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Carcinogenetic and Mutagenic action of 193-nm, 213-nm and 266-nm laser radiation.

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CustomVis Pulzar Z1 Solid State Refractive Laser System



 LASIK, LASEK, Epi-LASIK, PRK, PTK Customized Surgery

- Fast Treatment Times
 - Superior Eye tracking
- Easier to use, faster setup, very reliable.

Paul van Saarloos has a commercial interest in this product

Generation of Laser Wavelengths

Excimer 193 nm: Argon-Fluoride Gas Mixtures

Solid State 213 nm: Nd:YAG laser delivered through Non-linear Crystals for frequency conversion.

Histology

213 nm





193 nm





Absorbance Characteristics

Attenuation through BSS and NaCl at 213nm is significantly less than for 193nm



Absorption Coefficient

Dair, G.T., Ashman, R.A., Eikelboom, R.H., Reinholz, F., van Saarloos, P.P. (2001) Absorption of 193 and 213 nm laser wavelengths in sodium chloride solutions and balanced salt solution. *Archives of Ophthalmology*, *119: 533-537*.

Absorption: Clinical Significance

"The absorbance of 213nm light in BSS & Na Cl is much lower than 193nm"

More reliable, predictable treatment outcomes

Fluctuations in corneal hydration or environmental humidity are unlikely to have a significant effect upon the performance of the solid state laser

Colder, more efficient ablation with less energy wasted heating water.

UV Absorption Spectra of Human Cornea



The absorbance dependence on the wavelength for two human corneal samples between 260nm and 190nm

A "window of ablation" in the far UV region between 220nm-190nm.

Lembares, A., Hu, X., & Kalmus, G. W. (1997). Far ultraviolet absorption spectra of porcine and human corneas. SPIE 2971: 277-786

Cytotoxicity & Mutagenicity

Action spectra for cell inactivation and mutagenesis for bacterial and yeast cells after exposure to radiation in a



vacuum

190- 220nm range less sensitive

Nuclear DNA could be protected from this range of UV light by the surrounding cytoplasmic components

Munakata et al (1986) plus supplementary data (Hieda & Ito, 1986)

In Vitro: Cytotoxicity & Mutagenicity

In vitro study, Kaido et al., (2002) (Coherent Medical) compared mutagenic and cytotoxic potential between 193nm and 213nm

Showed 213nm had greater effect than 193nm

Technical issues with this in vitro study:-

- 1) Cells wet shielded 193nm radiation and not 213nm
- 2) Laser Fluence too low (4mJ/cm²) to dry cells
- 3) 213nm beam contaminated with 266nm radiation.

In vivo DNA Damage Study

300μm ablation of both eyes of fifteen live 1/2 lop rabbits

5 rabbits with 193nm Excimer as -ve control

5 rabbits with 266nm as +ve control
5 rabbits with 213nm

DNA Analysis

- Technique by Nuss, Puliafito and Dehm to detect unscheduled DNA synthesis (UDS)
- Enucleated eyes incubated in calf serum with radioactive thyamine into DNA for 3 days
- Fixed, embedded into 2-3mm sections
- Dipped into photographic emulsion, left for 2 weeks, fixed and stained
- All cells, partially labeled cells (disrupted DNA replication) and heavily labeled cells (normal cell division) counted

266nm Ablation

Partially labeled cells



Percentage of cells with Unscheduled DNA Synthesis



DNA Damage *In vivo* study: Unscheduled DNA Synthesis (UDS)

Positive Control

266nm: significantly more DNA damage.

For 193 & 213 nm: < 5% (average) of cells were affected.



266nm

<u>213nm</u>

193 & 213 nm SAME. Both produce Minimal DNA damage.



TUNEL Staining of Rabbit cornea following PRK

213nm 3 days

Ablated Zone

Non-Ablated

193nm 3 days

Ablated Zone

Non-Ablated

Solid State Lasers May Be An Improved Option for Refractive Surgery



Molecular and histological studies reveal similar results to 193nm

In vivo mutagenicity studies – 213 as safe as 193

PERFORMANCE ADVANTAGES

- Transmission through Aqueous solutions
- Higher Corneal Ablation Efficiency
- May Produce Less Thermal Effect

THANK YOU!!!!