

Starting spin off companies in at the Hungarian Academy of Sciences in the 90-es: hand in hands

Róbert Szipőcs, PhD

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HEPTECH Board Meeting
Tihany, 30 May, 2024

OUTLINE

1. Background

- Invention of chirped mirrors, patenting at MTA SZFKI (inventors: Robert Szipocs, Ferenc Krausz)
- Nobel Prize in Physics in 2023 – chirped mirrors for attosecond pulse generation

2. Founding spin off companies

- Founding spin off companies at MTA SZFKI and TU Wien, Austria (R&D Lézer-Optika Bt., Stingl OAG)
- Application of chirped mirror technology for science (e.g. at MPI Stuttgart, TU Wien, Uni. Groningen)
- Application of chirped mirror technology for industry (e.g. Spectra-Physics, Coherent, FemtoLasers)
- Founding R&D Ultrafast Lasers Ltd (broadly tunable Ti:sapphire lasers and OPO-s for life science)

3. R&D Ultrafast Lasers as an industrial partner of Wigner RCP

- Development of a 20 MHz repetition rate Ti-sapphire laser at MTA SZFI financed by R&D Ltd.
- FemtoBio project (National Technology Program 2006-2009) with Furukawa Electric (fiber lasers)
- FiberScope project (National Technology Program 2010-2014) with Genetic Immunity (nonlinear microscopy)
- Quantum microscopy (EUREKA Project on Quantum Optics, 2023-2026) with Ulm University (FLIM microscopy)

4. Hand in hands: how technology supports science, how science develops industry

The effect of dispersion in ultrafast laser systems and our solution for the problem

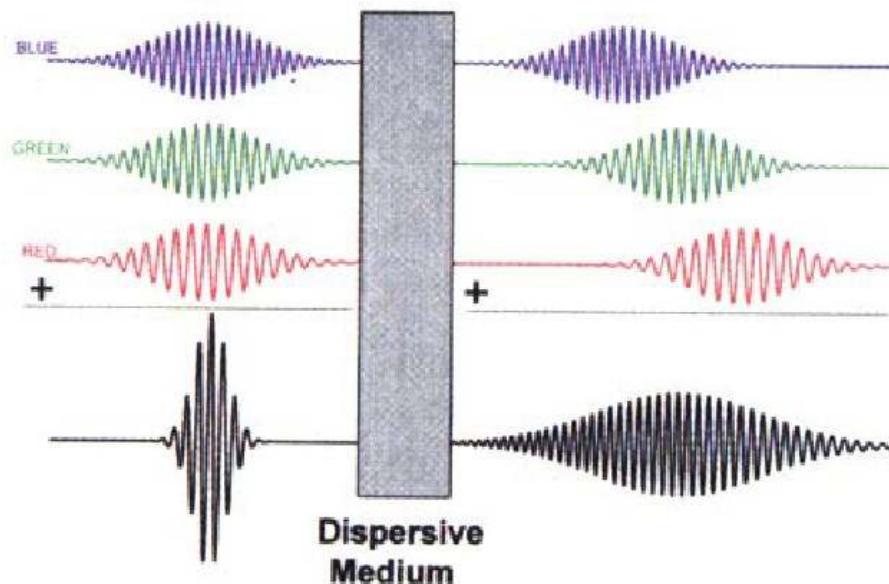
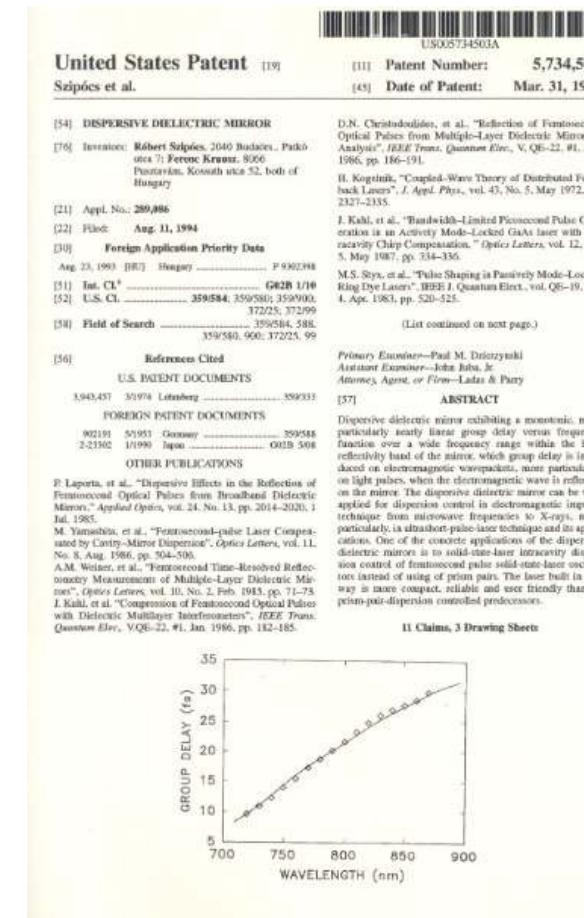


Figure 1. Optical pulse propagation through a dispersive medium: a frequency-dependent group delay leads to a pulse broadening and to a carrier frequency sweep.

"Pushing the Limits of Femtosecond Technology: Chirped Dielectric Mirrors," Optics & Photonics News 6(6), 16- (1995)
 R. Szipőcs, F. Krausz U. S. Pat. No.: 5,734,503



R. Szipőcs, A. Stingl, Ch. Spielmann, and Ferenc Krausz,

"Pushing the Limits of Femtosecond Technology: Chirped Dielectric Mirrors," Optics & Photonics News 6(6), 16- (1995)

The use of ultrabroadband chirped mirrors for attosecond pulse generation

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Ferenc Krausz wins the Nobel Prize!

Founder of attosecond physics wins the most important prize in science

06/08/2023

[Read more from the Institute](#)

Ferenc Krausz, Director at the Max Planck Institute of Quantum Optics and Professor at the Ludwig-Maximilian University of Munich, together with Pierre Agostini, and Anne L'Huillier, has been honoured with the 2023 Nobel Prize in Physics. The Nobel Committee is honouring the three researchers for the foundation of attosecond physics. An attosecond is a billionth of a billionth of a second. Laser pulses lasting only a few attoseconds can be used to track the movements of individual electrons. This provides fundamental insights into the behaviour of electrons in atoms, molecules and solids, but could also help to drastically speed up today's electronics.

In 2001, Ferenc Krausz generated light pulses in the attosecond range (1 attosecond – 10-18 seconds) for the first time. Their use for observing electron movements in atoms was honoured by Nature and Science as one of the 10 most important scientific achievements of 2002.

The basis for this was laid by Ferenc Krausz and his compatriot Robert Szöpös with the development of mirrors with which extremely intense laser pulses can be generated from a few oscillations of a light wave. In 2002, Ferenc Krausz and Theodor Hänsch, who is also director at the Max Planck Institute of Quantum Optics and professor at the LMU, succeeded in controlling not only the intensity of light pulses but also the phase, i.e. the exact course of a light wave, using the Ted Hänsch's frequency comb technique, which also won a Nobel Prize - in 2005.

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Márai (über Ferenc Krausz)

AZ IZOLÁLT ATTOSZEKUNDUMOS IMPULZUSOK ELŐÁLLÍTÁSÁT MEGALAPOZÓ LÉZERFIZIKAI FEJLESZTÉSEK A KILENCVENESEN ÉVEK KÖZEPÉN A BÉCSI MŰSZAKI EGYETEMEN

Szöpös Róbert

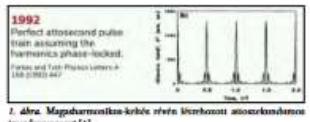
HUN-REN Wigner Fizikai Kutatóközpont, Budapest

Irásmóban a következő pár oldalon azt szeretném bemutatni, hogy az izolált attoszekundumos impulzusok előállítása miatt éppen Krausz Ferencnek sikerült előzör a világban, és hogy ez az eredményt milyen alapvető fontosságú lézerfizikai, lézertechnológiai fejlesztések előzte meg a kilencvenes években – többek között a Bécsi Műszaki Egyetemen, az ószakmai vezetésben.

Ahhoz, hogy az egész fejezetben folyamat megtörténjen és érthetődjön, tanán a legegyszerűbb elszemelésekkel, vagyis ott, hogy egy magas rendű nemlineáris optikai folyamat, a magasharmonikus keltés (IHG) révén hogyan lehet rövid, nagy intenzitású lezertimpulzusokkal attoszekundumos impulzusokat előállítani – például nemegyszázmillió attól függően, hogy a lézerimpulzus hossza hogyan viszonyul annak (esetükben tipikusan 800 nm körülű) középhullámhosszához. A nemlineáris optika rejtélyéhez, azon belül a magasharmonikus-keltés elméleteihez állapíthatunk Domki Péter és Tóth Csaba írásai vezetik le az olvasót ebben a Nobel-díjhoz kapcsolódó körönkből, utóri vonás csak összefoglalja a legfontosabb megoldásokat, törvényszégeket.

Elsőször is, a gázatomok polárisálhatási képessége szintetizálásig megtárt a páratlan sorozatú harmonikus kusok általános elét a IHG során. Másodszor, a legmagasabb sorozatszám (legtöbbet hullámhosszú) harmonikus rendjét az att elérőlétező impulzus maximális térfürdője (E_f) határozza meg, vagyis minden nagyobb a fókuszált térfürdő az atomokban, amik magasabb harmonikusokat tudnak előállítani. A harmadik, és az izolált attoszekundumos impulzusok előállítása szempontjából rendkívül fontos megoldás, hogy a magas harmonikusok minden térfürdőszámmal egyidőben, egymással azonos fizikai tulajdonságokkal, így egy most már vi-

szonylag egyszerűen megírható, 20–30 fs-os Ti-zafir léserverziós impulzusainak nemegyszéba (pl. Kr) történő fókuszálásával esetén egy attoszekundumos impulzussortozatot ($I_{\text{ábra}}$), nem pedig egy időfhamisítás mérésekre jogosult használható, izolált attoszekundumos impulzust kaphatunk.



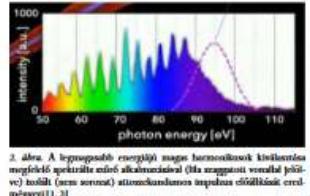
1. ábra: Magasharmonikus-keltés révén izolált attoszekundumos impulzusok [1]

löholt, egyedi attoszekundumos impulzusokat a magasharmonikus-keltés során kizároló olyan lézergörbötők esetén tudunk előállítani, amikor a lézertimpulzust leíró $E(t)$ előfüggvény csak egy, a többször lényegesen magasabb maximummal rendelkezik, és így az előben keletti magas harmonikusok rendje, energiajára lényegesen megnöveljük a többször maximumban keletti vonatkozó értékeit. Ebben az esetben, ha megfelelő spektrális szűrő alkalmazva ($I_{\text{ábra}}$) kiválasztja a legmagasabb energiával rendelkező XIV. (komány ultrahalvány) komponensét, akkor ezek egyedül, attoszekundumos lézertimpulzusokként jelennek meg a színes után [1, 2].

A magasharmonikus-keltéshez használt, nagy ener-

giájú lézertimpulzusnak tünetek – izolált attoszekundumos impulzusok előállítások – praktikusan kör fel-

tételeknél kell megfelelnie: az első, hogy a lézertimpul-



2. ábra: A legmagasabb energiájú maga harmonikusok kiválasztása megfelelő spektrális szűrő alkalmazásával (ha szaggatott vonalat jelöljük) [3]



Szöpös Róbert, PhD, okleveles végzettségről (EMI, MSc), kísérleti (SZTE, PhD), a HUN-REN Wigner Fizikai Kutatóközpont továbbtanulmányos, az R&D Üzlet vezetője. Kitüntetései: 2002-ben Ferenc Krausznal, a fentemelkedőként ismertetett először a lézertimpulzusokról szóló eredményekről kizárt Németország Céh Díja, 2005-ben Ted Hänsch díja a MTA Fizikai Tudományos Oktatási Díjak capa.

PAST: INVENTING CHIRPED MIRRORS IN 1993 (MTA SZFKI / TU Wien) THE SOLUTION FOR ULTRAFAST SOLID STATE LASERS

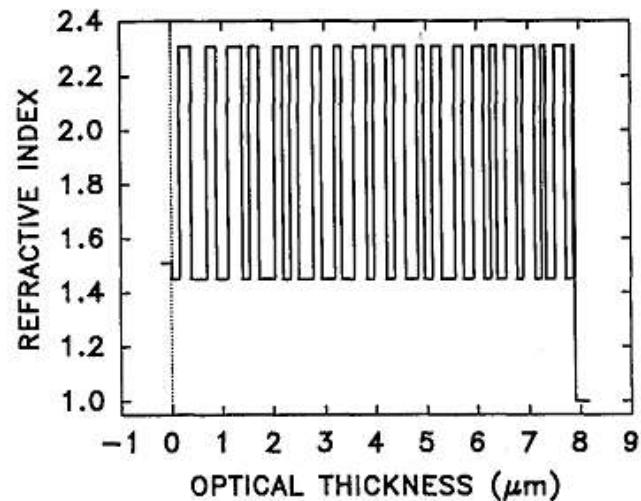


Fig. 1. Theoretical refractive-index profile of a high-reflectivity $\text{TiO}_2\text{-SiO}_2$ multilayer coating designed specifically for broadband GDD control in femtosecond lasers.

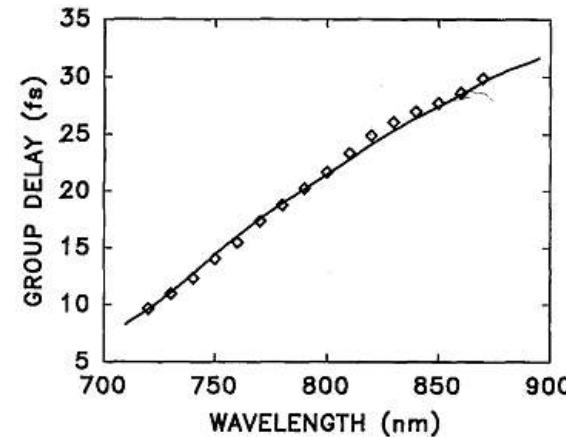
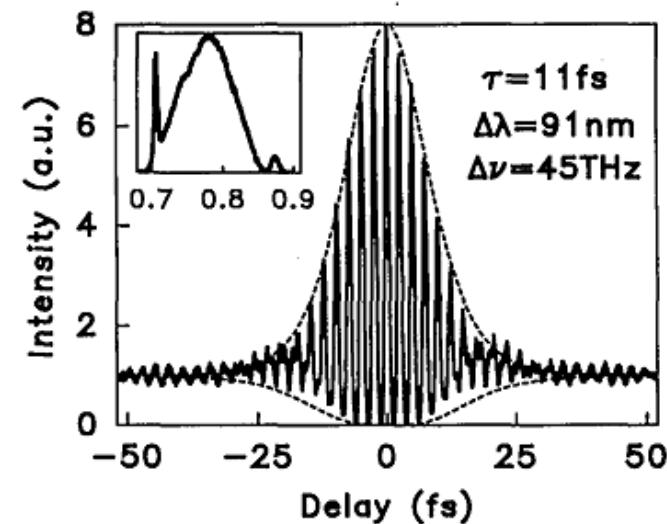
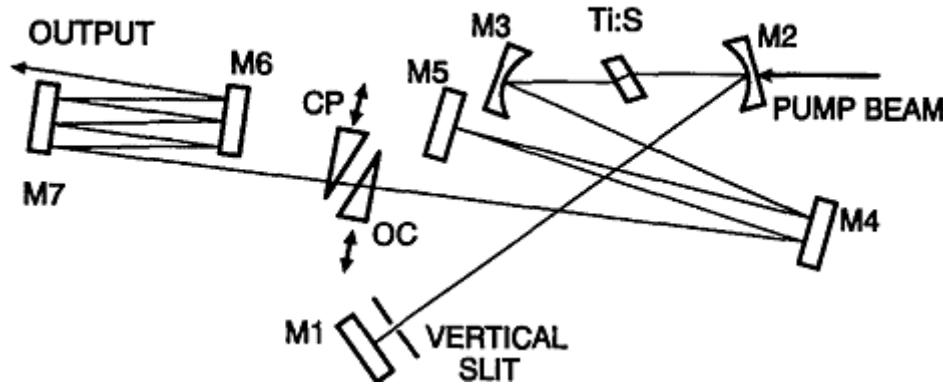


Fig. 3. Computed group delay as a function of wavelength (solid curve) together with experimental data (squares) for the multilayer design of Fig. 1. Note that the absolute delay could not be measured; therefore a wavelength-independent constant delay was added to the measured relative data.

R. Szipőcs, K. Ferencz, Ch. Spielmann, F. Krausz, *Opt. Lett.* 19, pp. 201-203 (1994)
R. Szipőcs, F. Krausz: Dispersive dielectric mirror; U. S. Pat. No.: 5,734,503 (1993)

MIRROR DISPERSION CONTROLLED Ti:SAPPHIRE LASER

Founding Stingl OAG in Vienna by Ferenc Krausz



LINEAR CAVITY

- ☺ Highly stable femtosecond pulses with duration of ~ 11fs

A. Stingl, Ch. Spielmann, F. Krausz, R. Szipőcs, *Opt. Lett.* 19, pp. 204-206 (1994)

R. Szipőcs, F. Krausz: U. S. Pat. No.: 5,734,503 (1993)

Development of a Ti:sapphire oscillator + CPA system at TU Wien using chirped mirrors for generating sub-mJ, ~18 fs pulses for pulse compression in a hollow fibre filled with noble gases (Ar/Kr)

> ULTRAFAST LASERS

High-quality seed pulses from mirror-dispersion-controlled Ti:sapphire system allow chirped pulse amplification without a pulse stretcher.

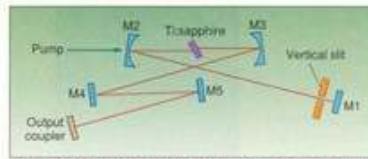


FIGURE 1. Femtosecond Ti:sapphire laser uses chirped dispersive mirrors (M1-M5) for intracavity dispersion compensation.

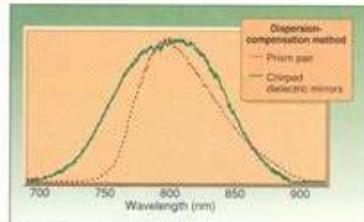


FIGURE 3. Comparison of spectra from prism-controlled and chirped-mirror Ti:sapphire oscillators reveals enhanced performance of chirped-mirror system, which yields a more symmetric pulse.

Chirped dielectric mirrors improve Ti:sapphire lasers

Ch. Spielmann, M. Lenzner, F. Krausz, R. Szipőcs, and K. Ferencz

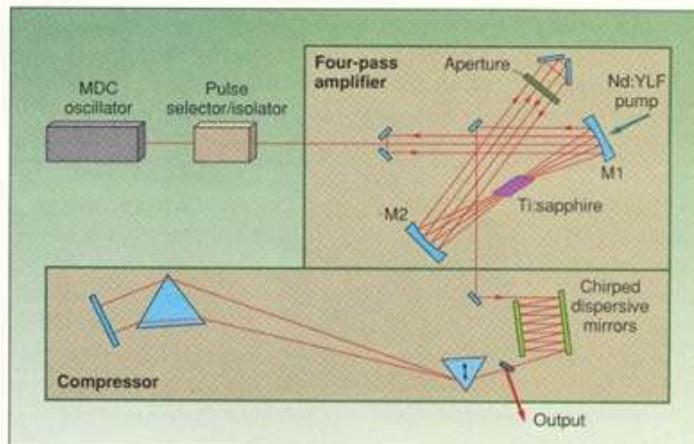
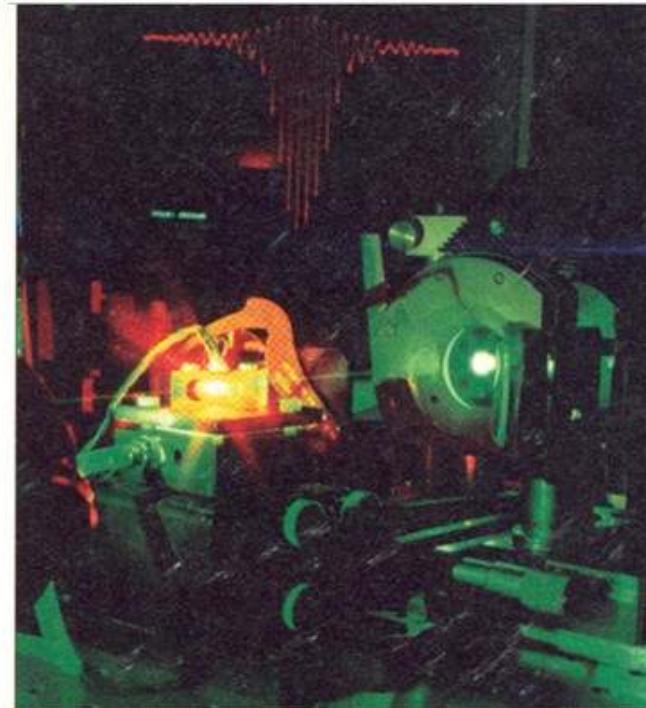


FIGURE 4. Four-pass kilohertz Ti:sapphire amplifier produces 18-fs pulses with an energy of 100 μ J (after compression) when pumped with 6-mJ pulses from a frequency-doubled Nd:YLF laser.



Multipass Ti:sapphire amplifier is seeded with high-quality 8-fs pulses generated by a Ti:sapphire oscillator incorporating chirped dielectric mirrors for dispersion compensation.

TECHNISCHE UNIVERSITÄT WIEN

Ch. Spielmann, M. Lenzner, F. Krausz, R. Szipőcs, K. Ferencz,
„Chirped dielectric mirrors improve Ti:sapphire lasers,” Laser Focus World, 1995 December, pp. 55-60 (1995).

Compression of high-energy laser pulses below 5 fs

M. Nisoli, S. De Silvestri, and O. Svelto

Centro di Elettronica Quantistica e Strumentazione Elettronica—Consiglio Nazionale delle Ricerche, Dipartimento di Fisica, Politecnico, Piazza L. da Vinci 32, 20133 Milano, Italy

R. Szipöcs and K. Ferencz

Szilárdtestfizikai Kutatóintézet, Pf. 49, H-1525 Budapest, Hungary

Ch. Spielmann, S. Sartania, and F. Krausz

Abteilung Quantenelektronik und Lasertechnik, Technische Universität Wien, Gusshausstrasse 27, A-1040 Wien, Austria

Received October 25, 1996

High-energy 20-fs pulses generated by a Ti:sapphire laser system were spectrally broadened to more than 250 nm by self-phase modulation in a hollow fiber filled with noble gases and subsequently compressed in a broadband high-throughput dispersive system. Pulses as short as 4.5 fs with energy up to $20\text{-}\mu\text{J}$ were obtained with krypton, while pulses as short as 5 fs with energy up to $70\text{ }\mu\text{J}$ were obtained with argon. These pulses are, to our knowledge, the shortest generated to date at multigigawatt peak powers. © 1997 Optical Society of America

- Spectral broadening in hollow core fibre filled with noble gas (Ar/Kr) (M. Nisoli, S. De Silvestri, O. Svelto, Appl. Phys. Lett. 68, 2793 (1996))
- Dispersion compensation by ultrabroadband chirped mirror (E.J. Mayer, J. Möbius, A. Euteneuer, W.W. Röhle, R. Szipöcs: Opt. Lett. 22, 528 (1997))
- Compressed pulse duration ~5 fs at ~800 nm
- ~1.5 oscillation of electromagnetic field > allows generation of **isolated attosecond pulses** by high harmonic generation (HHG)

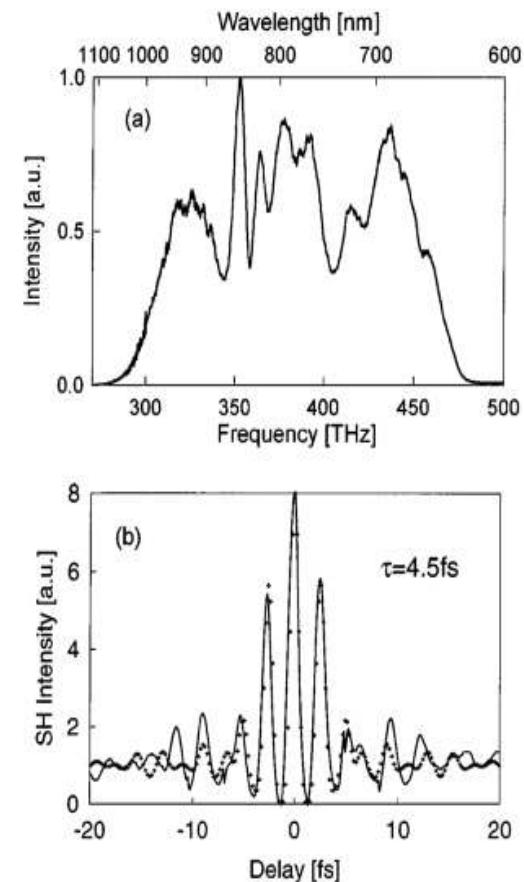
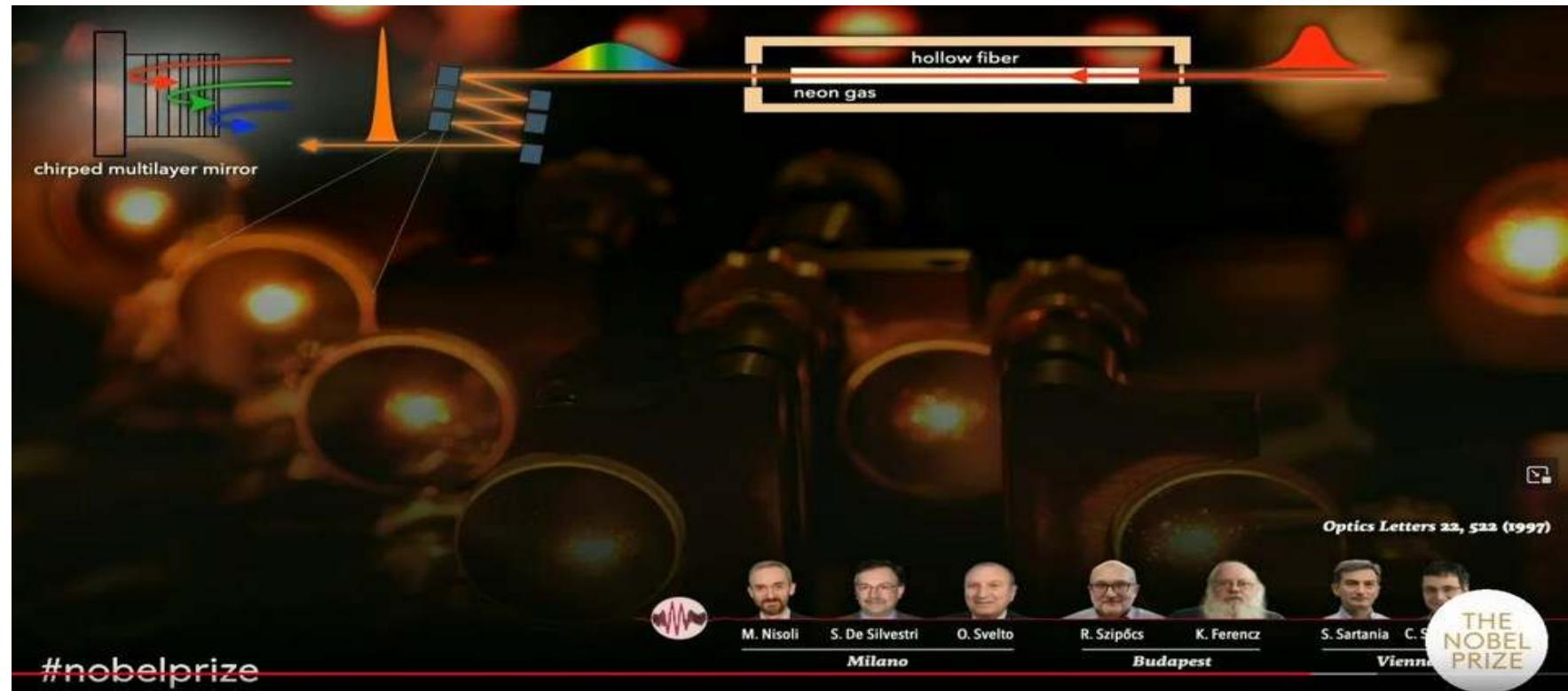


Fig. 2. (a) Spectral broadening in krypton at $p = 2.1$ bars and $P_0 = 2$ GW. A low-intensity pedestal (~1% of the peak) extends below 600 nm. (b) Measured (solid curve) and calculated (crosses) autocorrelation trace; an evaluation of the pulse duration (FWHM) is also given.

Generating isolated attosecond pulses by HHG of a sub-mJ, 5 fs pulse



Generation of ~5 fs, sub-mJ laser pulses using spectral broadening of ~18 fs amplified laser pulses in a noble gas filled hollow core fiber and chirped mirrors for dispersion compensation

2023 Nobel Prize lectures in physics, Pierre Agostini, Ferenc Krausz and Anne L'Huillier, <https://www.youtube.com/watch?v=xVXjFBW-2kI>

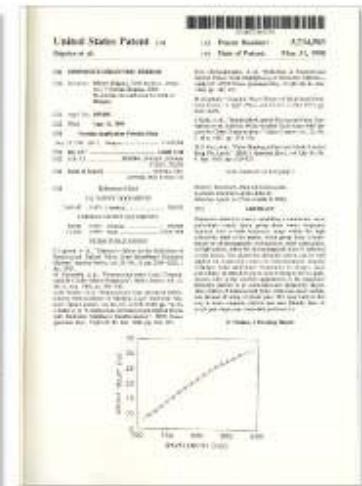
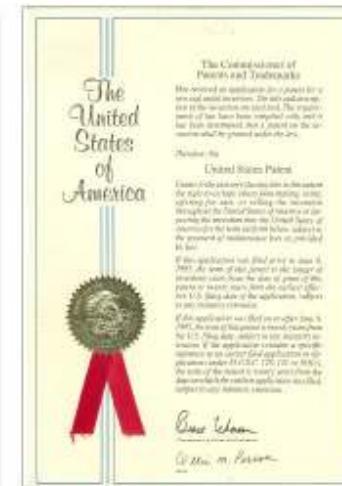
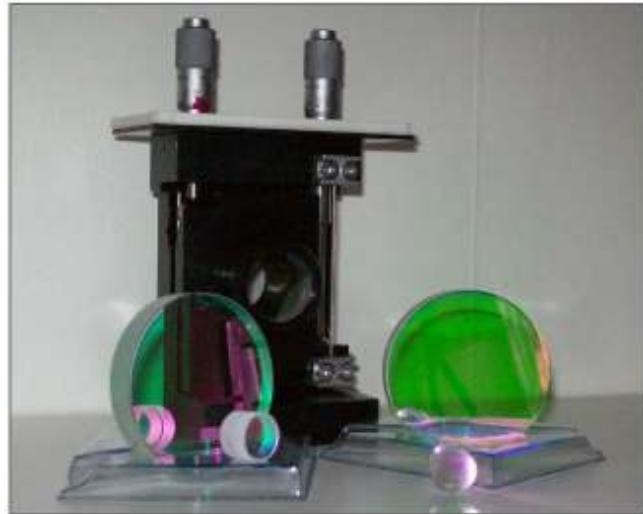
M. Nisoli, S. De Silvestri, O. Svelto, R. Szipőcs, K. Ferencz, Ch. Spielmann, S. Sartania, and F. Krausz,
"Compression of high-energy laser pulses below 5 fs," Opt. Lett. 22, 522-524 (1997)

OPTICS

Pioneering Ultrafast LaserTechnology by R&D
INVENTING CHIRPED MIRRORS

Femtosecond Dispersive
and Broadband Optics by IBS technology

Patents



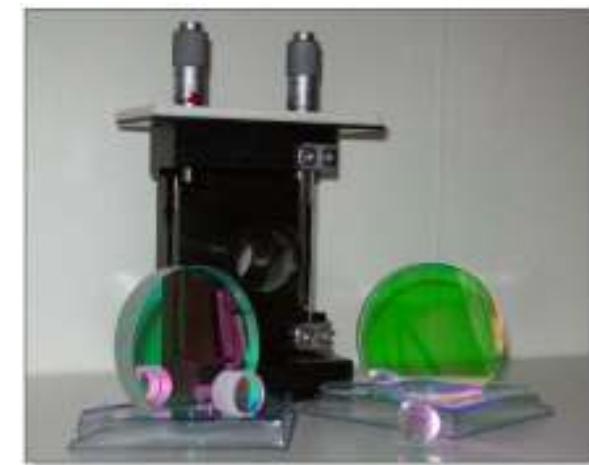
Founding R&D Lézer-Optika Bt. in 1995 for commercialization of chirped mirror technology

Related patent submitted in August 1993 by Róbert Szipőcs and Ferenc Krausz at MTA SZFKI

R&D ULTRAFAST LASERS LTD.

Femtosecond Dispersive and Broadband Optics by IBS technology

- Chirped mirrors (CM)
- Low dispersion ripple, highly dispersive negative dispersion mirrors (MCGTI)
- Ultrabroadband chirped mirrors (UBCM)



DISPERSIVE MIRRORS, CHARACTERIZATION: WHITE LIGHT INTERFEROMETRY AND CCD

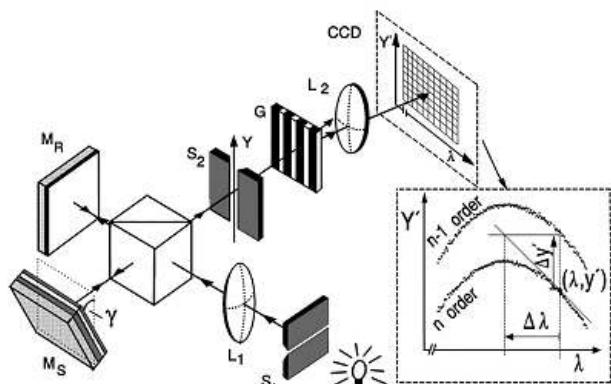
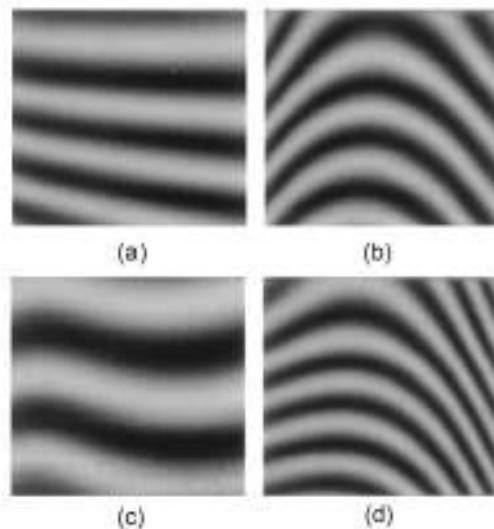


Fig. 1. Spectrally resolved white-light interferometer for group-delay measurement of dielectric mirrors. L_1 , L_2 : achromatic lenses; S_1 , S_2 , slits; M_S , sample mirror; M_R , reference mirror; G , transmission grating.



- (a) Low dispersion sample (linear phase shift)**
- (b) Chirped mirror sample (quadratic phase shift)**
- (c) Gires-Tournois Interferometer mirror (cubic phase shift)**
- (d) (c)+(d)**

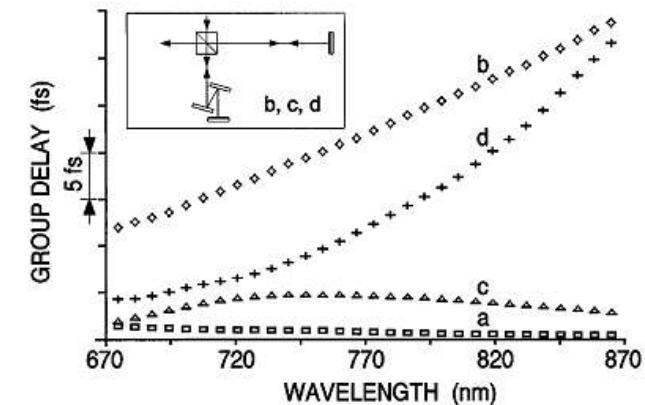


Fig. 4. Measured group-delay functions obtained by computer processing of the images shown in Fig. 3 (every fifth point is plotted). The curves correspond to a single reflection. Inset: four-reflection arrangement used for measuring curves b-d.

R&D ULTRAFAST LASERS LTD.

Femtosecond Dispersive
and Broadband Optics by IBS technology

Services

- **Custom design** of femtosecond laser mirrors for dispersion compensation (Ti:S, Cr³⁺, Yb³⁺, etc.) IR OPO, Vis-OPO, OPA, etc.
- **Dispersion measurement** on laser mirrors and other optical components.



Chirped mirrors for fs optical parametric oscillators

April 15, 1995 / Vol. 20, No. 8 / OPTICS LETTERS 919

Chirped-mirror dispersion-compensated femtosecond optical parametric oscillator

J. Hebling, E. J. Mayer, and J. Kuhl

Max-Planck-Institut für Festkörperforschung, Heisenbergstrasse 1, D-70569 Stuttgart, Germany

R. Szipöcs

Optical Coating Laboratory, Research Institute for Solid State Physics, P.O. Box 49, H-1525 Budapest, Hungary

Received January 3, 1995

We describe the operating characteristics of a femtosecond optical parametric oscillator employing chirped mirrors for intracavity group-velocity dispersion compensation. Pumped by 760 mW of power from a self-mode-locked Ti:sapphire laser, this device provides 100-fs near-transform-limited pulses continuously tunable from 1.18 to 1.32 μm with an average power of 100–180 mW. The limitations of the present setup and strategies for further pulse shortening are discussed.

Ultrabroadband chirped mirrors for ultrafast lasers

528 OPTICS LETTERS / Vol. 22, No. 8 / April 15, 1997

Ultrabroadband chirped mirrors for femtosecond lasers

E. J. Mayer, J. Möbius, A. Euteneuer, and W. W. Röhle

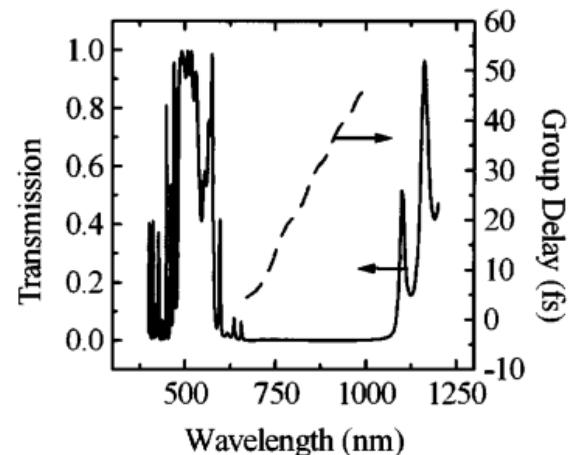
Department of Physics, Philipps University, Renthof 5, D-35032 Marburg, Germany

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R&D Lézer-Optika Bt., P.O. Box 622, H-1530 Budapest, Hungary

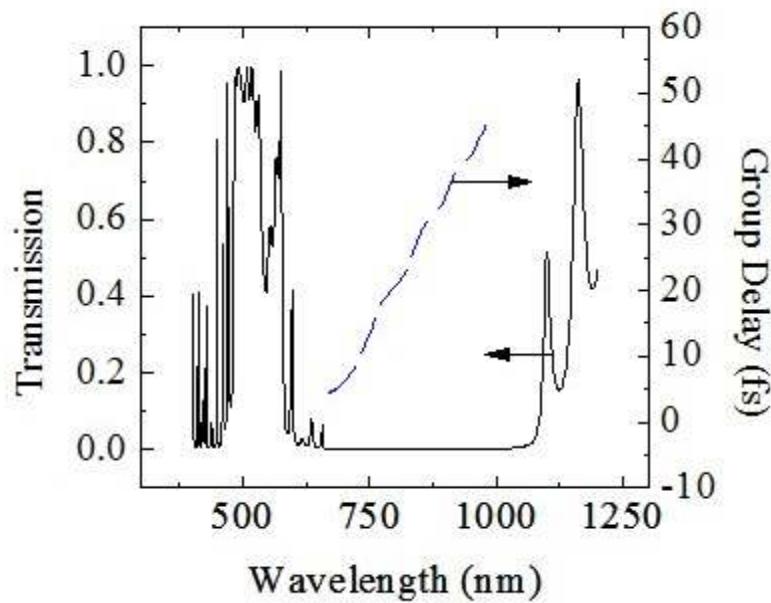
Received November 25, 1996

We report on the performance of widely tunable femtosecond and continuous-wave Ti:sapphire lasers that use a newly developed ultrabroadband mirror set. The mirrors exhibit high reflectivity ($R > 99\%$) and smooth variation of group delay versus frequency over a wavelength range from 660 to 1060 nm. Mode-locked operation with pulse durations of 85 fs was achieved from 693 to 978 nm with only one set of ultrabroadband mirrors. © 1997 Optical Society of America



- The first widely tunable femtosecond pulse Ti:sapphire laser
- High reflectivity ($R . 99\%$) and smooth variation of group delay over a wavelength range from 660 to 1060 nm
- Mode-locked operation from 693 to 978 nm using one set of mirrors

Ultrabroadband chirped mirrors for ultrafast lasers



Increased reflectivity band and smooth dispersion over the 680-1060 nm wavelength range.



"Broadband Optics with Λ -track Extend the Reach of Multiphoton Microscopy"

 **Spectra-Physics**

The Solid State Laser Company™

E.J. Mayer, J. Möbius, A. Euteneuer, W. Rühle, R. Szipócs: Opt. Lett. 22, 528-530 (1997)

Compression of laser pulses down to 4.6 fs

Optics in 1997

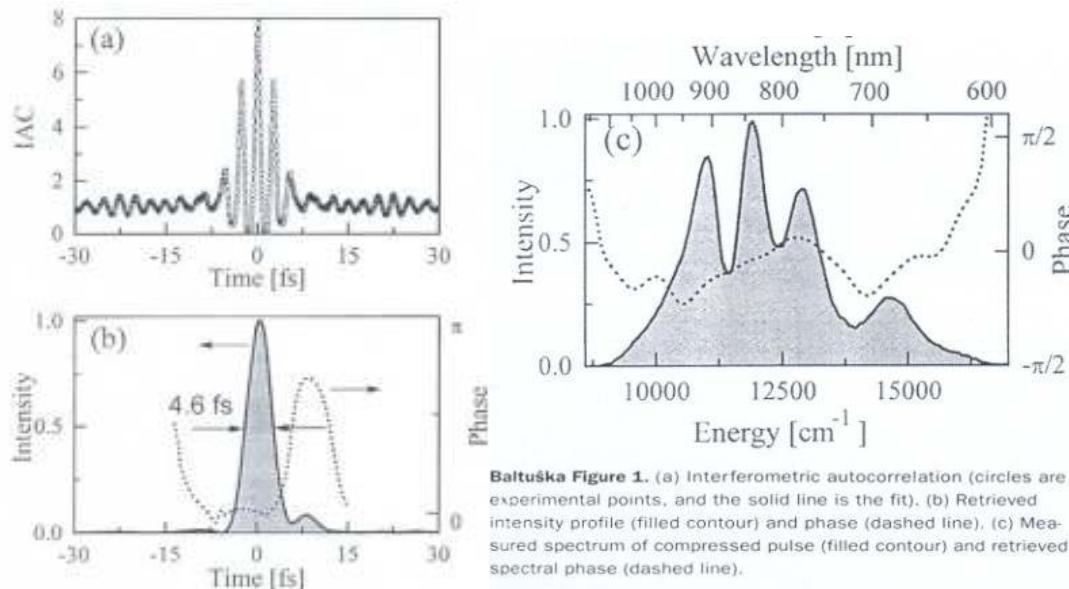
Ultrafast Technology

ULTRAFAST TECHNOLOGY

A Compact All-Solid-State Sub-5-fsec Laser

Andrius Baltuška and Maxim S. Pshenichnikov, Ultrafast Laser and Spectroscopy Laboratory, Dept. of Chemistry, Univ. of Groningen, Groningen, The Netherlands; Róbert Szűcs, Research Institute for Solid State Physics, Budapest, Hungary; and Douwe A. Wiersma, Ultrafast Laser and Spectroscopy Laboratory, Dept. of Chemistry, Univ. of Groningen, Groningen, The Netherlands.

Recent developments in solid-state lasers,¹ chirp-mirror technology,² and methods of pulse characterization³ made it possible to design an all-solid-state laser that delivers sub-5-fsec pulses at a 1-MHz repetition rate.⁴ Such extremely short light pulses at a high



Baltuška Figure 1. (a) Interferometric autocorrelation (circles are experimental points, and the solid line is the fit). (b) Retrieved intensity profile (filled contour) and phase (dashed line). (c) Measured spectrum of compressed pulse (filled contour) and retrieved spectral phase (dashed line).

R&D ULTRAFAST LASERS LTD.



Founding

R&D Ultrafast Lasers Research and Development Ltd. in 1997

BOOTH NUMBER: 8109

R&D ULTRAFAST LASERS LTD.



Company Description

Featured Product: Dual wavelength fs laser system for 3D CARS imaging including tunable Ti:sapphire and Yb fiber laser

Manufacturer of single or double wavelength ultrafast laser systems including ultrashort (ps or fs) pulse, ultrabroadband or broadly tunable Ti:sapphire lasers, Yb-doped fiber lasers, amplifiers and optical parametric oscillators. Their typical applications include time resolved or CARS spectroscopy or nonlinear (2P, SHG or SRS/CARS) microscopy. Manufacturer of ultrafast laser optical coatings including different dispersive mirrors such as chirped mirrors. Complete laser laboratory construction.



Founded:	in 1997
Location:	1121 Budapest, Konkoly Thege út 29-33. 6. ép. I. em. (KFKI Campus)
Infrastructure:	3 laser-optical laboratories, 1 electronic and 1 mechanical workshops, offices
Staff:	Engineering (2), Software (1), Optics/Lasers (1), Administration (0.5 + 0.5)
Web-site:	www.sziopcs.com

1. Scientific background

1.1 Optics (ultrafast – ps, fs)

1.2 Lasers (ultrafast – ps, fs)

 1.2.1 Solid state

- **Ti:sapphire (680 – 1040 nm)**

- **OPO-s (signal: 1020- 1240 nm, idler: 2 – 2.5 μm)**

 1.2.2 Fiber

- **Yb-fiber (1020-1060 nm)**

- Er-fiber (1520-1600 nm), SHG: 760 – 800 nm)

1.3 Microscopy

- 2PEF, SHG, **CARS for Life Science**

- **FLIM (confocal, 2P) for Life Science and Quantum Optics**

2. Technical background

2.1 Optics (thin films, optical design – Zemax)

2.2 Mechanics (ProE, SolidWorks)

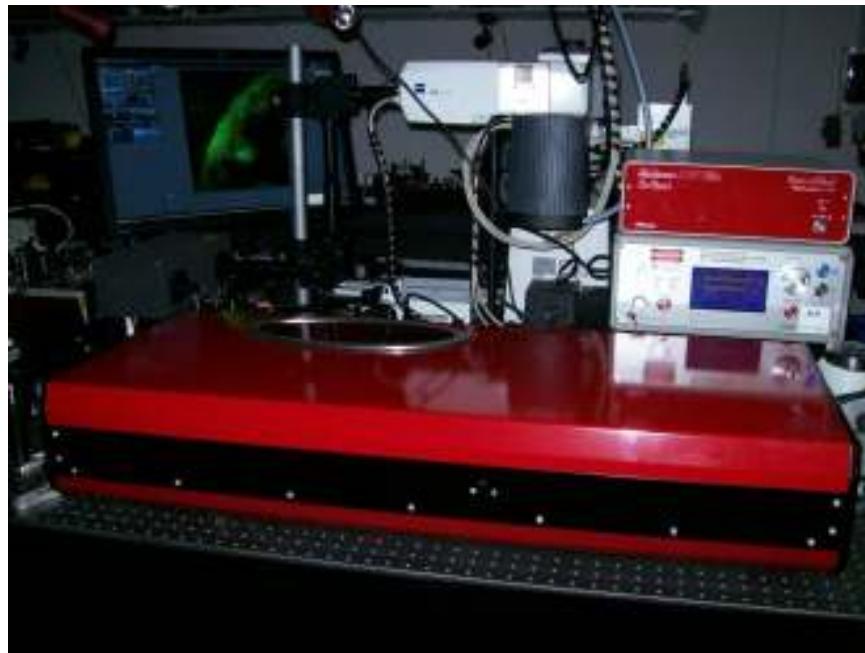
2.3 Electronics (Eagle)

2.4 Software (C# (ARM, PIC), Vivado (FPGA), Delphi, LabView)

FemtoRose 100 TUN/NoTouch

The First

Broadly Tunable, femtosecond pulse Ti:sapphire laser in 1998



Pioneering Ultrafast LaserTechnology by R&D

INTRODUCING THE FIRST ULTRABROADBAND CHIRPED MIRRORS
FOR BROADLY TUNABLE FEMTOSECOND LASERS

528 OPTICS LETTERS / Vol. 22, No. 8 / April 15, 1997

Ultrabroadband chirped mirrors for femtosecond lasers

E. J. Mayer, J. Möhns, A. Endesuer, and W. W. Röhle

Department of Physics, Philipps University, Renthof 5, D-3500 Marburg, Germany

R. Szepics

R&D Laser-Optics Kft., P.O. Box 622, H-1339 Budapest, Hungary

Received November 25, 1996

We report on the performance of widely tunable femtosecond and continuous-wave Ti:sapphire lasers that use a newly developed ultrabroadband mirror set. The mirrors exhibit high reflectivity ($R > 99\%$) and smooth variation of group delay versus frequency over a wavelength range from 690 to 1050 nm. Mode-locked operation with pulse durations of 85 fs was achieved from 690 to 978 nm with only one set of ultrabroadband mirrors. © 1997 Optical Society of America

PPLN OPO FOR 1 to 1.4 MICRON (signal) AND 2.0-2.5 MICRON (idler)

R&D ULTRAFAST LASERS LTD.

FemtoRainbow 100 OPO

Femtosecond tunable synchronously pumped optical parametric oscillator



- *Ti:Sapphire laser wavelength conversion*
- *Synchronously pumped at ~76 MHz*
- *Output is widely tunable from 1010 to 1260 nm*
- *Output power from up to 100 mW*
- *15 nm to 30 nm spectral width (FWHM)*
- *KTP or PPLN crystal based conversion*
- *Wavelength stabilization by computer control*

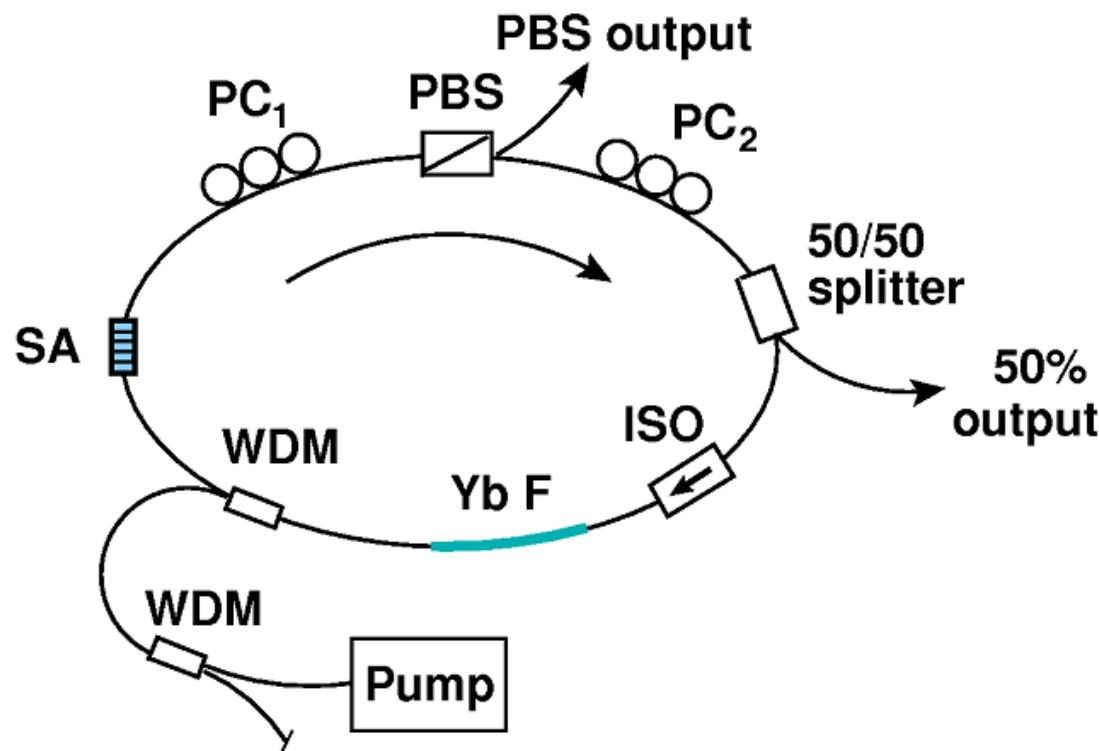
Development of Yb-fiber laser technology in the FemtoBio NTP project (2006-2009)
MTA SZFKI – R&D Ultrafast Lasers Ltd – Furukawa Electric



J. Fekete, A. Cserteg, Szipőcs; All-fiber, all-normal dispersion ytterbium ring oscillator;
Laser Physics Letters 6, 49-53, 2009

Development of Yb-fiber laser technology in the FemtoBio NTP project (2006-2009)

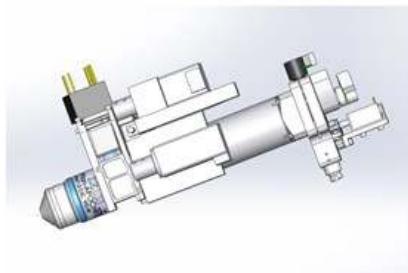
MTA SZFKI – R&D Ultrafast Lasers Ltd – Furukawa Electric



J. Fekete, A. Cserteg, Szipőcs; All-fiber, all-normal dispersion ytterbium ring oscillator;
Laser Physics Letters 6, 49-53, 2009

Our *FiberScope* project for Biomedical imaging applications (2010-2014)

- Reduce the cost of the pulsed laser applied (Yb-fiber instead of Ti:sapphire)
- Deliver the light for measurements through optical fiber (**fiber delivery**)
- Small size scanning microscope head (handheld device)
- In vivo 3D measurements: laser safety issues
- Applications in dermatology and nanomedicine

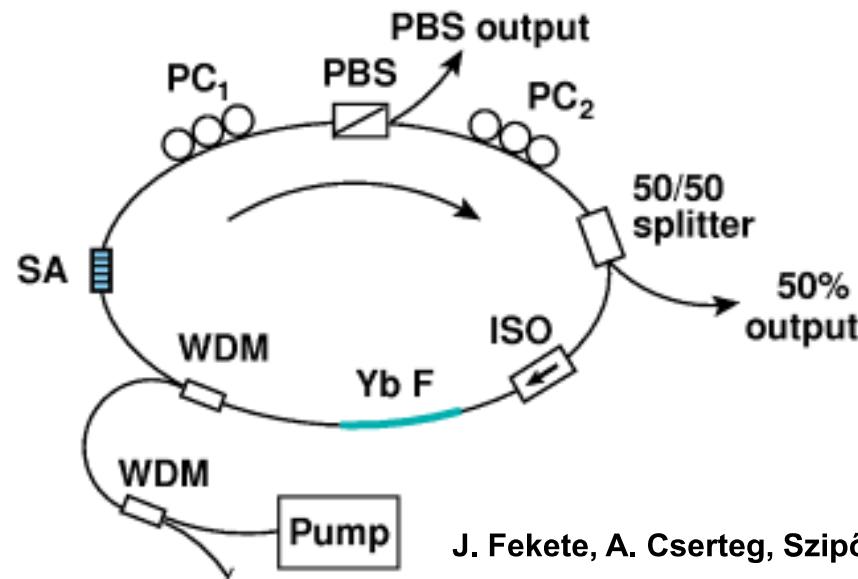


FemtoFiber + scanning head for confocal/2PF imaging = FiberScope

FIBER LASERS

All-Fiber, All-Normal-Dispersion Ytterbium Ring Oscillator

- Determined by interplay between **gain, self-phase modulation, dispersion and filtering** effects
- Pulse shaping is based on **nonlinear polarization rotation** in the fiber together with **spectral and temporal filtering** by a polarizing element



PC: polarization controller

PBS: polarizing beam splitter

ISO: isolator

Yb F: Ytterbium doped fiber

WDM: wavelength division multiplexer

SA: saturable absorber

J. Fekete, A. Cserpeg, Szipőcs; All-fiber, all-normal dispersion ytterbium ring oscillator,
Laser Physics Letters 6, 49-53, 2009



FiberScope system description, for details, see:



Handheld nonlinear microscope system comprising a 2 MHz repetition rate, mode-locked Yb-fiber laser for *in vivo* biomedical imaging

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<https://www.szipocs.com>

FiberScope bloch scheme

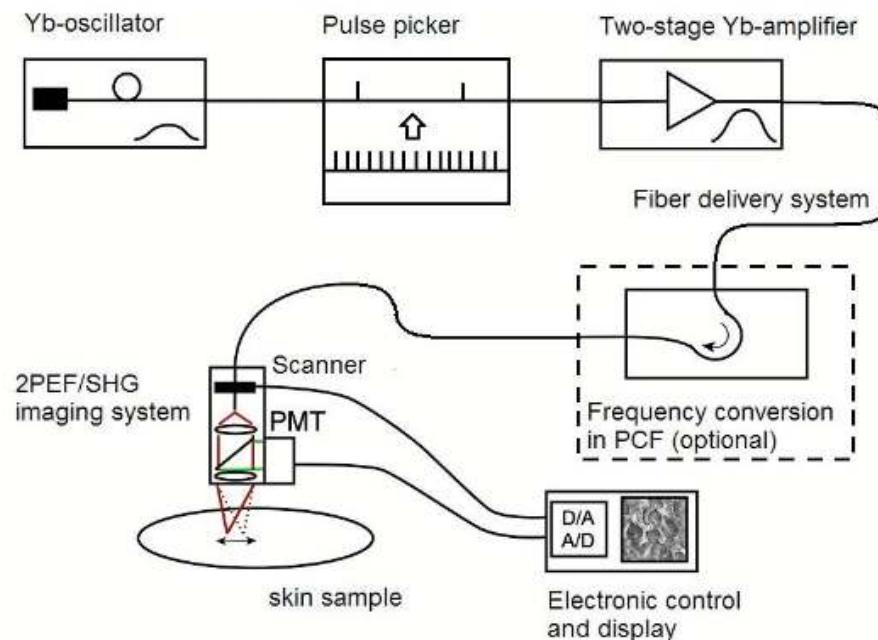
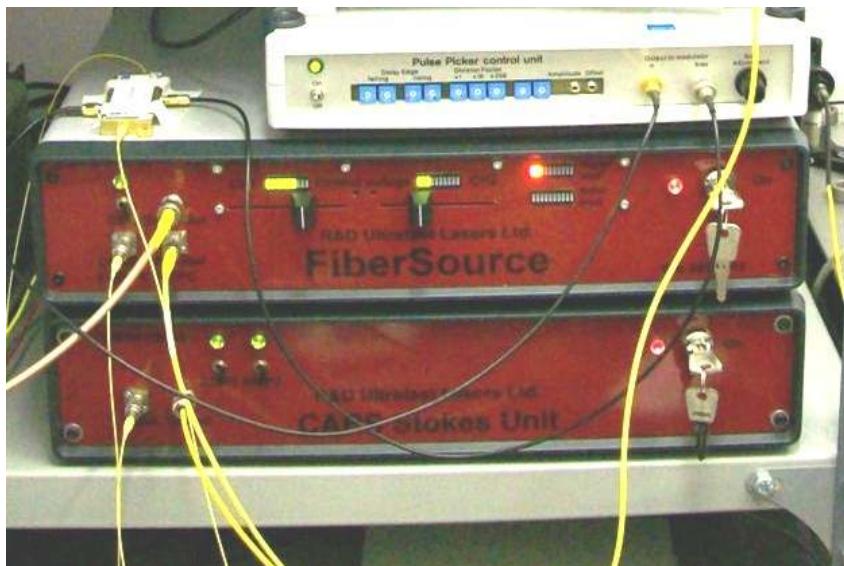


Fig. 1. Bloch scheme of the handheld 2PEF/SHG microscope imaging system comprising a 2 MHz mode-locked Yb-fiber laser.

FiberScope nonlinear microscope

Photo of the 2MHz Yb-fiber laser system used as a pulsed laser light source of our *FiberScope* device

Photo of the scanning microscope head of the *FiberScope* device with plastic covering



The low cost, flexible, ~2 MHz pulsed laser source

Laser components and performance

- All-fiber, AND type, mode-locked Yb-fiber oscillator (*R&D*)
- Pulse picker (*JenOptik*, Jena, Germany)
- Two-stage Yb-amplifier (*R&D Ultrafast Lasers Ltd.*)

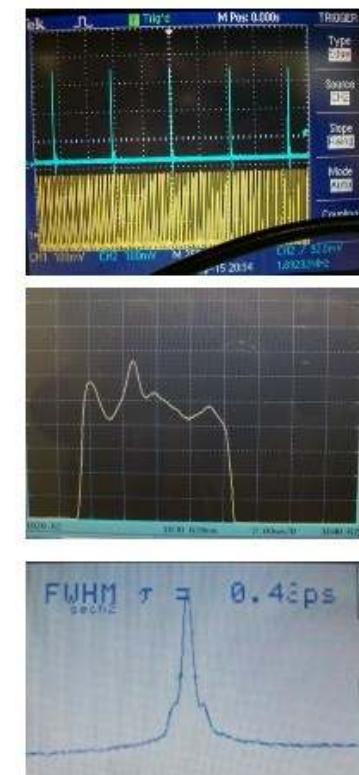
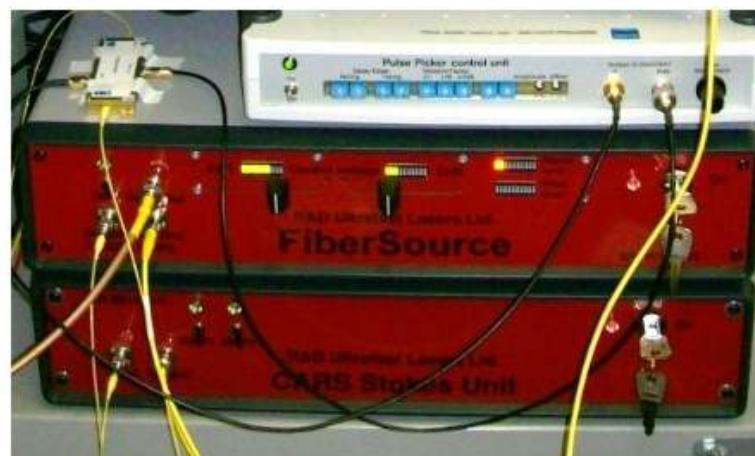
Pave: ~ 200 mW

$\tau < 0.5$ ps

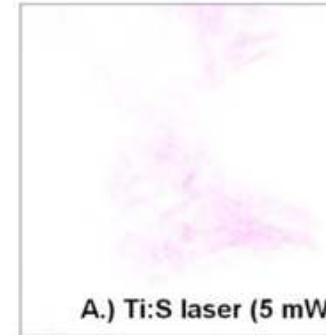
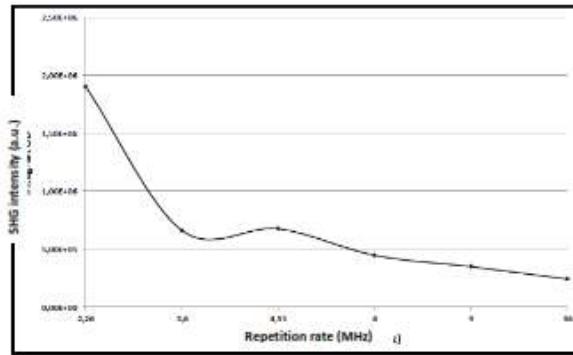
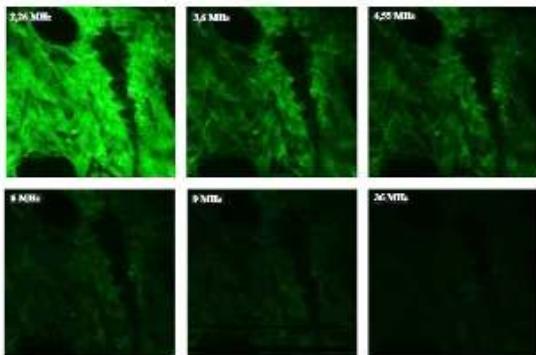
$\lambda_0 \sim 1030$ nm

$\Delta\lambda$: 8-12 nm

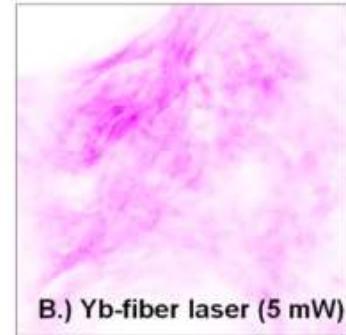
$v: \sim 1.89$ MHz



Low repetition rate [7] and IR excitation [4] advantage



A.) Ti:S laser (5 mW)

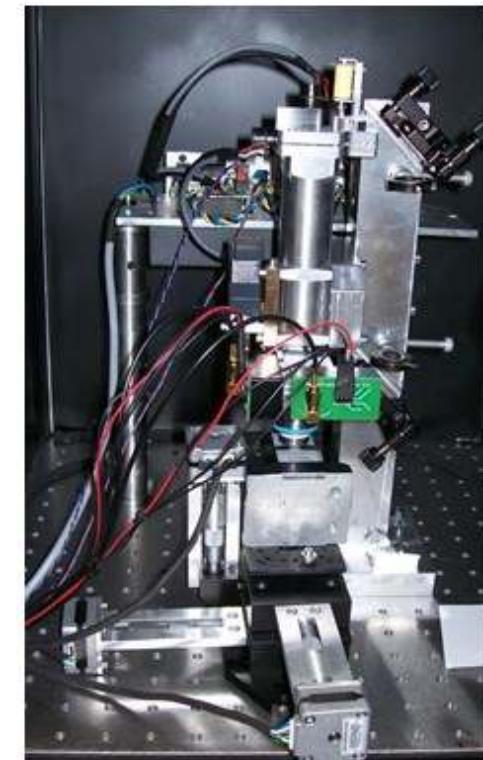
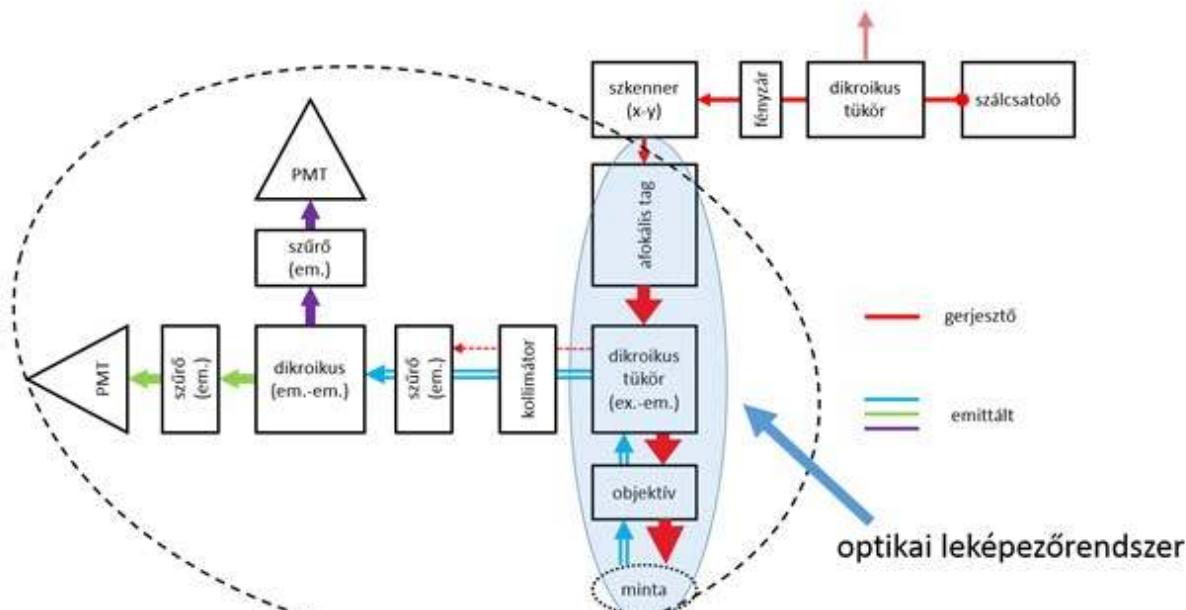


B.) Yb-fiber laser (5 mW)

SHG signal intensity as the function of repetition rate of the Yb-fiber laser. Average power: 5 mW (measured after the microscope objective). Pixel dwell time: 5 μ s. Operation wavelength: 1030 nm. Fixed transmission grating distance of \sim 80 mm. Ex-vivo murine skin sample.

Comparison of SHG imaging performance of different mode-locked lasers having the same average power of 5 mW (on the sample) for nonlinear microscopy. Ex-vivo murine skin sample, imaging depth: $z = 30 \mu\text{m}$, same microscope settings. Collagen distribution measured by A) an industry standard, 80 MHz Ti:sapphire laser operating at 800 nm, and by B) our 2MHz Yb-fiber oscillator and amplifier system operating at 1030 nm.

Optical design of the scanning microscope head of the *FiberScope*



A. Csákányi, R. Szipőcs R: Design of optical imaging system of a scanning nonlinear microscope „Fiberscope”

Kvantumelektronika 2014, Budapest, 28.11.2014, Paper P64.

Investigation of skin tumor basalioma (BCC)

- Collagen, as marker
- Ex vivo samples, immediately after surgery



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doi: 10.1111/j.1600-0846.2012.00641.x

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Skin Research and Technology

Diagnosis of BCC by multiphoton laser tomography

Stefania Seidenari¹, Federica Arginelli¹, Sara Bussoli¹, Jennifer Cautela¹, Anna Maria Cesinaro², Mario Guanti¹, Davide Guardo¹, Cristina Magnoni¹, Marco Manfredini¹, Giovanni Ponti¹ and Karsten König^{3,4}

¹Department of Dermatology, University of Modena and Reggio Emilia, Modena, Italy

²Department of Pathology, University of Modena and Reggio Emilia, Modena, Italy

³Department of Biophotonics and Lasertechnology, Saarland University, Saarbrücken, Germany and ⁴AntLab GmbH, Schillerstraße 3, 6745, Ettlingen, Germany

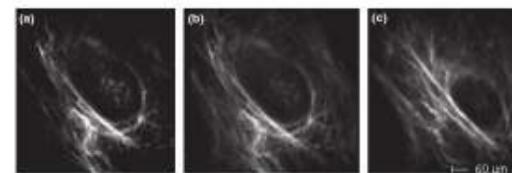
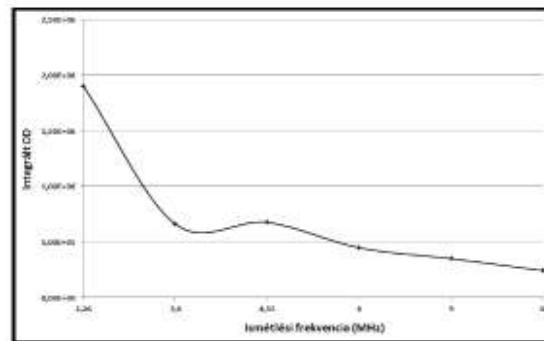
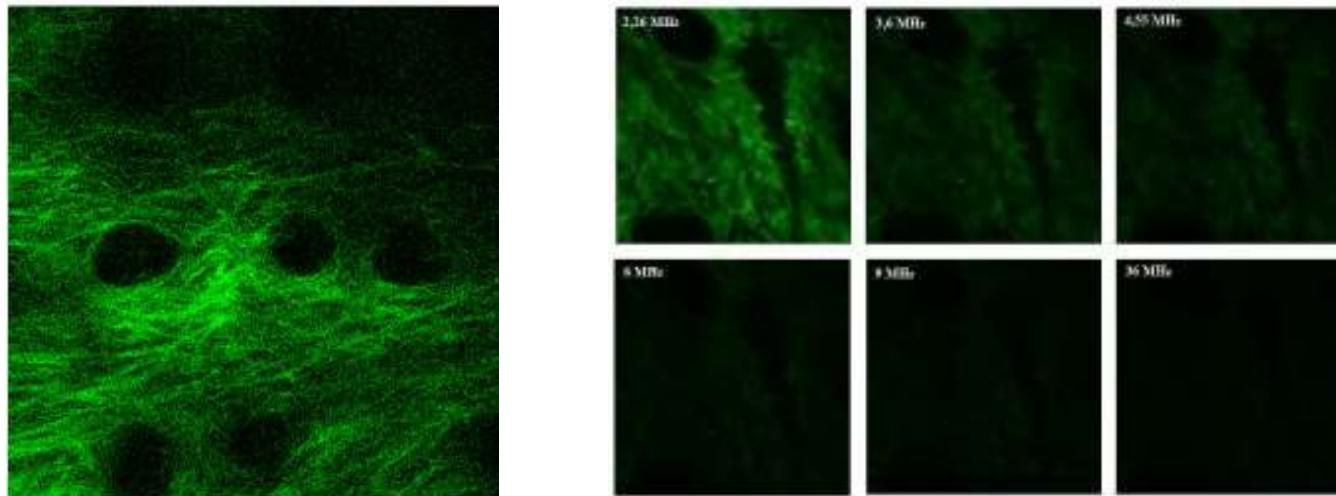


Fig. 4. Basal cell carcinoma. (a) 100 μ m depth, excitation wavelength 800 nm. Shifting the wavelength to 820 nm, basaloid cells become less visible; employing an excitation wavelength of 820 nm basaloid cells disappear and it is possible to observe empty spaces surrounded by collagen fibres (phantom island); (b) 200 μ m depth; (c) 320 μ m depth.

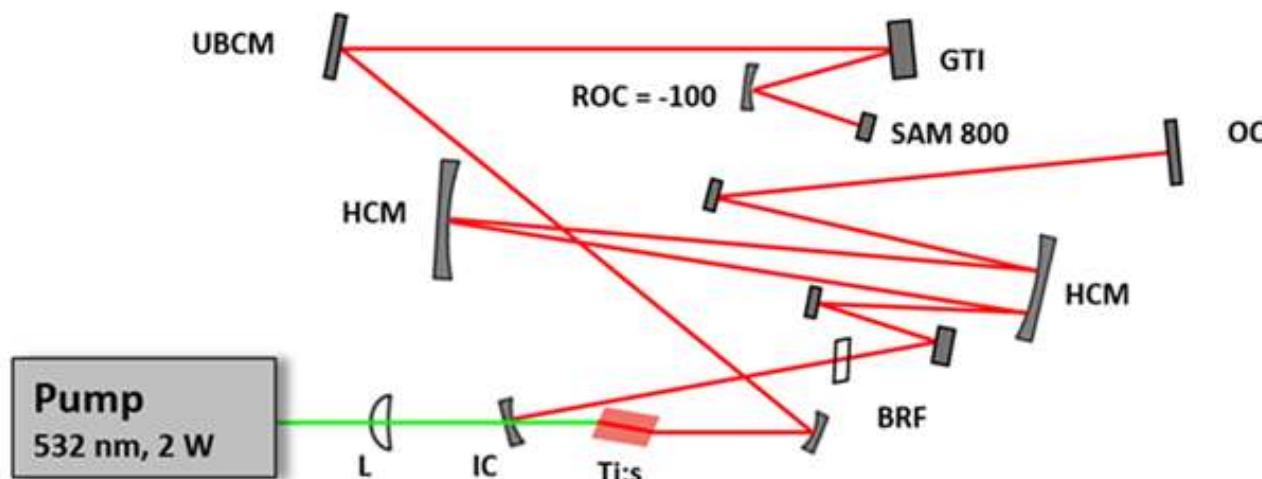
Using *FiberScope* for collagen detection



SHG intensity of collagen as the function of repetition rate of our Yb-fiber laser
(average power: ~5 mW, pixel dwell time: 5 μ s)

20 MHz, tunable, sub-ps Ti:sapphire laser

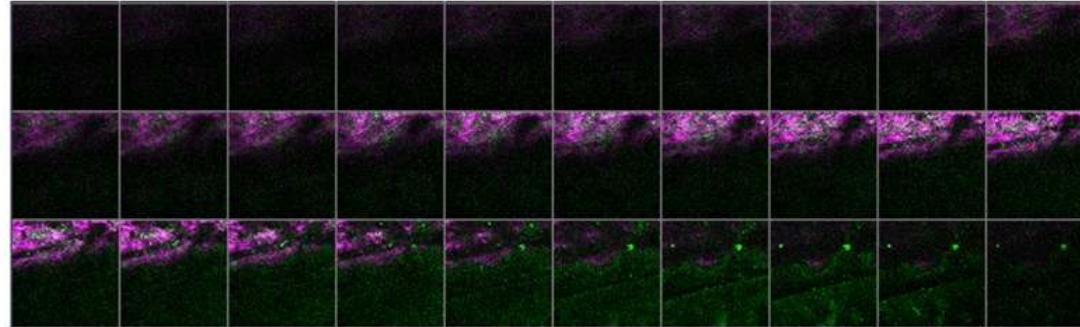
Setup upgraded for DVRF CARS and fiber delivery



- Prism pair is replaced by GTI mirrors
- SESAM for mode-locking instead of hard aperture Kerr-lens mode-locking
- Ultrabroadband chirped mirrors

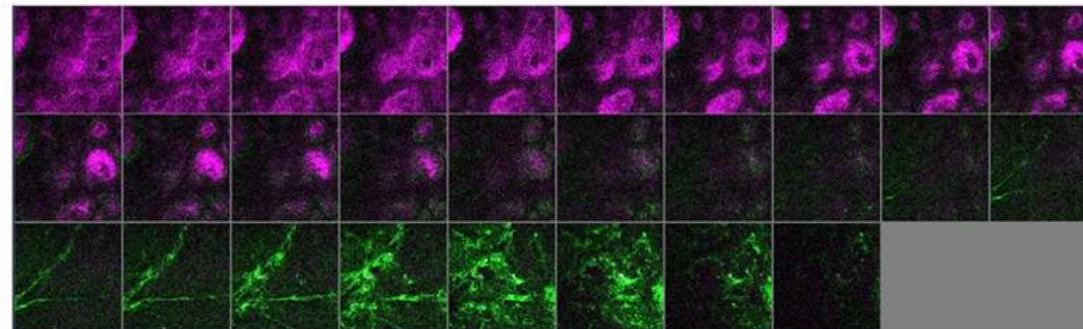
- Pump power at 532 nm: ~ 2.1 W
- Repetition rate: ~ 20 MHz
- Pulse duration: 0.6 – 1.0 ps (adjustable)
- Spectral bandwidth: < 2 nm

Z-stack imaging of basal cell carcinoma (BCC) biopsy



In case of BCC, we could detect the border of the tumor highlighted by a strong contrast of AF (cells with basal cell morphology) and SHG (collagen fibers around tumor nest).

Z-stack imaging of hemangioma biopsy



Hemangioma and angiokeratoma are vascular proliferations. **Hemangiomas** can be either congenital or acquired, spontaneously evolving with age. These do not require further treatment. **Angiokeratomas (AK)** can not be distinguished from hemangiomas macroscopically, and even though these have dermatoscopic features, dermatoscopic differentiation is often challenging. The distinguishing between hemangiomas and AK is important as AK may indicate the presence of other severe diseases. It is the most characteristic sign of **Fabry-disease** leading to **severe heart, lung and kidney damage**. Early diagnosis could be promoted by the **recognition of AKs**. AKs can be a sign of other storage diseases.

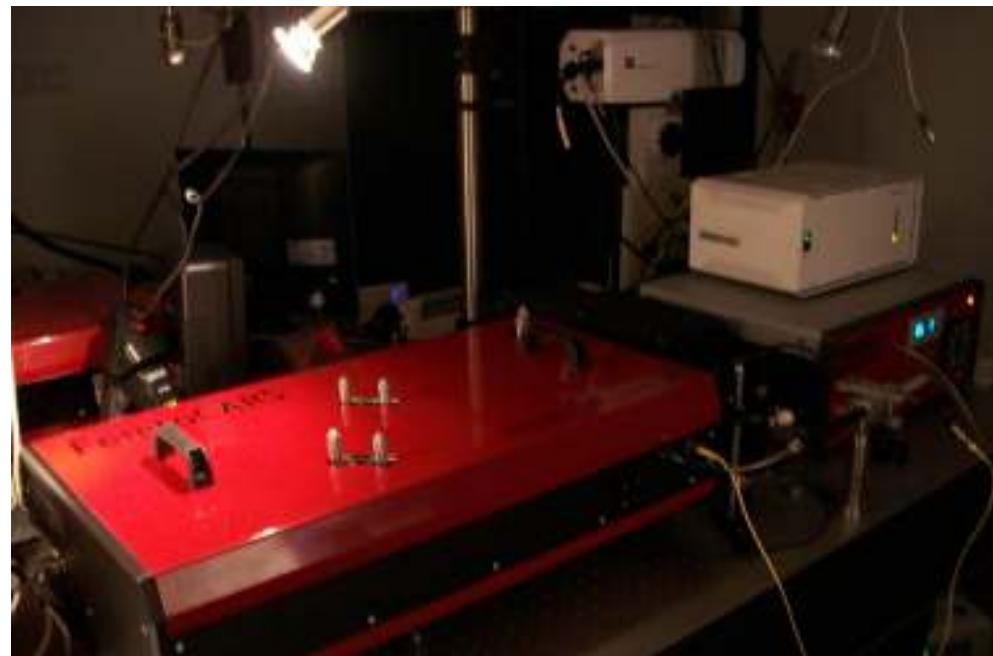
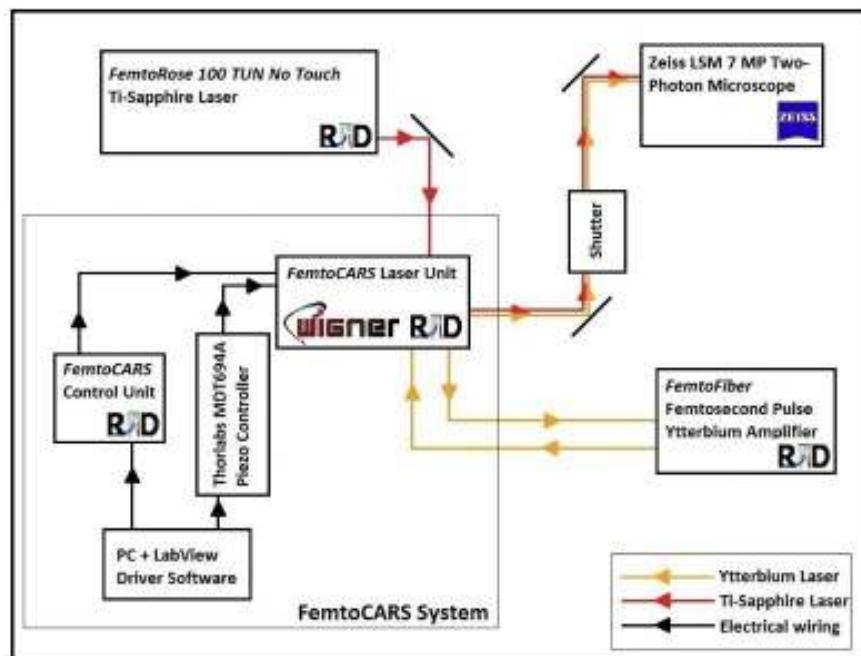
Yb-FIBER AMPLIFIER FOR CARS IMAGING



FemtoCARS

the

Label-free, 3D Microscopic Imaging System for Real-time in vivo Diagnostics



Dual vibration resonance frequency CARS microscopy imaging of basal cell carcinoma to achieve stain free histopathology

Previously, we utilized CARS microscopy to visualize lipids in the adipocytes of murine skin

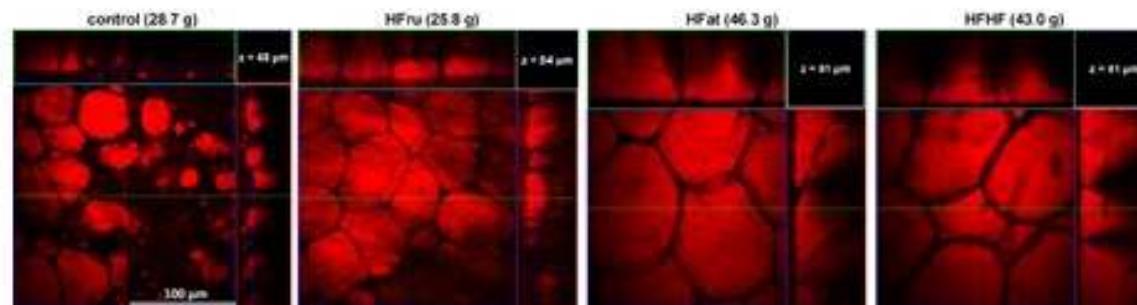


Biomed Opt Express. 2016 Nov 1; 7(11): 4480–4489
Published online 2016 Oct 7. doi: [10.1364/OE.7.11.4480](https://doi.org/10.1364/OE.7.11.4480)

PMCID: PMC5119669

Diet-induced obesity skin changes monitored by *in vivo* SHG and *ex vivo* CARS microscopy

Dóra Haluszka,^{1,2} Kende Lőrincz,¹ Norbert Kiss,^{1,2} Robert Szöpös,^{2,3} Enikő Kurov,¹ Nőra Gyöngyösi,¹ and Norbert M. Winkler^{1,*}



BUILDING LABORATORIES



CARS SYSTEM INSTALLED AT UNIVERSITY OF SZEGED

[Related articles](#)

CARS imaging system installed at the University of Szeged, Department of Neurology (Prof. Gábor Tamás lab), June 2014

Photo Gallery





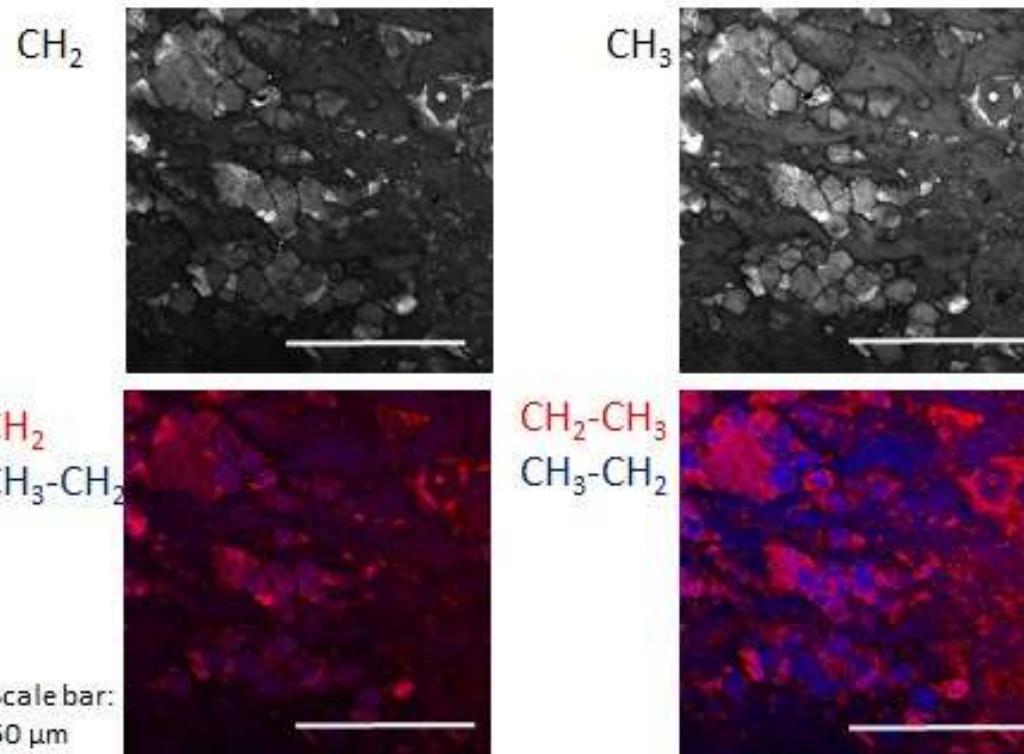
Stain-free Histopathology of Basal Cell Carcinoma by Dual Vibration Resonance Frequency CARS Microscopy

Authors

[Authors and affiliations](#)

Norbert Kiss, Ádám Kroopp, Kende Lőrincz, András Bánvölgyi, Róbert Szipőcs , Norbert Wikonkál

CARS images of human basal cell carcinoma



Real time DVRF-CARS (double channel, lipid/protein detection) using a 20 MHz, sub-ps Ti:sapphire laser

NTHOA.4.pdf

Biophotonics Congress, Biophotonic Optical 2020
(Tunisiania, Montevideo, OCT 07-09, BHAN) © CLEA,
2020

A 20 MHz, sub-ps, Tunable Ti:sapphire Laser System for Real Time, Stain Free, High Contrast Histology of the Skin

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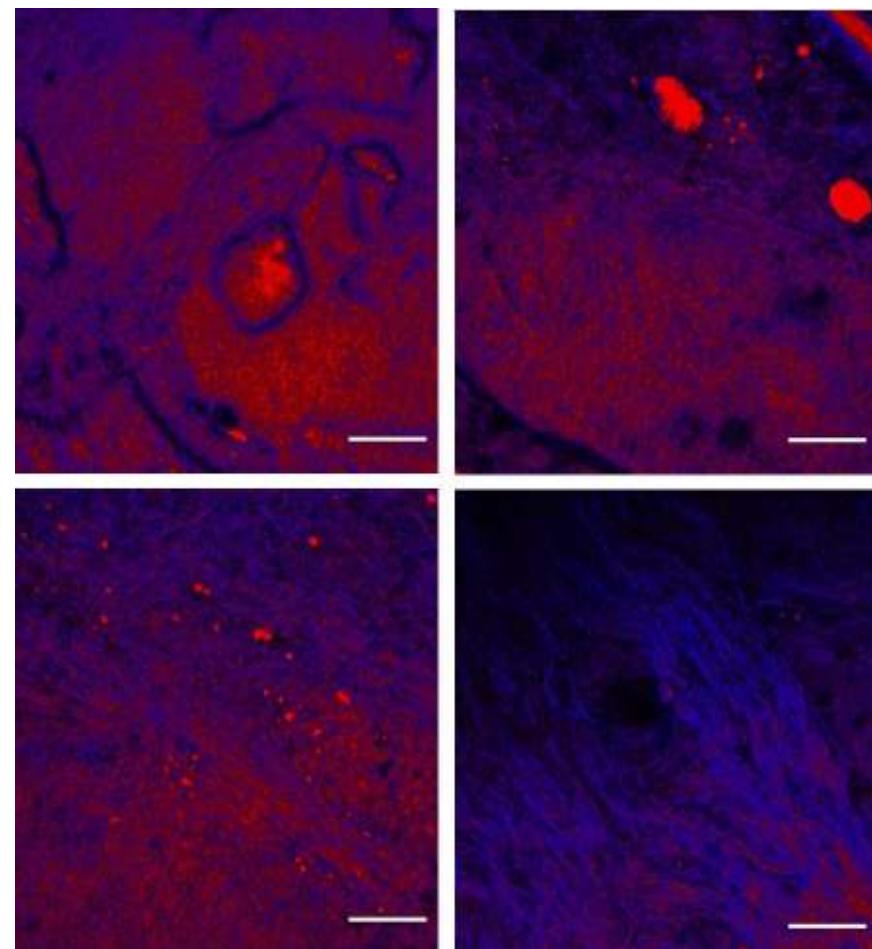
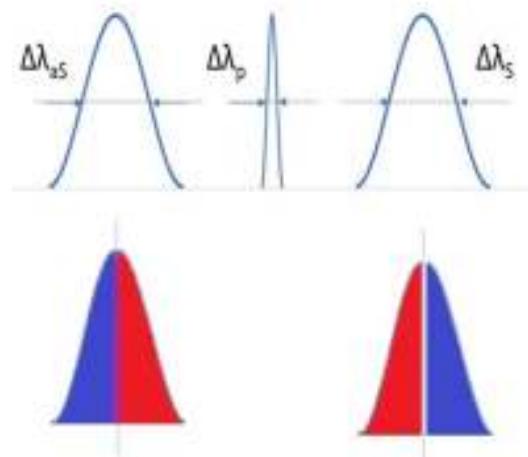
³BOLD Optical Laser Ltd, Kossuth Lajos str 26-31, H-1121 Budapest, Hungary

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Abstract: A 20 MHz repetition rate, sub-ps Ti:sapphire (Ti:S) laser system is proposed for real time, high chemical contrast dual vibration resonance frequency (DVRF) CARS imaging of the skin suitable for *in vivo* histology. © 2020 The Author(s)

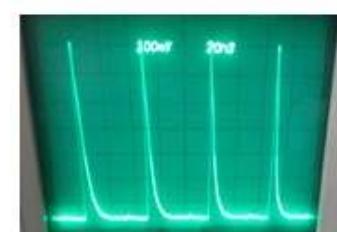
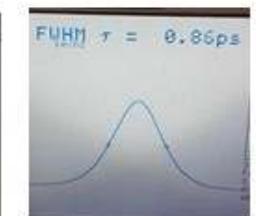
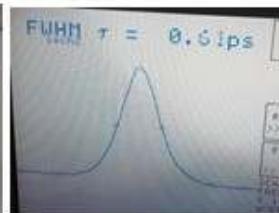
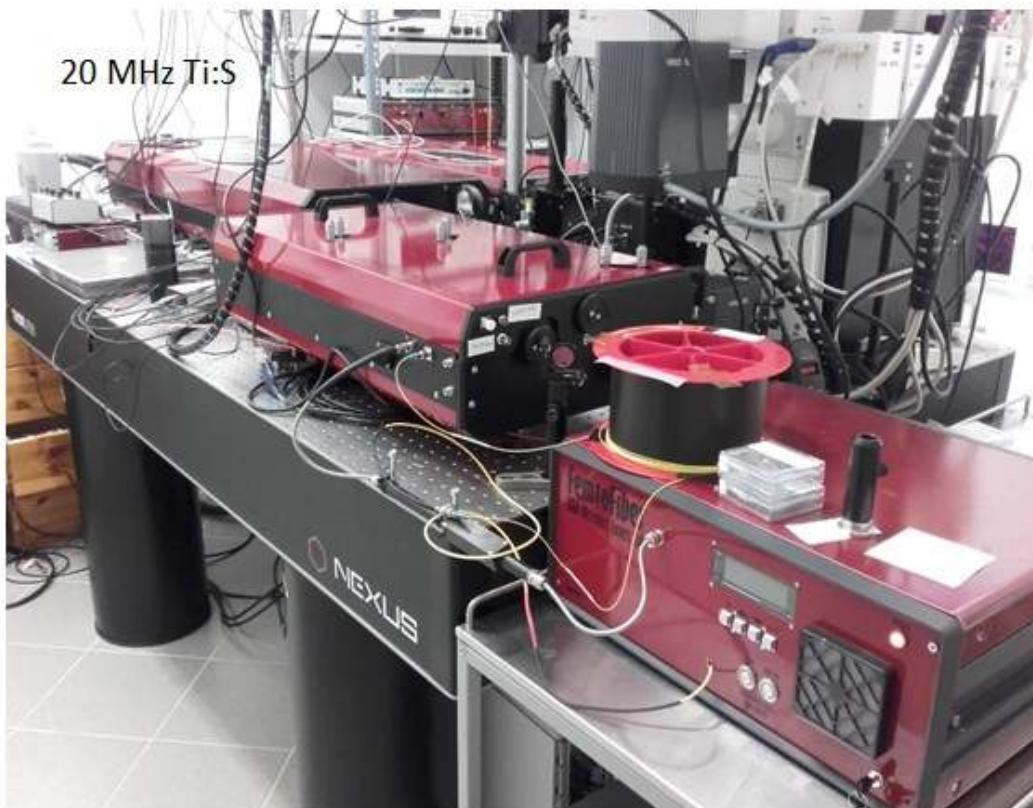
OCIS codes: (140.7900) Ultrafast lasers; (180.6115) Nonlinear microscopy;



Femtosecond Lasers for Nonlinear Microscopy Research Group
(Principal investigator: Robert Szipőcs, Ph.D.)

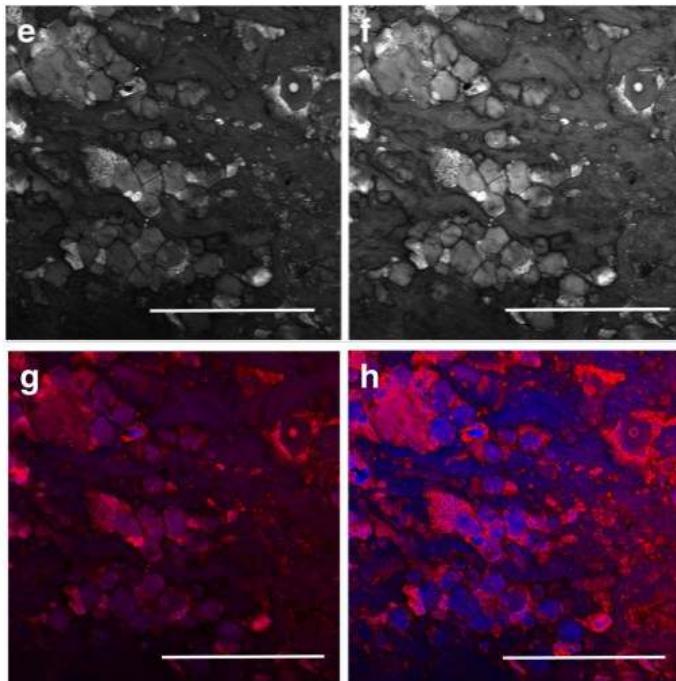


20 MHz, sub-ps, Tunable Ti:sapphire Laser for DVRF CARS imaging



For real time, *in vivo* CARS imaging: **P_{pump}** (794 nm) = **10 mW**, **P_{stokes}** (1030 nm) = **5 mW** (**measured on sample**), PixelDwellTime: **6 μs**

Nonlinear optical microscopy for imaging basal cell cancer



Lipid and protein channels (red and blue color encoding)

Single channel detection

Measurement takes at least 3-5 minutes (changing pump wavelength, set zero delay for pump and Stokes pulses), processing takes also 5-10 minutes

CARS on *ex vivo* basal cell cancer sample

Cite as: C. Cserép *et al.*, *Science* 10.1126/science.aax6752 (2019).

Microglia monitor and protect neuronal function via specialized somatic purinergic junctions

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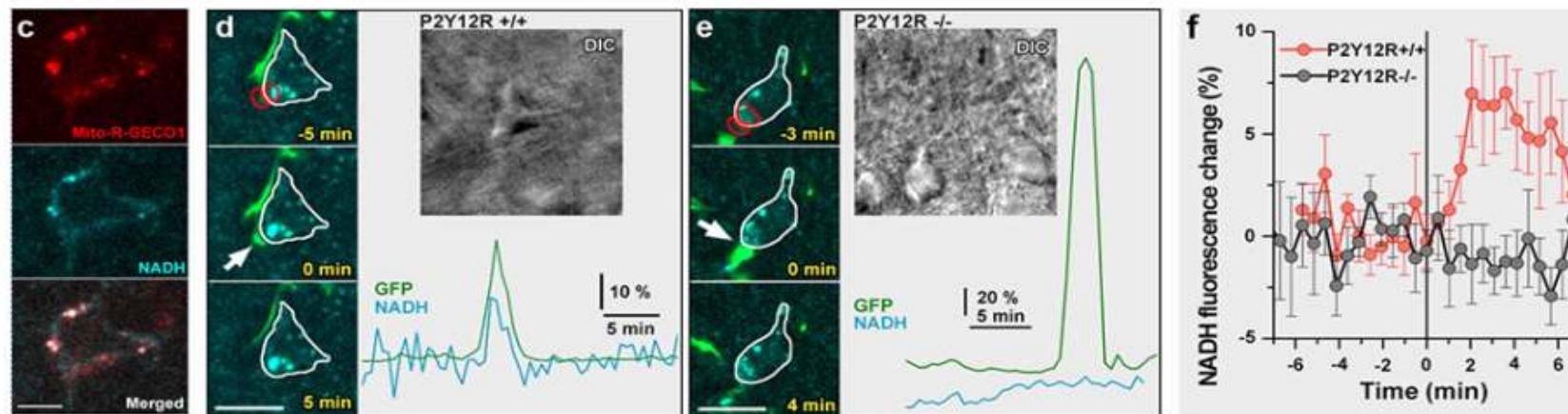
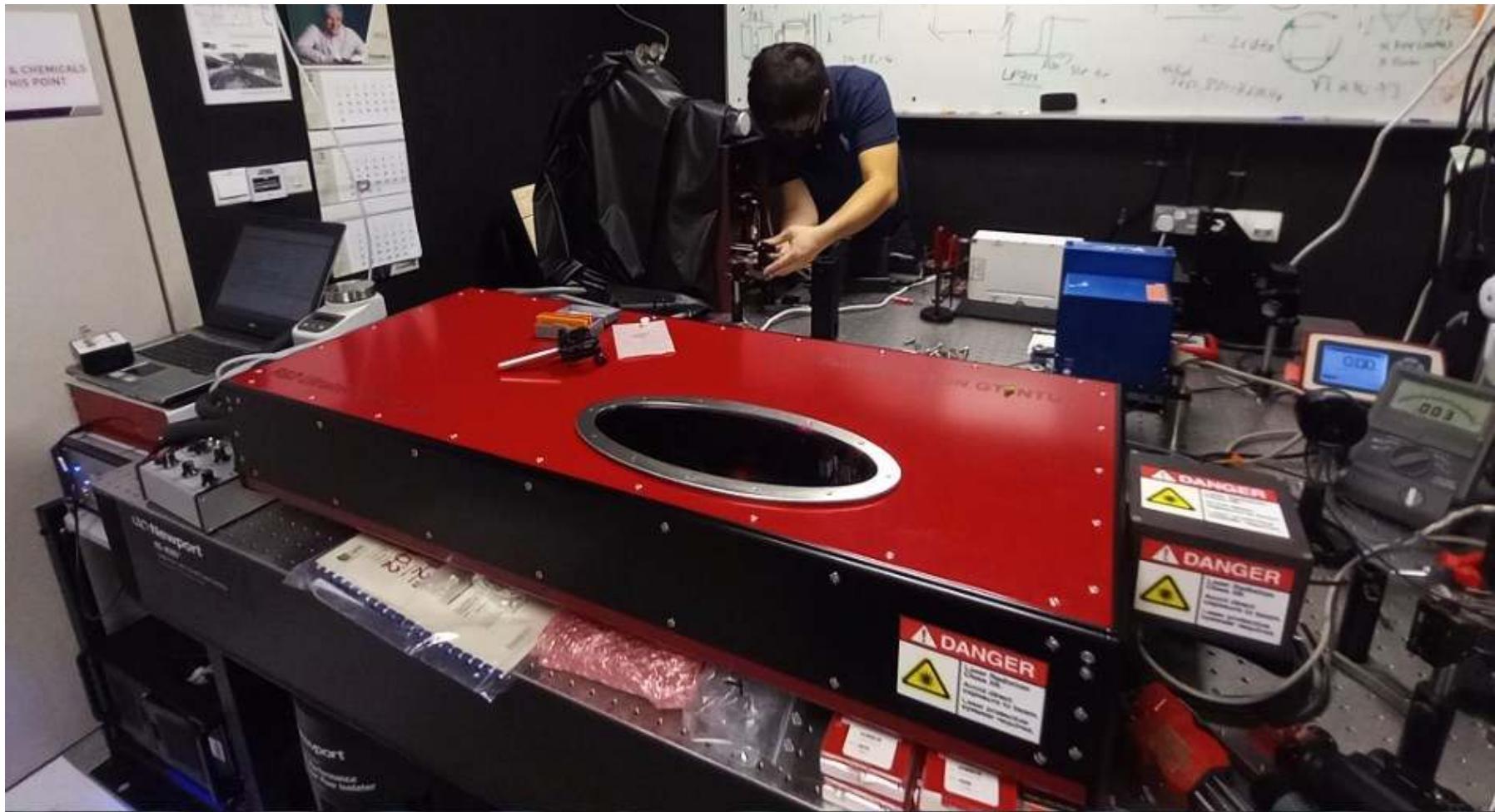


Figure 4. Physiological microglia-neuron communication at the somatic junction site is P2Y12R-dependent and is linked with neuronal mitochondrial activity. a) b) are not shown here. c) Mito-R-Gec01 expression co- nicotinamide adenine dinucleotide (NADH) intrinsic fluorescence. d-e) Representative samples from time-lapse imaging of microglia show processes extend and contact neuronal soma in CX3CR1+/GFP/P2Y12R+/+ (d) and CX3CR1+/GFP/P2Y12R-/- (e) mice. White arrow indicates the contact site of microglia. DIC images of the imaged neurons and the fluorescence signal of GFP (green) and NADH (dark cyan) of red outlined areas are shown. f) Average of NADH intrinsic fluorescence of all neurons in P2Y12R+/+ (red, n=10) and P2Y12R-/- mice (black, n=11).

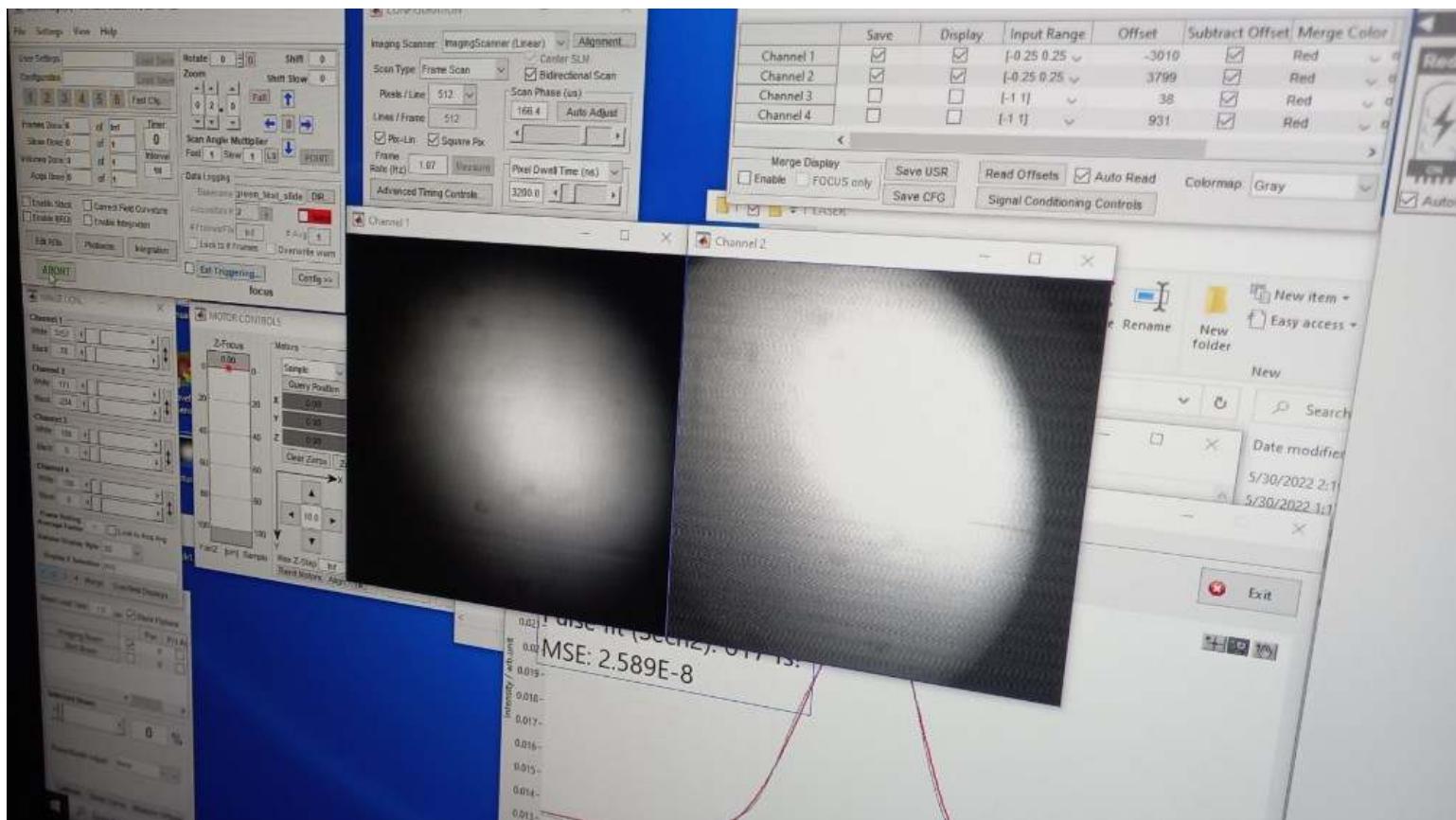
R&D ULTRAFAST LASERS LTD.

Testing for fiber delivery and 2-photon imaging
developed for Alzheimer's disease research for at NTU, Singapore



R&D ULTRAFAST LASERS LTD.

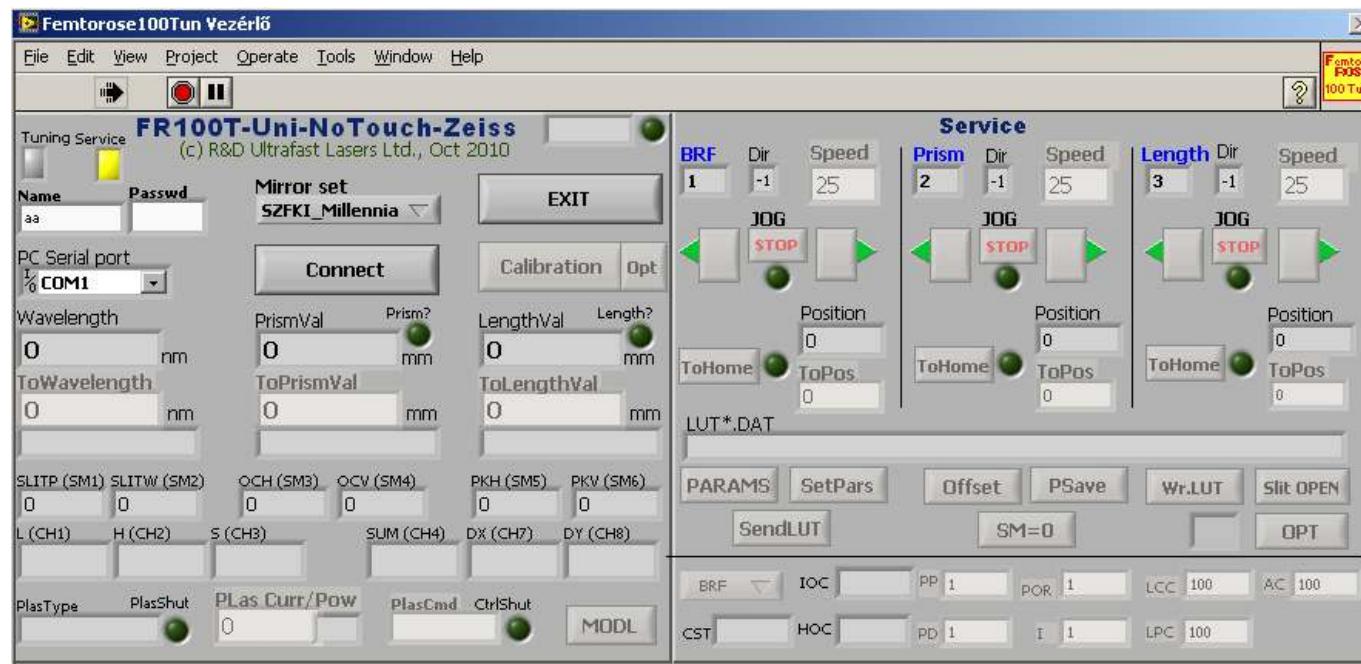
Testing for fiber delivery and 2-photon imaging
at NTU NOBIC lab, Singapore



LASER CONTROL (HW + SW)



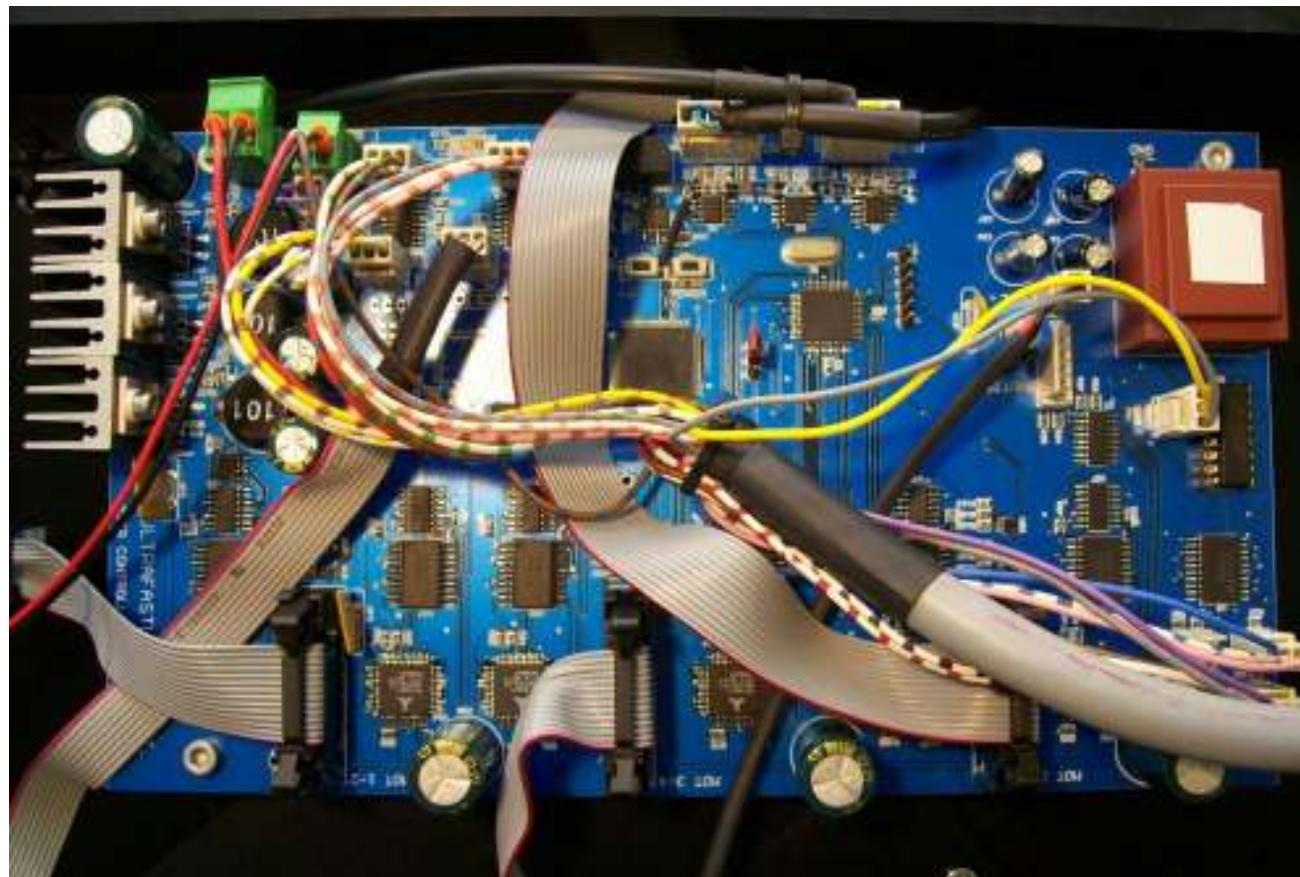
STEP and DC micromotor drivers, SW
Photodiodes, quadrant detectors for beam position sensing
PIC and ARM microcontrollers



ELECTRONICS

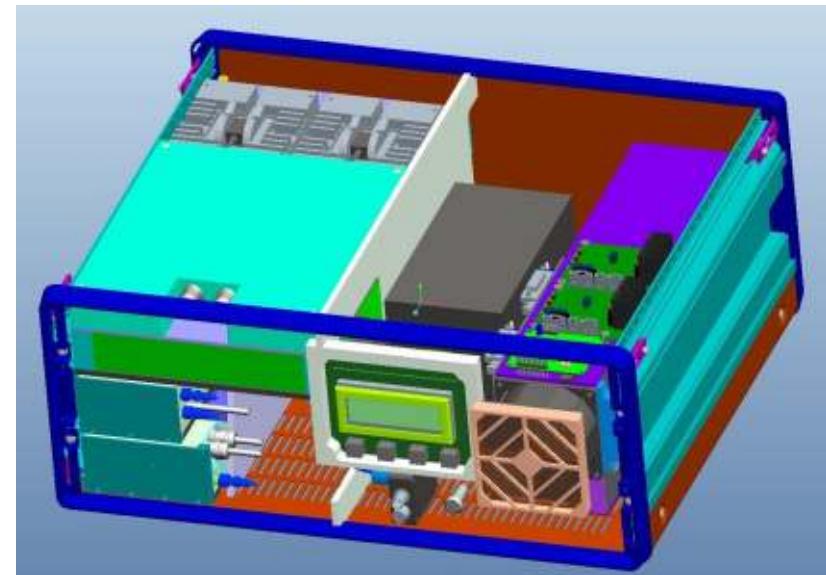
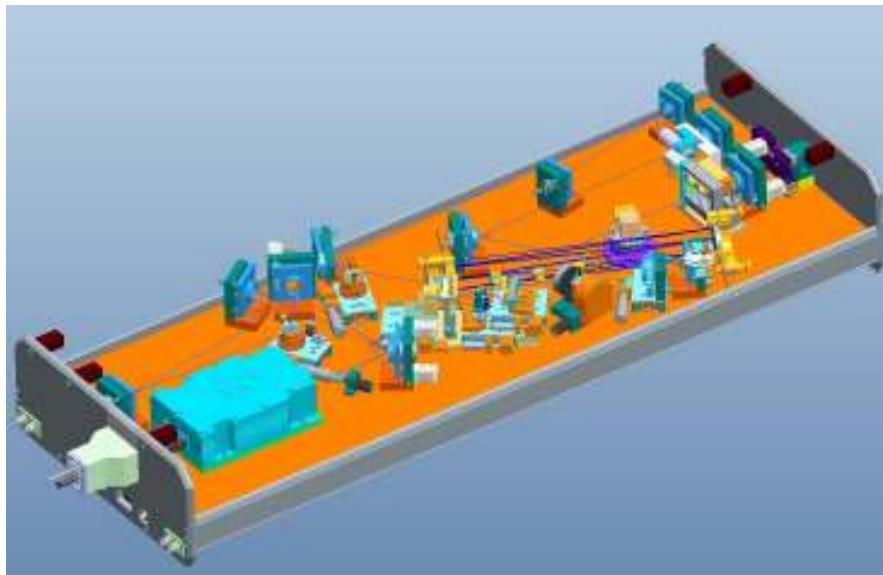


PCB-s with ARM and PIC microcontrollers



MECHANICS

Mechanics for solid state and fiber lasers



ProEngineer models for CNC manufacturing



- **Research and development of ultrafast (ps, or fs) pulse solid state or optical fiber lasers for applications in *in vivo* 3D nonlinear microscopy**

Our aim is to develop laser systems that improve the quality of imaging (e.g. imaging depth, resolution), support easier handling (held-held imaging systems, fiber integration) or minimize the risk of thermal or photochemical damage of the living tissue.

- **Main R&D results:**

- **Development of a long cavity, sub-ps pulse, tunable Ti-sapphire laser operating at ~22 MHz repetition rate utilizing a piezo controlled GTI for intracavity dispersion control**
- **Development of an all-fiber Yb-fiber oscillator and amplifier laser operating at ~2 MHz repetition rate and delivering ~0.5 ps pulses**
- **Development of a two-wavelength sub-ps pulse laser system for CARS imaging applications.**

The research group involved in **development and applications of different label free imaging techniques (2PEF, SHG, CARS, FLIM)** in the fields of dermatology, neurology and metabolic research with its partners at **Semmelweis University (SE)**, **Szeged University (SZTE)** and **University of Sports Science (TF)**, respectively.



Publications of the group

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- [2] **L. Fésüs, Á. Krolopp, G. Molnár, N. Kiss, G. Tamás, and R. Szipőcs**, "A 20 MHz, sub ps, Tunable Ti:sapphire Laser System for Real Time, Stain Free, High Contrast Histology of the Skin," in **Biophotonics Congress: Biomedical Optics 2020**, OSA Technical Digest (Optica Publishing Group, 2020), paper MTh3A.4. (2020).
- [3] **Á. Krolopp, L. Fésüs, G. Szipőcs, N. Wikonkál, and R. Szipőcs**, "Fiber coupled, 20 MHz Repetition Rate, sub ps Ti:sapphire Laser for in vivo Nonlinear Microscopy of the Skin," in **Biophotonics Congress 2021**, OSA Technical Digest (Optical Society of America, 2021), paper DF2A.5. (2021).
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- [7] **G. Szipőcs, Á. Krolopp, S. P. Chong, P. Török, and R. Szipőcs**, "Low Reflection Loss Dispersion Compensation Scheme for Broadly Tunable sub-ps Solid State Lasers," in **CLEO 2023**, Technical Digest Series (Optica Publishing Group, 2023), paper JTh2A.121. (2023).
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