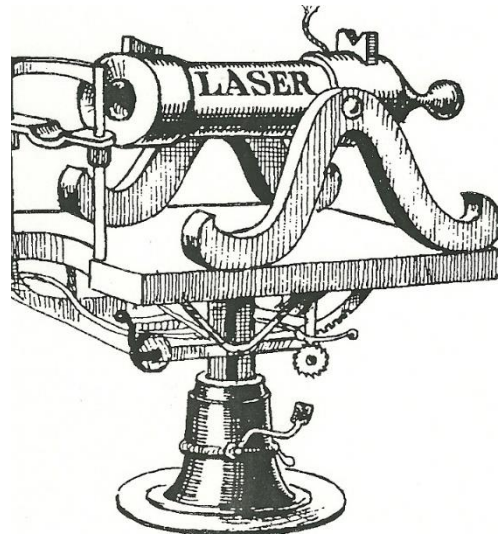


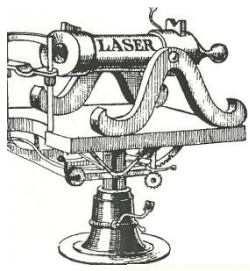
Market requirements for next generation fiber lasers for medical applications



MERIDIAN

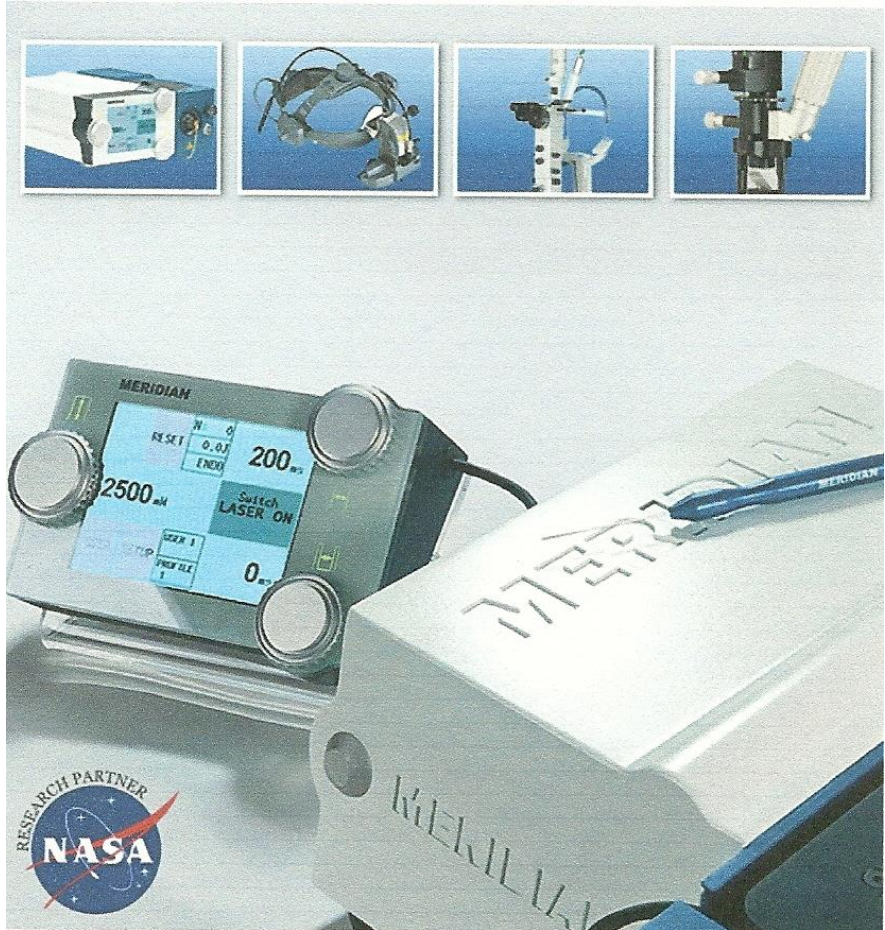


SWISS MADE

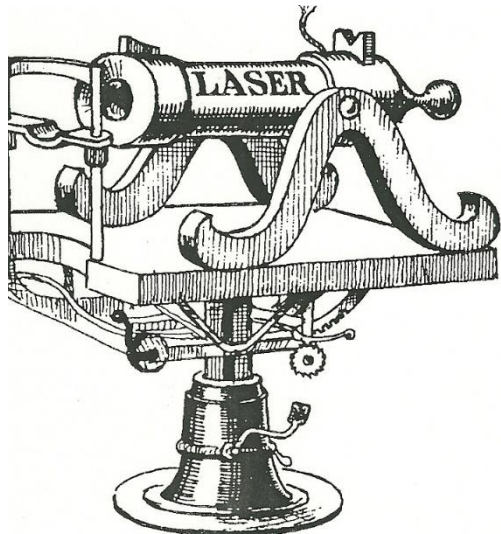


MICRORUPTOR 6

MERIDIAN



Market requirements for next generation fiber lasers for medical applications

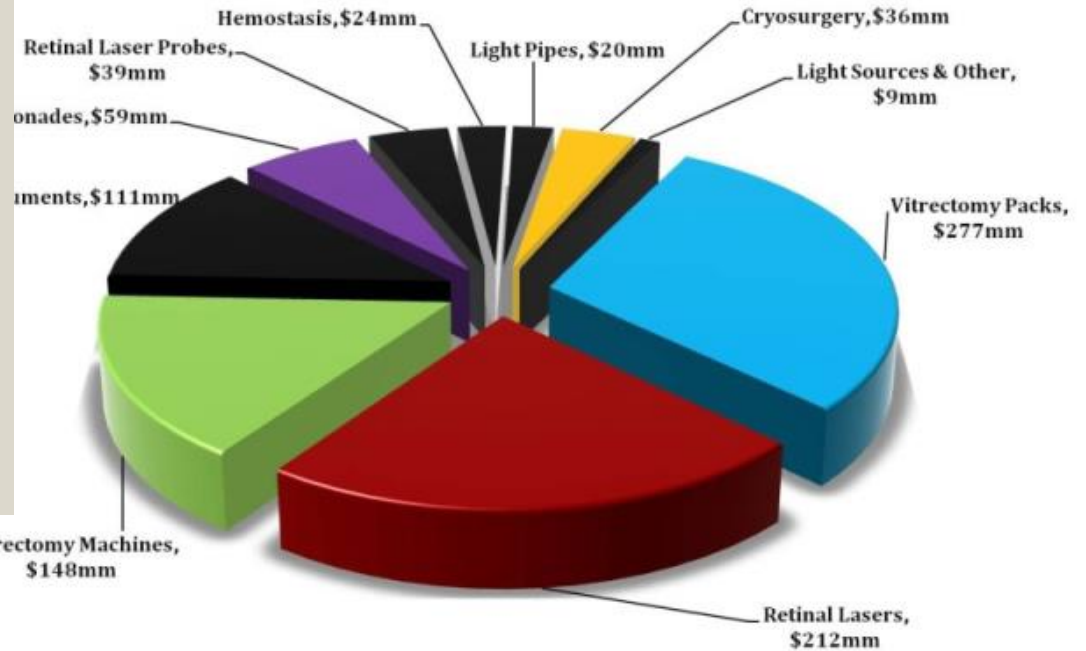
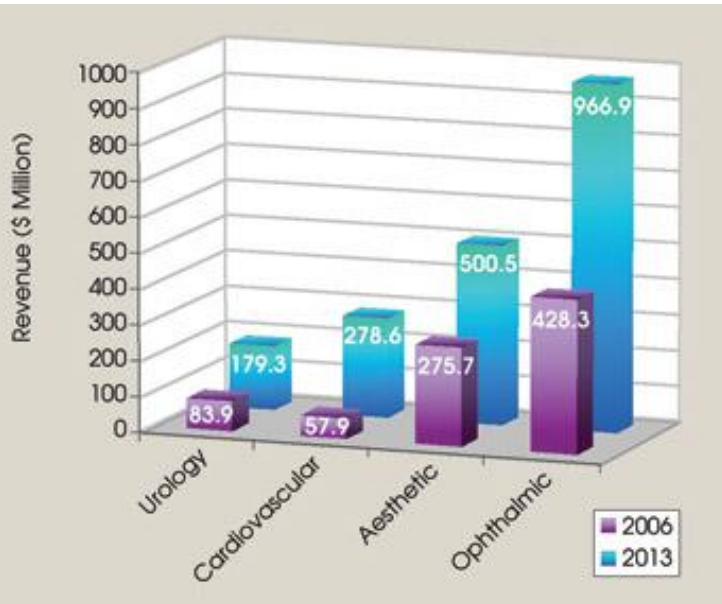


- Suitable for selective treatment of tissue with minimal side effects
- Reliable and stable laser beam
- low cost and compact
- High MTBF / easy plug and play (splice and play)
- Flexible in wavelength
- Scalable

Medical Laser Market

2011 Global Retinal Surgery Device Market

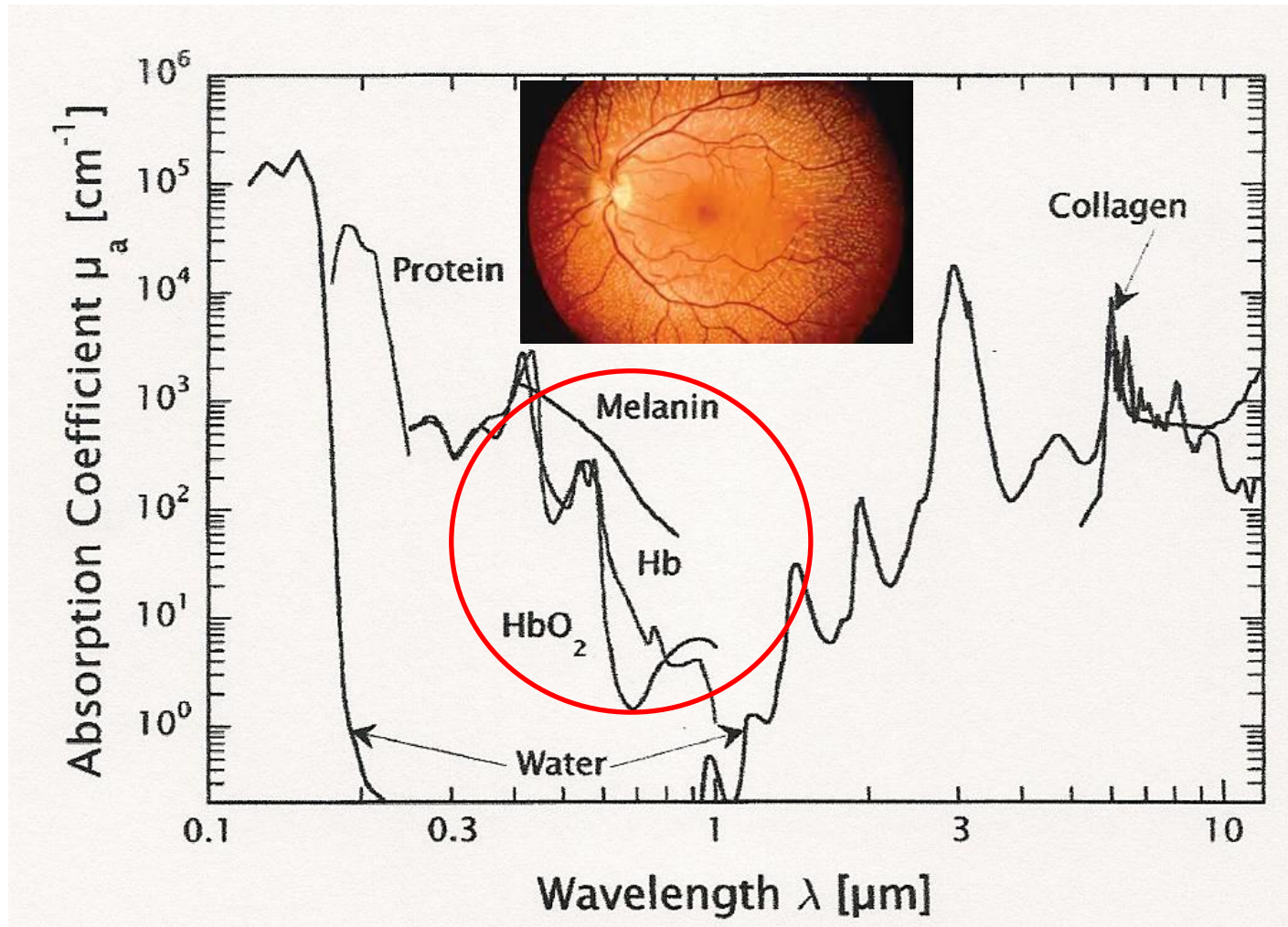
Market Size = \$935 mm
Annual Rate of Growth* ~7%



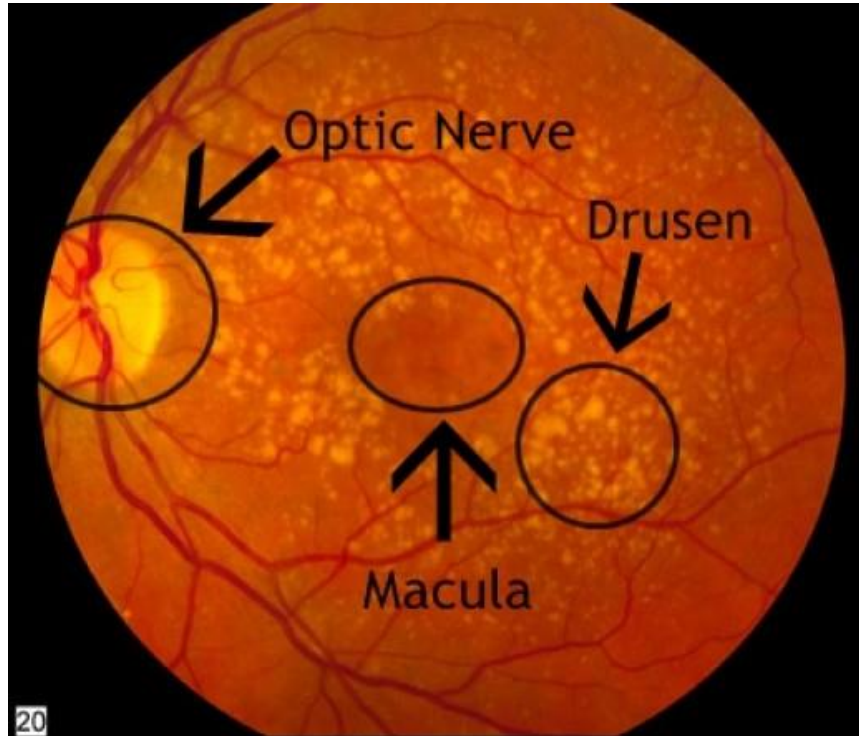
*Source: Synergetics USA Form10-Q for period end April 30, 2012. We estimate that the vitreoretinal market grew approximately 7 percent to \$935 million in 2011, as compared to 2010.

Medical Laser Market approx 15% of 9.4 B\$ world Market (2014)

Low Power Optical properties of tissue



Standard treatment: Thermal coagulation of Retina



In the laser spot a whitening of the Tissue is observed

Typical pulse duration: 0.5-1ms

Typical pulse power: 1-3W

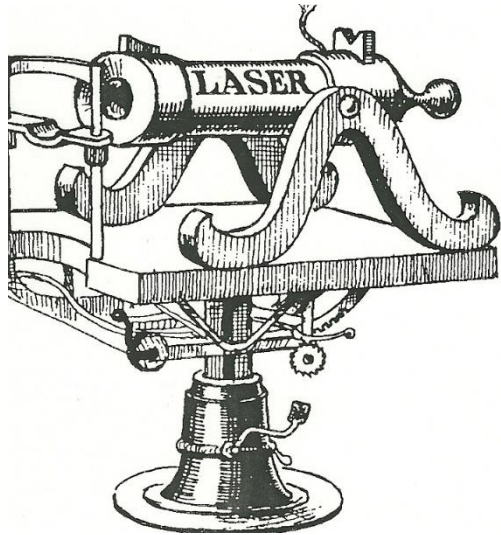
Typical spot size 0.1-0.5mm

Standard laser: Nd:YAG 2f(532nm)

Argon (514nm)

l'Esperance

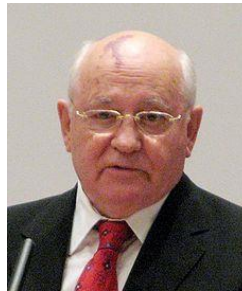
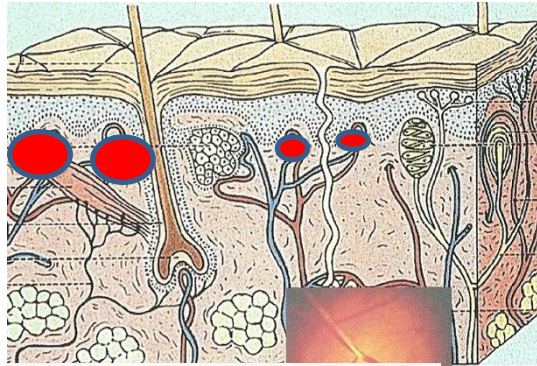
Market requirements for next generation fiber lasers for medical applications



- **Suitable for selective treatment with minimal side effects**
- Reliable and stable
- low cost and compact
- High MTBF / easy plug and play (splice and play)
- Flexible in wavelength
- Scalable

Selektive Laser Treatment

Example: Portwine stain



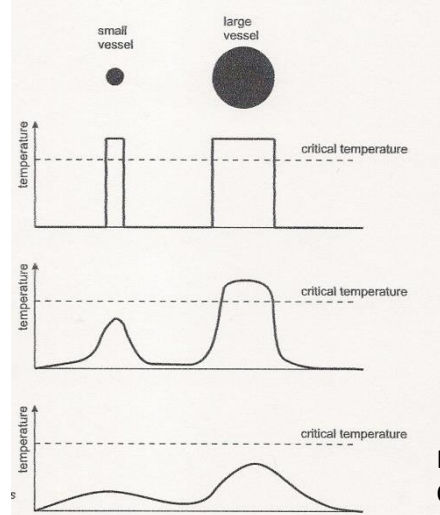
How to achieve selective irradiation:
-right wavelength
-right irradiation time
-right energy density

Heat relaxation time

$$t = d^2/16D$$

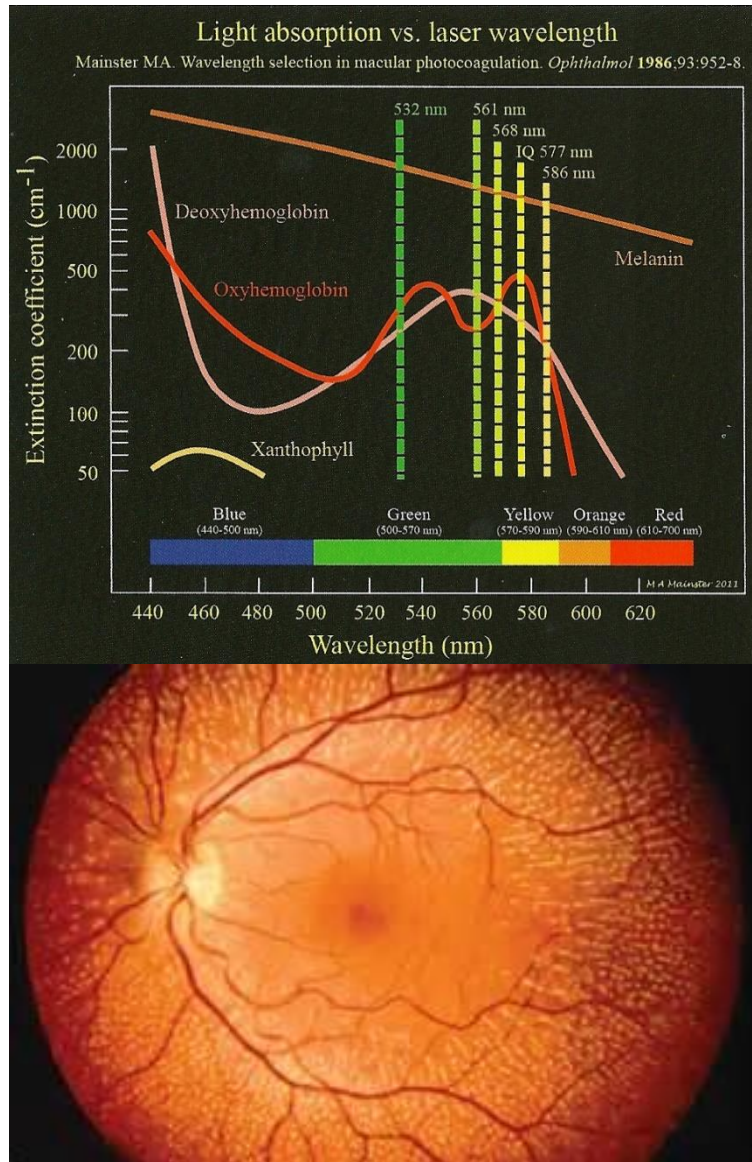
d= diameter of heated cylinder
D= diffusion constant
tissue appr. $10^{-7} \text{m}^2/\text{s}$

Diffusion velocity in tissue: $0.7 \mu\text{m}$ in $1 \mu\text{s}$



B.Cox 2013
Optics in Medicine

Medical Markets: advantage yellow radiation

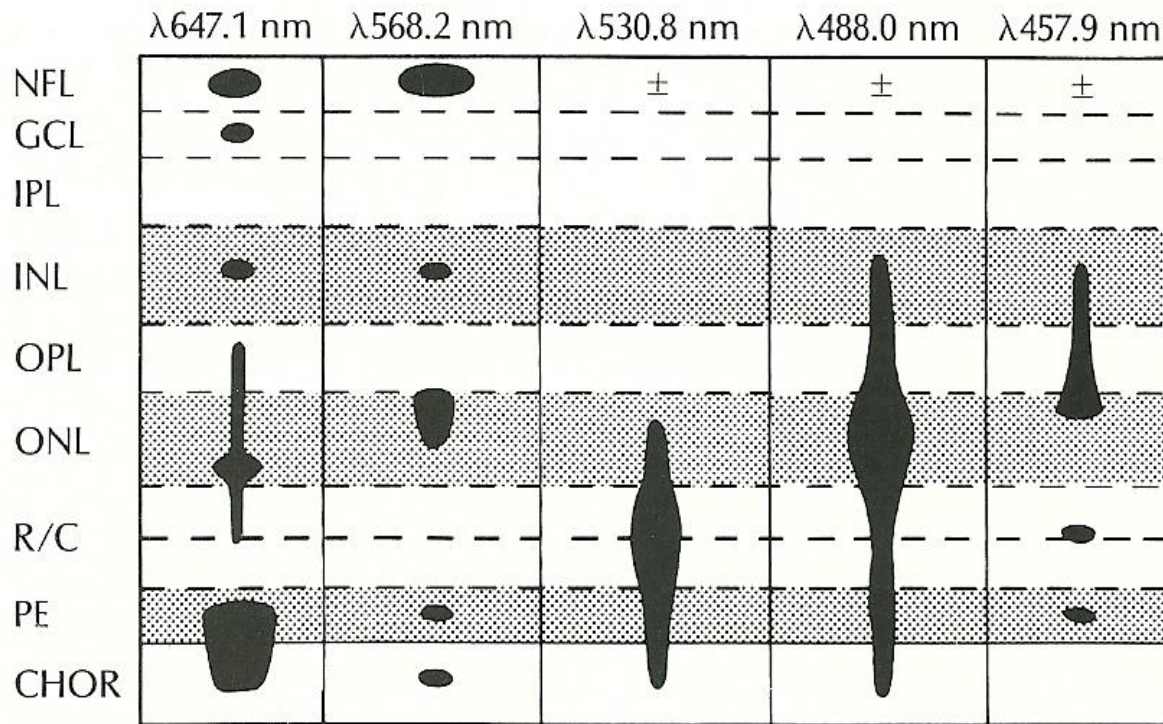


Advantages yellow:

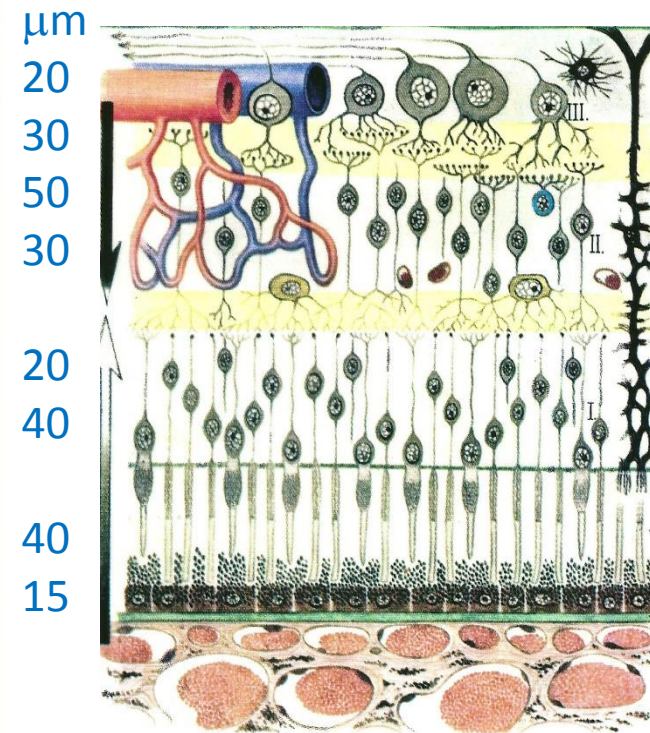
- less scattering in ocular media and aging lenses
- no absorption of Xanthophyll and therefore treatment close to macula possible
- highest ratio HbO/Melanin optimal for vascular structures
- low scattering in retinal layers leads to smaller volumes, needs less power, and less side effects but requires shorter pulses
- high choriocapillaris absorption leads to more uniform effects in irregular fundus pigmentation
- less scarring
- less discomfort of patient

Yadav et al, Ind. J. Ophthalmol. 62(1), 2014

Optical properties of retina: Local Selective Absorption

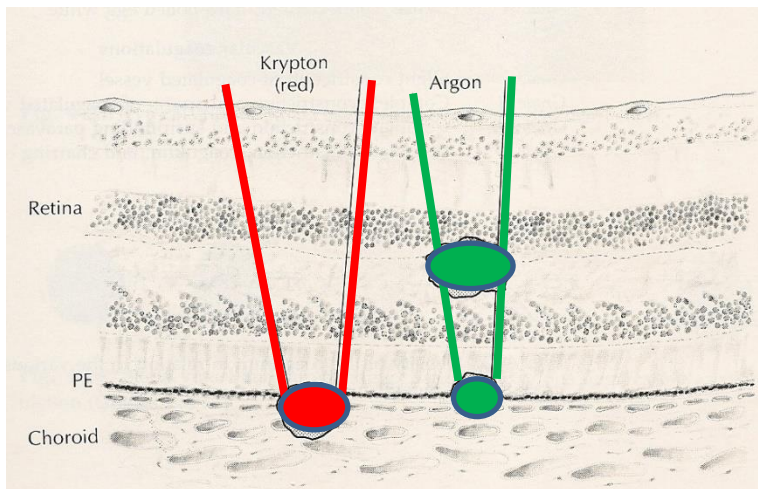


1989 I'Esperance (with Dye laser)

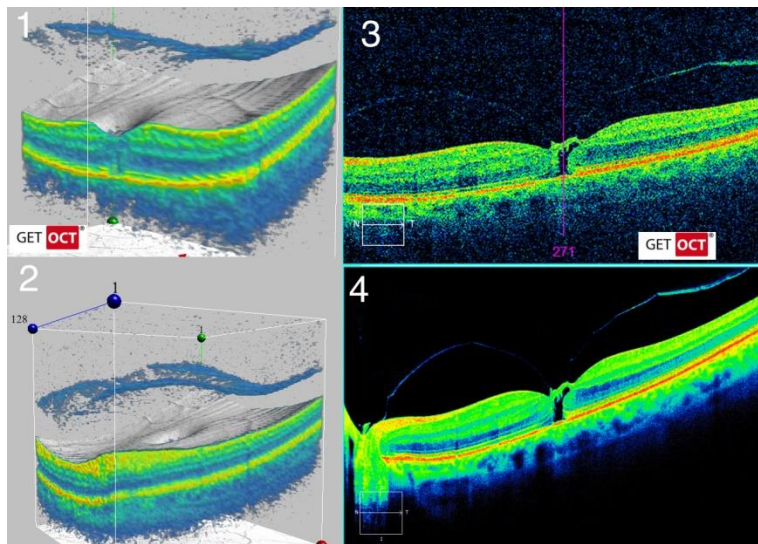


(Thiel)

Selektive «Subthreshold» Laser Treatment



Îesperance(1989)

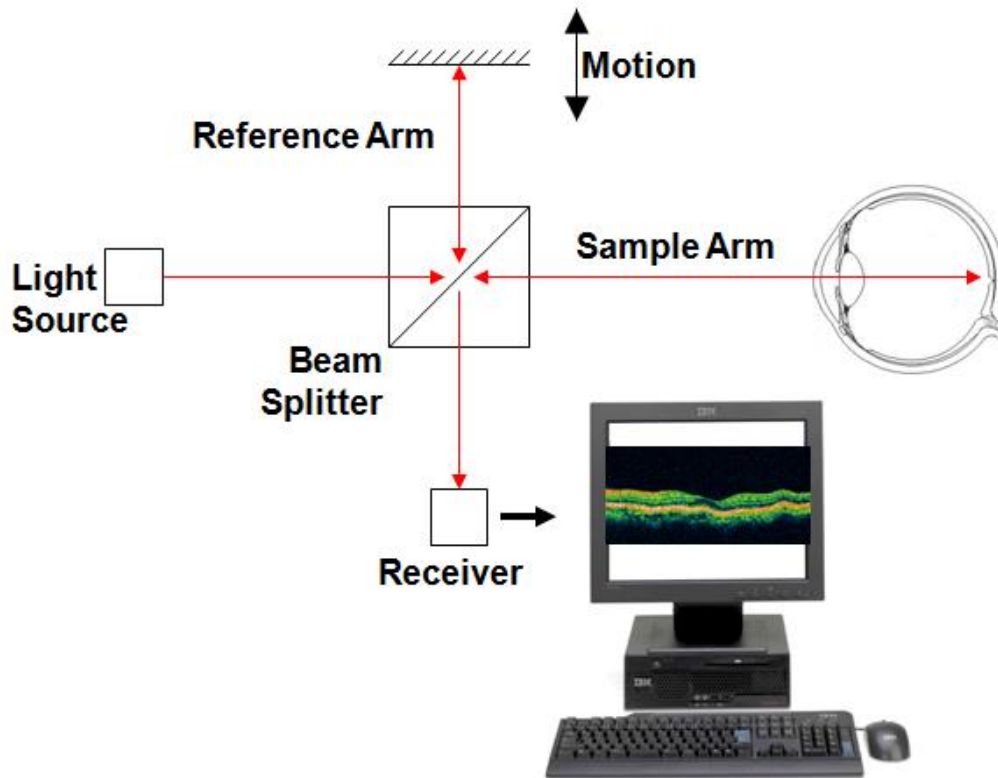


(Google Pictures)

Effect not visible with microscope

Solution:

OCT (Optical Coherence Tomography)

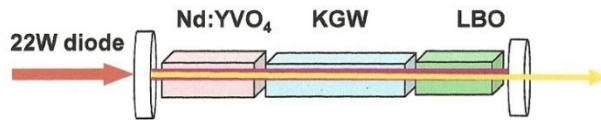


Medical lasers for the yellow-orange spectral region

Raman-DPSSL

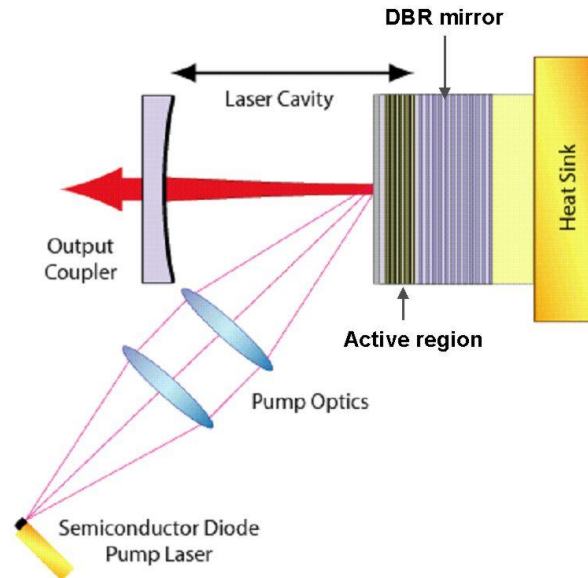
OPDL

FL/Raman-FL



Dekker, Pask and Piper (submitted to *Optics Letters*) report 700mW CW output at 588nm by intracavity SHG of 1196nm 1st -Stokes of KGW pumped intracavity by 1064nm from diode-pumped Nd:YAG, at diode-yellow efficiency ~5%.

KGW: $\text{KGd}(\text{WO}_4)_2$



e.g. Coherent/Philips



«old» lasers: Ar, Kr, Dye Lasers ; lasers to come: Diode lasers

Fiberlaser for the yellow-orange spectral region

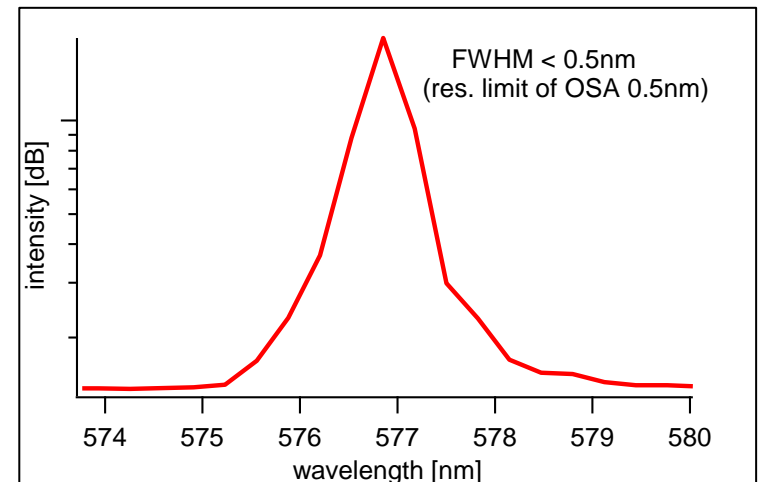
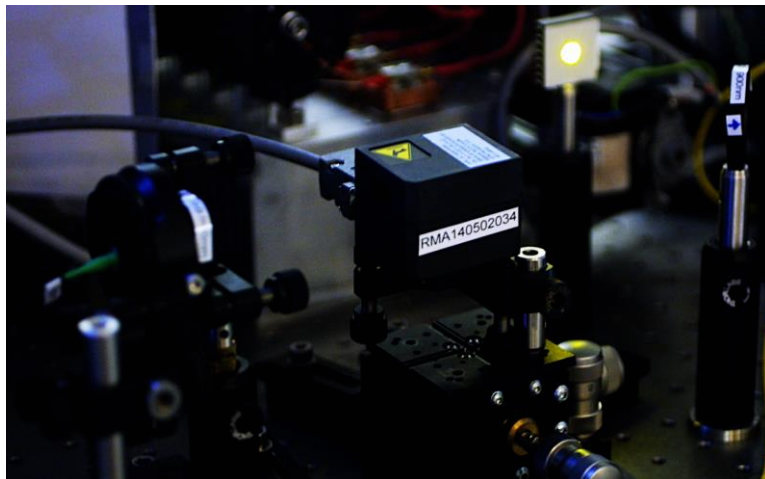
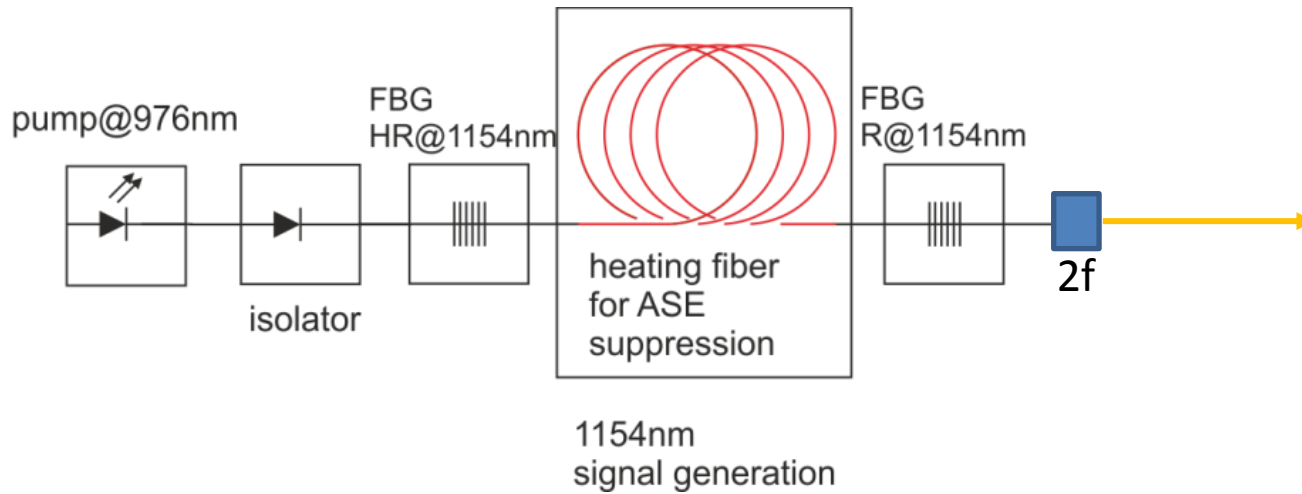
Raman-DPSSL
Diode Pumped Solid State

OPDL
Optically Pumped Diode/VECSEL

FL/Raman-FL
Fiberlaser

- What is the problem?
 - 1154 nm light required to frequency double it to 577 nm
 - Lasing of ytterbium doped fibers occur preferentially at 1030 nm ($2f = 515\text{nm}$)
- Possible solutions:
 - Yb fiber laser at 1100 nm and subsequent Raman shift to 1154 nm
 - Yb fiber laser at 1100 nm with Raman fiber cavity at 1154 nm
 - Yb fiber laser at 1154 nm with heated fiber

Yellow Fiberlaser (Romano, Ryser et al SNFL)



First Results: Combination of A-Scan OCT with 577nm Laser Beam

Steiner, Kowal et al

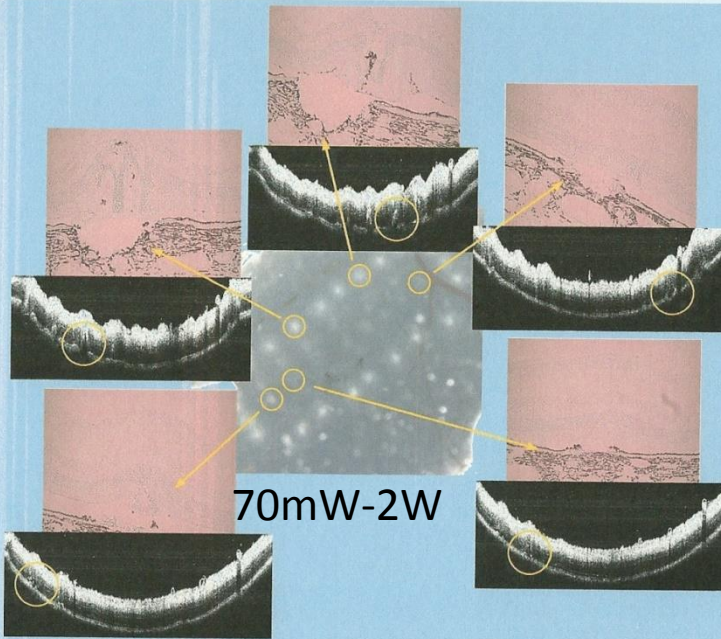
OCT Aufnahmen mit Histologie

Sample 7

u^b

UNIVERSITÄT
BERN

ARTORG CENTER
BIOMEDICAL ENGINEERING RESEARCH



17.03.14

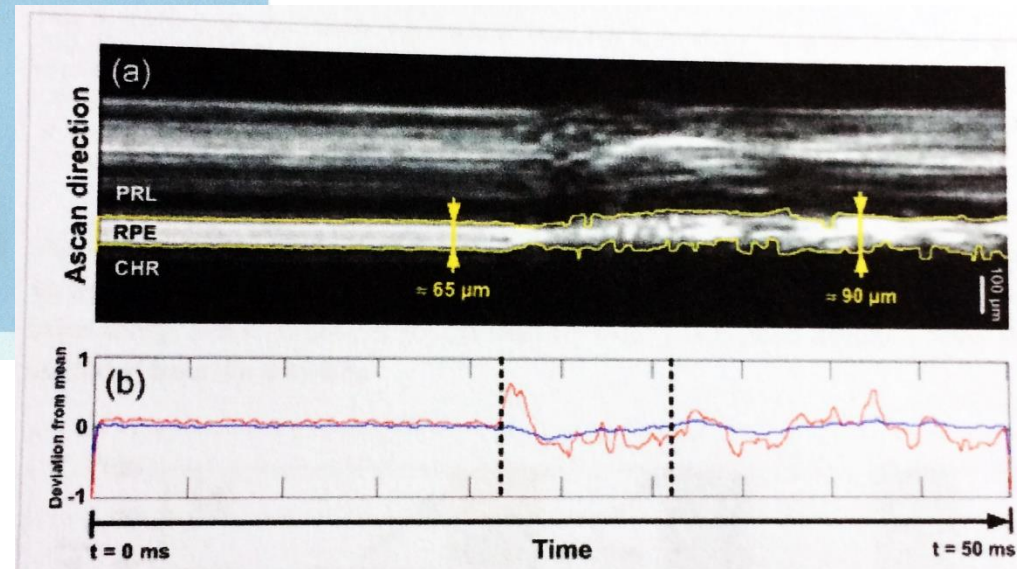
M scan

A scan 60kHz

Resol. approx $2\mu\text{m}$

Integ.time $14\mu\text{s}$

Steiner et al, Phot. Journ. IEEE Dec 2014



Advantages of FL

Performance

Stable SM laser

Efficient Frequency Doubling

Flexibility

Power scalable (limits are ASE and nl-effects in SM Fiber-improvement: large core fibers)

Simple change of wavelength

Combiner and other fiber modules available on market (increasing component market)
(e.g. integration of fiber sensors)

Repair by Splicing

Investment Costs/Cost of ownership

Low costs of fibers (Bragg grating mask)

Compact, modular assembly

Lifetime, MTBF

Vision: «All in Fiber» Ophthalmic Treatment system

(12) Patent Application Publication (10) Pub. No.: US 2008/0058780 A1
 Vogler (43) Pub. Date: Mar. 6, 2008

(54) LASER SYSTEM FOR REFRACTIVE SURGERY

(30) Foreign Application Priority Data
 Aug. 7, 2006 (EP)..... 06 016 465.4

(75) Inventor: Klaus Vogler, Eckental (DE)

Publication Classification

Correspondence Address:
 J. Andrew Lowes
 Attorney for Applicants
 Haynes and Boone, LLP
 901 Main Street, Suite 3100
 Dallas, TX 75202-3789 (US)

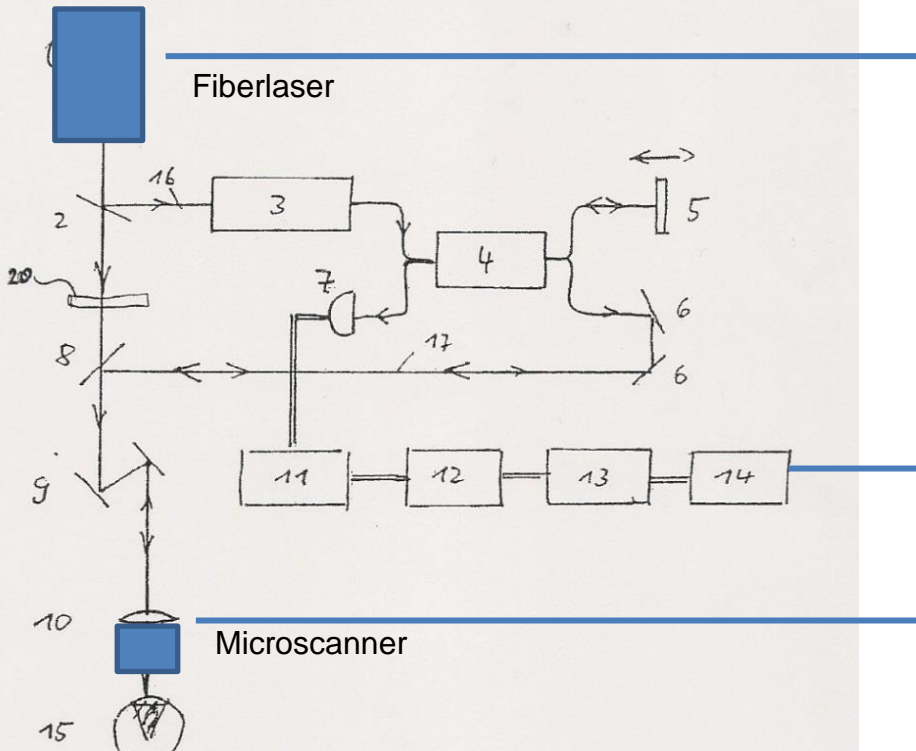
(51) Int. Cl. A61F 9/008 (2006.01)
 (52) U.S. Cl. 606/5

(57) ABSTRACT
 A laser system for refractive surgery comprises a laser beam for generating laser beam pulses and optical means for directing these laser beam pulses as a working beam onto an eye. Some of the working radiation is extracted for optical coherence tomography, in order to measure geometrical structures in the cornea.

(73) Assignee: WaveLight AG, Erlangen (DE)

(21) Appl. No.: 11/835,283

(22) Filed: Aug. 7, 2007

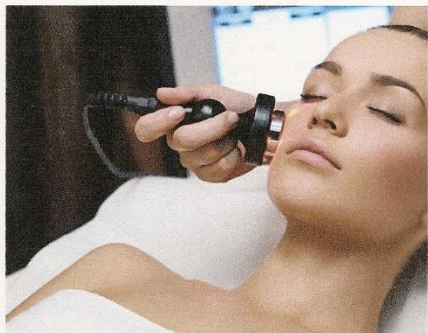
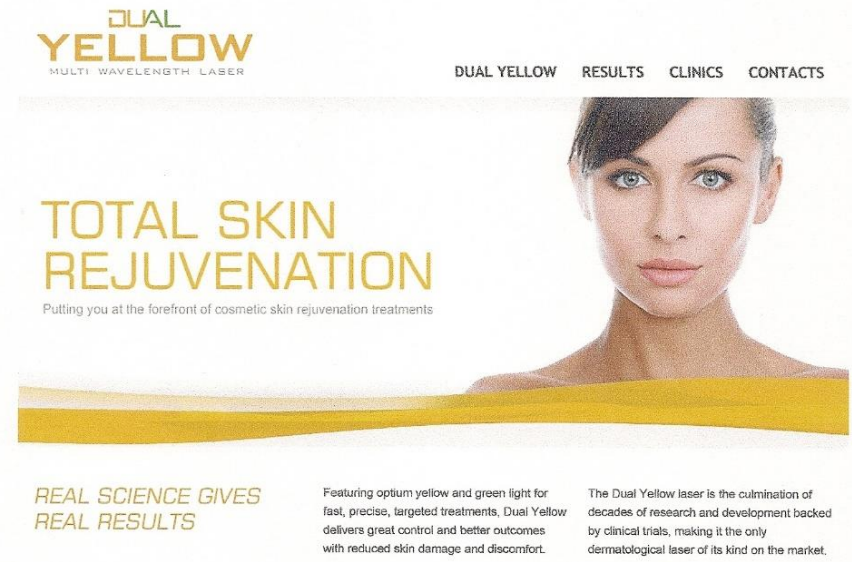


Offline workstation for identifying and Marking of laser treatment Sites in OCT-Tomogram
 Transfer of position to scanner and release of laser pulse and online control

Future: Combination of FL and F-Sensors(OCT) in Dermatology for selective Treatments



DermaLumics(2015)



Fraxel™ Re:Store Dual Laser

Dual FL
1550nm(Er-FL)/
1972nm(Th-FL)

At the moment DPSSL

611nm(DPSSL)

578nm(DPSSL)

But potential for FL

Possible Support of a small company by SNFL

Is a small Swiss company able to develop FL for medical niche applications and to produce sustainable without larger investments costs?

Possible support by SNFL

A) Research and pre-development:

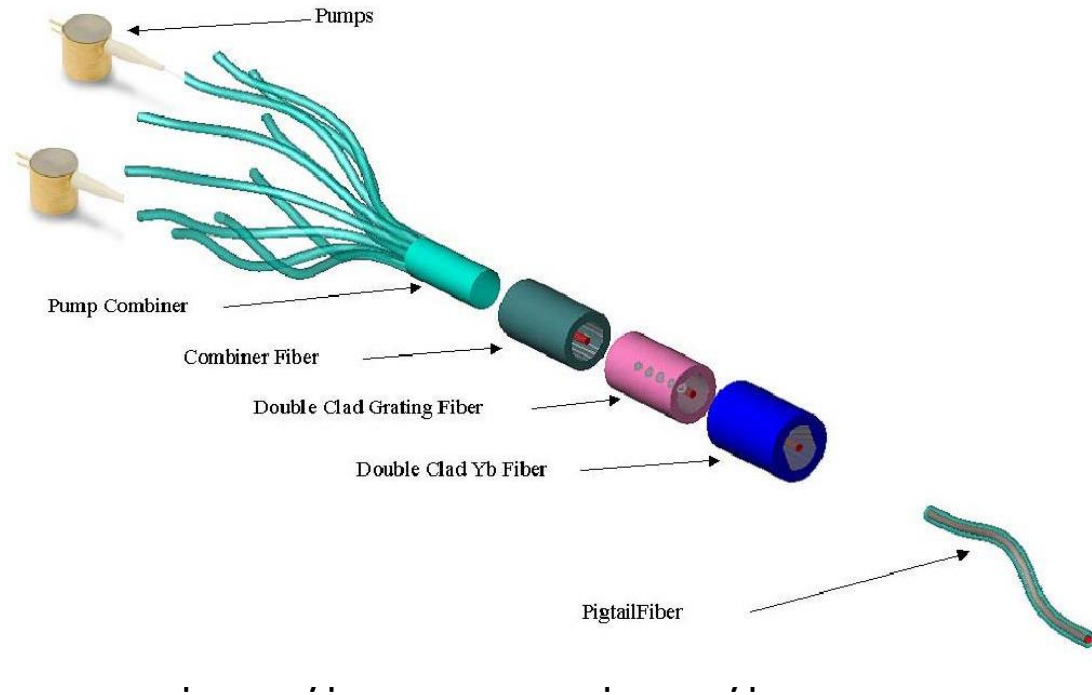
- Identification of solutions with low cost components and modules on the market
- consideration of MP requirements,
- ability to finish/optimize fibers

B) Product development

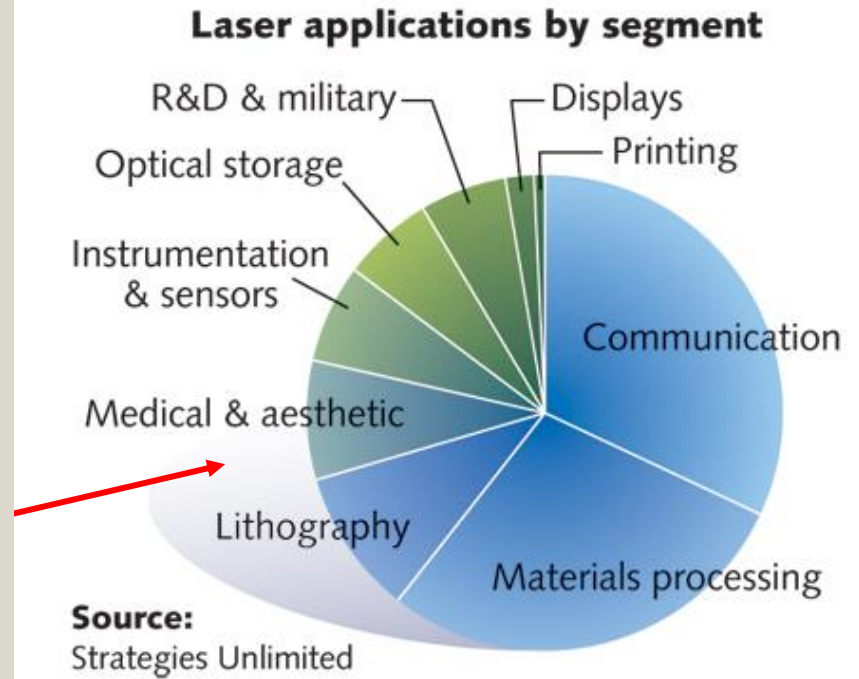
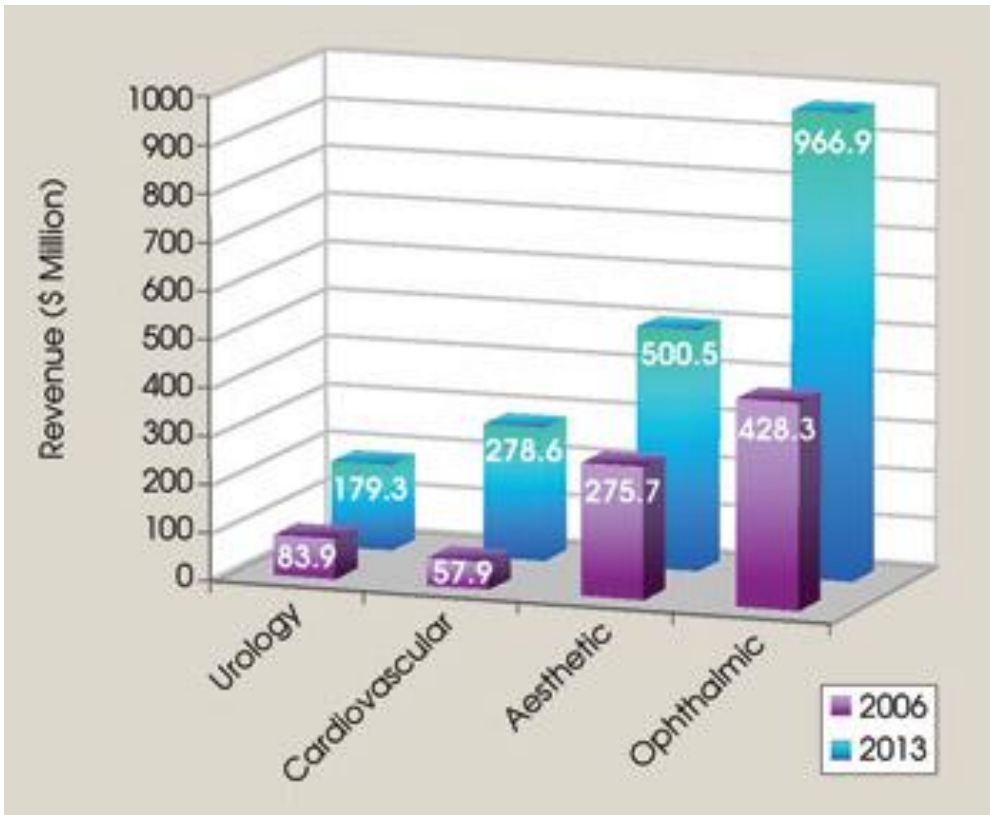
Technical support for trans

C) Production of small number of units

Possibility to share expensive tools (e.g. Splicing)



Medical Laser Market

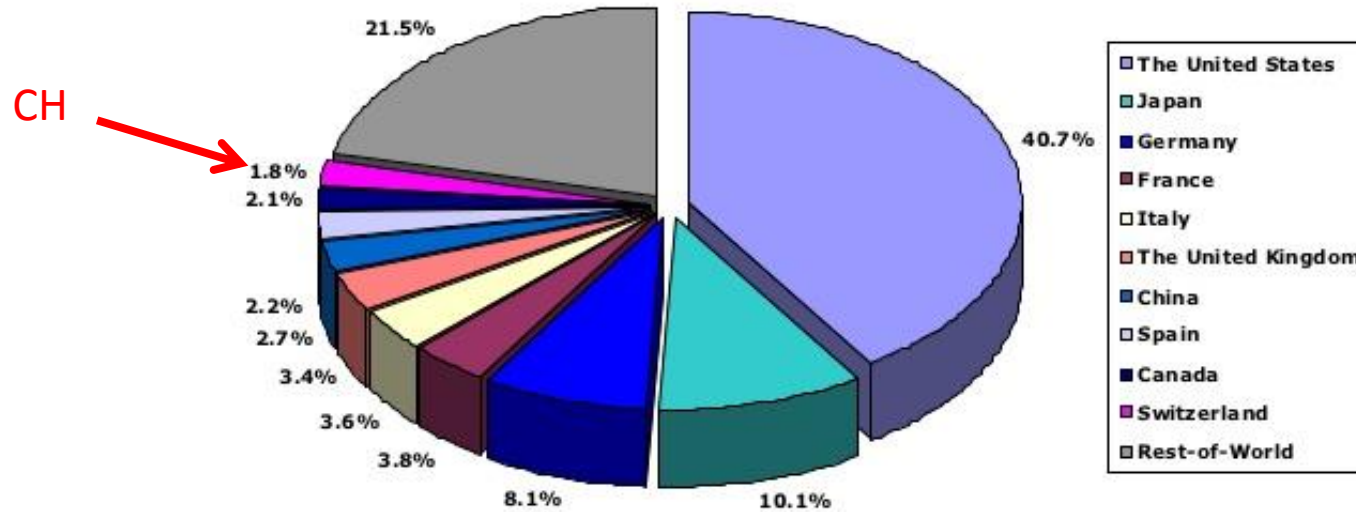


2014 approx 9.4 B \$

Source: Laser Market Place 2014

2010 Medical Device Market Regional Analysis

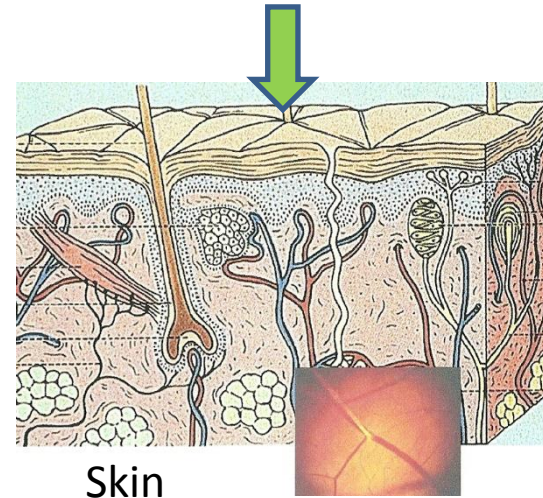
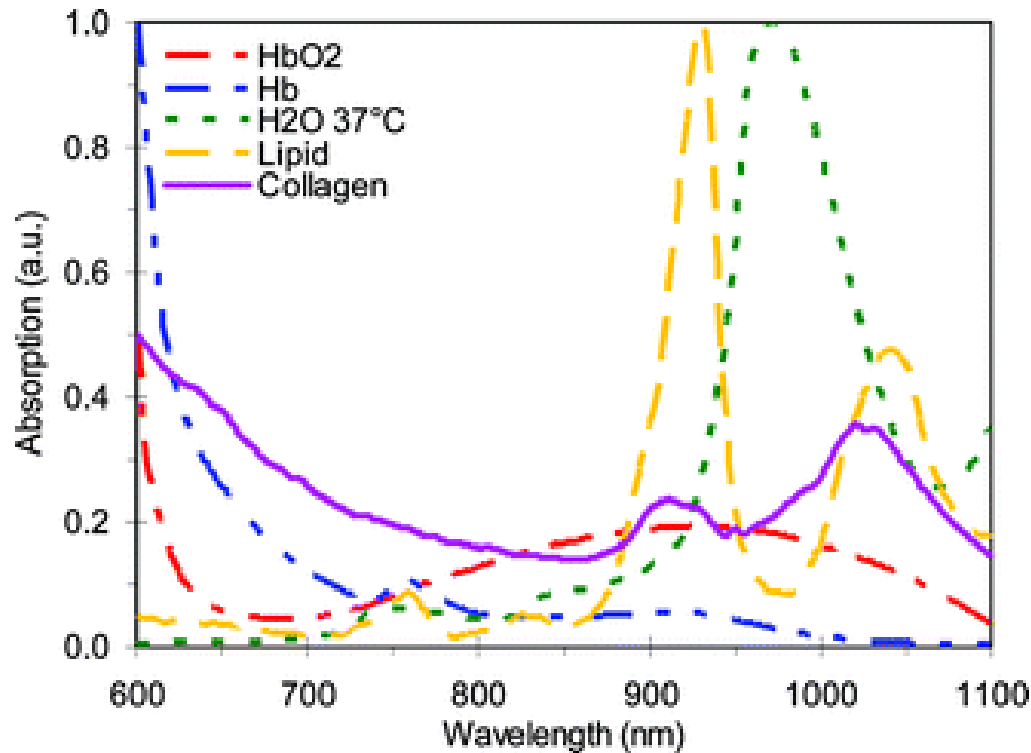
Medical Device Market: Percent Split of Revenues by Geographical Region (World), 2010



Source: Frost & Sullivan

The majority of medical device manufacturers' markets are set on a global basis and manufacturers are seeking expansion into new regions, such as Asia, Eastern Europe and Latin America, in order to achieve greater profits. In Europe, local companies usually have subsidiaries, direct sales offices or distributors in several European countries. Non-European companies tend to establish a head office in Europe and direct sales offices in major markets. The intensive coverage of the European sales territory has resulted in strong competition in all regional markets. Accordingly, local manufacturers do not have monopolies in their own countries, and smaller companies are forced to market their products on an international level.

Medical Markets: example skin



Skin

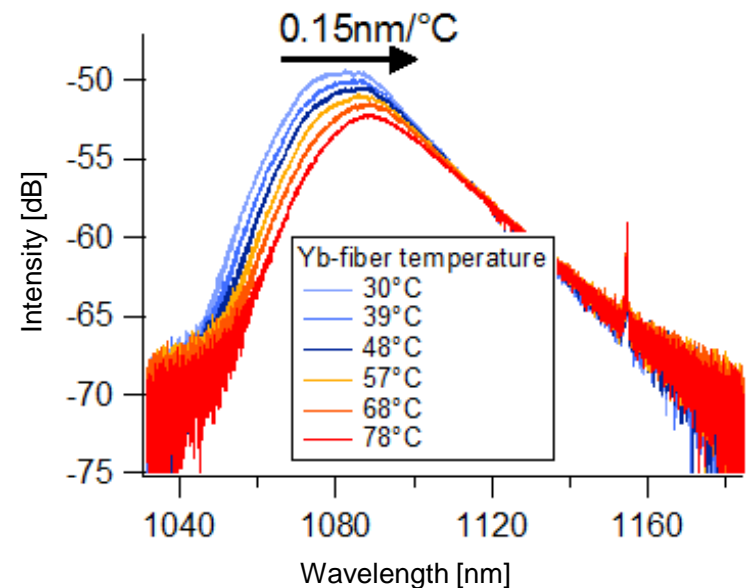
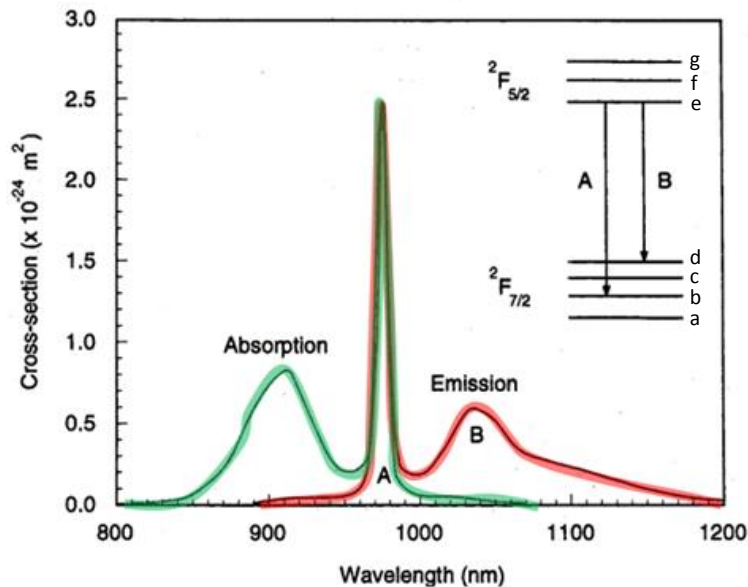
Biological results of (Laser) Irradiation

Temperature	37-60 °C	60-65 °C	90-100 °C	some 100 °C	some 1000 °C	
Effect	Heat / increased diffusion	Denaturisation Coagulation	Phase Change	carbonisation	vaporisation Verbrennung	Shockwave plasma
Optical change	Scattering refraction reversible	whitening Scattering Destruction of structures	shrinking Scattering	Darkening Increased Absorption.	Gases Vapor	Vapor debrises
Biological result	Stimulation	Damage	Shrinking	heavy Damage	mechanical Damage	

How to operate a cavity at 1154 nm

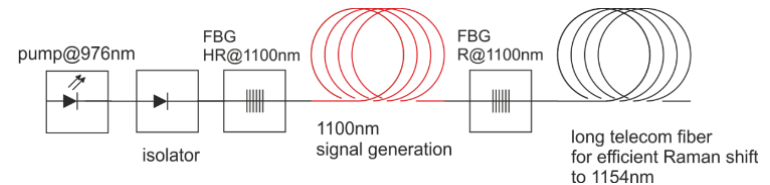
Romano, Ryser et al(SNFL)

- If an Yb-doped fiber is heated, the lowest energy levels «a» and «e» are less populated
- This causes a shift in the emission spectrum towards higher wavelengths
- This allows to run a cavity directly at 1154 nm

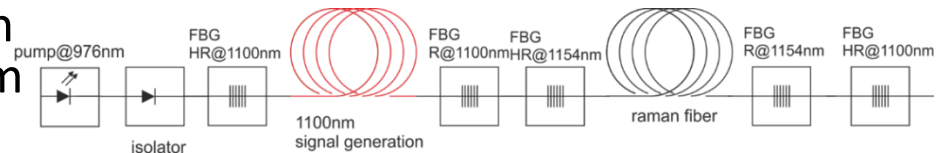


Yellow laser

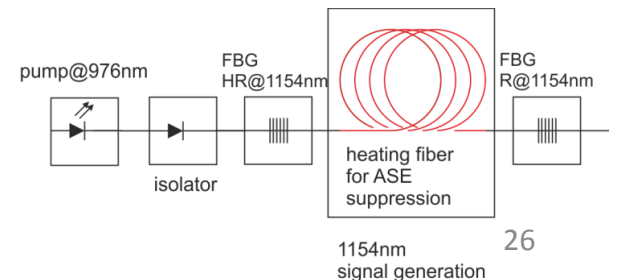
- Yellow laser at a wavelength of 577 nm hardly exist
- What is the problem?
 - 1154 nm light required to frequency double it to 577 nm
 - Lasing of ytterbium doped fibers occur preferentially at 1030 nm
- Possible solutions:
 - Yb fiber laser at 1100 nm and subsequent Raman shift to 1154 nm



- Yb fiber laser at 1100 nm with Raman fiber cavity at 1154 nm



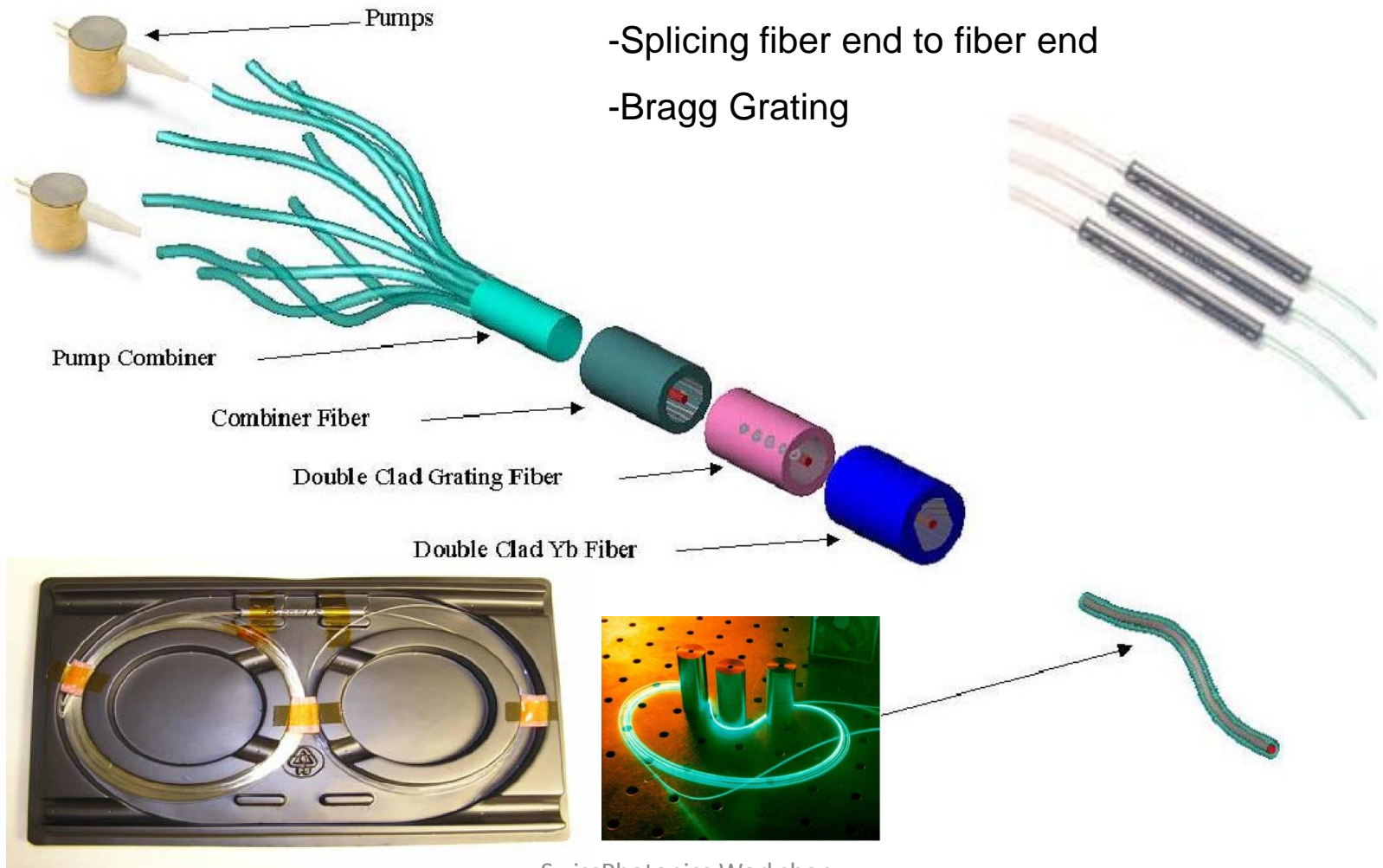
- Yb fiber laser at 1154 nm with heated fiber



Fiber Laser: Principle

Key Technology:

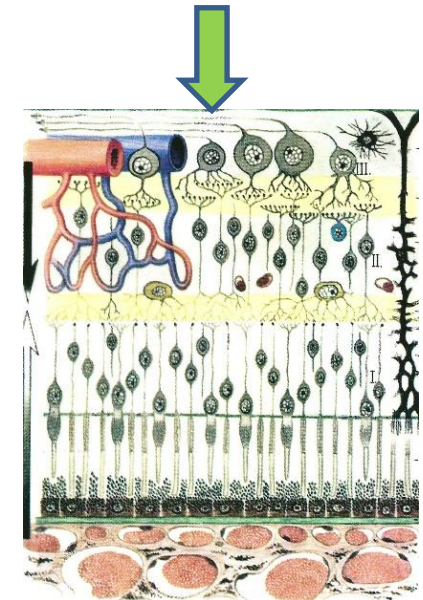
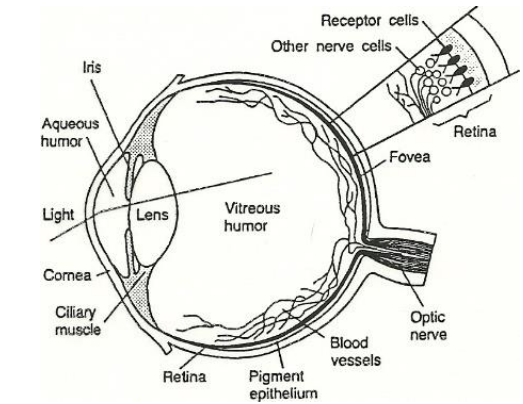
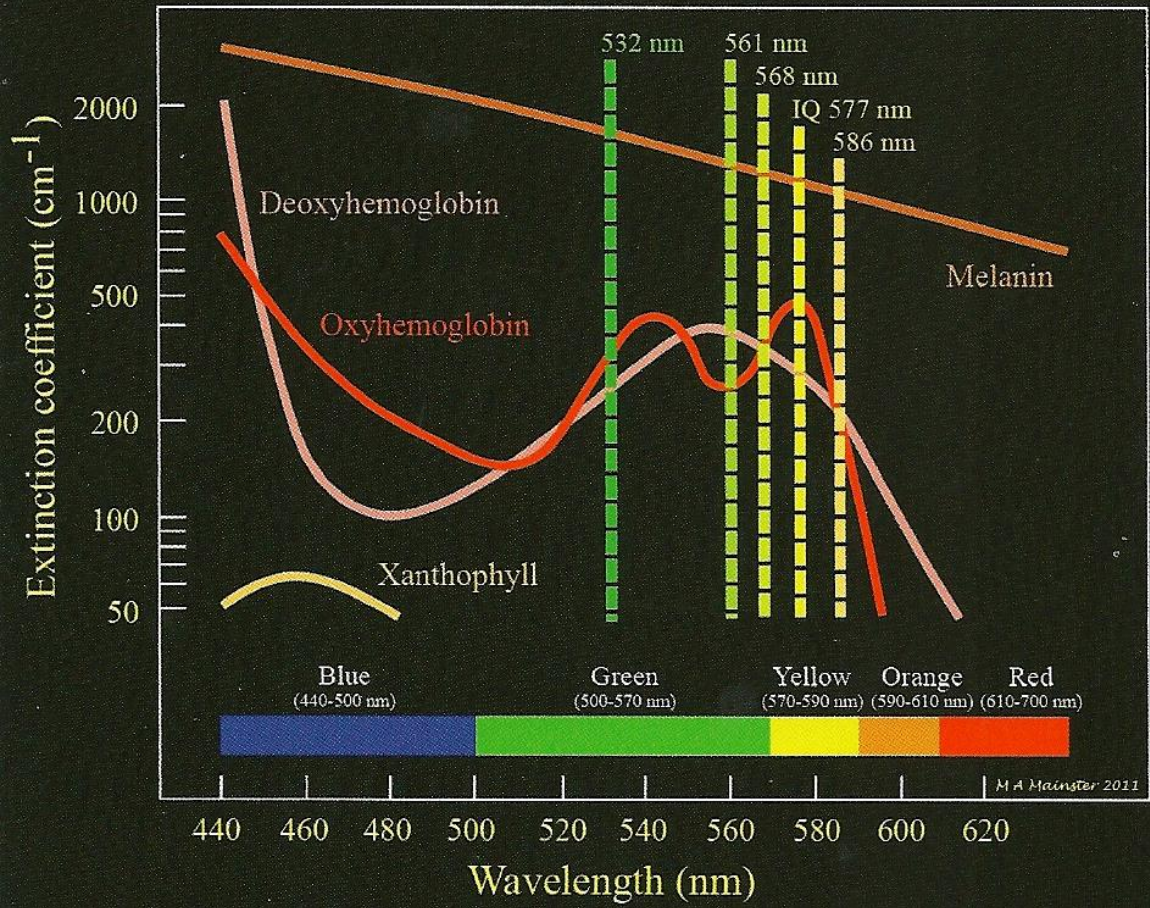
- Splicing fiber end to fiber end
- Bragg Grating



Absorption properties of Retina

Light absorption vs. laser wavelength

Mainster MA. Wavelength selection in macular photocoagulation. *Ophthalmol* 1986;93:952-8.



(Thiel)