Hydroeye
- Pain Ibuprofen 400mg TID
- Pain Vicodin 5mg hydrocodone/ 300 mg APAP q4-6hrs
- Glaucoma Diamox 500mg BID
- AMD Macula Protect Complete / AREDS 2
- Inflammation Prednisone / Dosage varies

# Conclusions

- Many prescription options available to treat a variety of ocular condition
- Consider patient age, history, drug interactions, compliance, cost
- Important to treat and monitor
- Practice to the fullest extent of our education!

145 146

# THANK YOU

wwhitley@cvphealth.com

147