



uTIRF Biodetection Station and iDiagnostics Application Development Kit

iDiagnostics - the Handheld Future of Precision Medicine *accurate, rapid, personalized, yet affordable molecular diagnosing*

Develop *your* custom-design molecular diagnostics using uTIRF and iDiagnostics Application Development Kit

Early detection and prognosis of cancer and other diseases, personalized medicine, prevention of pandemics, food and water safety, and other healthcare tasks require a highly sensitive, accurate, rapid, multiplexed, and yet affordable technology. Eleven years of research at TIRF Labs resulted in the development of such technology - handheld-based platform, termed - iDiagnostics. It is a thousand-fold more accurate and more sensitive than traditional methods. iDiagnostics is rapid and robust, yet inexpensive.

The goal of TIRF Labs is to make iDiagnostics available and affordable to every family on the globe. To facilitate the development of applications, TIRF Labs offers uTIRF biodetection station and Application Development Kit (ADK). We are offering these products to the diagnostic community, including academic, industrial, and government research groups. We will use the Open Innovation Approach to achieve synergy for collaborative development of molecular diagnostics products.

TIRF Labs offers: 1) uTIRF station - turnkey TIRF system, which includes lightguide-, prism-, and objective-TIRF with open perfusion and closed flow chambers, low light CMOS camera, 465-520-637 nm illuminator, 1X, 4X and 60X objectives, and 2) iDiagnostics Application Development kit, which consists of TIRF handheld cradle, cartridge blanks, manual microarrayer, reagents kit, and experimental protocols. This set of products is developed for facilitating biodetection stages of the biomarker discovery stage and bioassay development for existing and emerging biomarkers. We anticipate that collaboration with research groups that are developing and have developed panels of assays for molecular diagnosing will allow for creating superior clinical applications for accurate diagnosis and prognosis of cancer and other diseases.

TIRF Labs will provide comprehensive support for the development of iDiagnostics applications. We have incorporated prior feedback and will incorporate future comments from our users in the next releases of the uTIRF and ADKs. We will supply advanced hardware, software, reagents, development tools, protocols, videos, and other technical support to accommodate the requirements for different diagnostic applications.

Figs. 1, 2 show the optical scheme and photos of uTIRF, IgTRIF accessory, and manual microarrayer. Fig.3 shows iDiagnostics handheld cradle, and reusable cartridges designed for the development stage.

The arrays of bioassays printed at the surface of TIRF slide include assays for detecting protein, nucleic acid, and metabolite markers. The arrays contain internal controls to ensure reliability. The cartridge is equipped with a 20-microliters flow cell, which encompasses the TIRF microarray. iDiagnostics requires no or minimum sample preparation. Whole blood can be analyzed after addition of an anticoagulant. Simple sample preparation modules are included in the cartridge. Additional space is reserved for complex sample preparation modules as will be required for applications developed in the future. See iDiagnostics brochure and visit *i-diagnostics.net* website for more information.

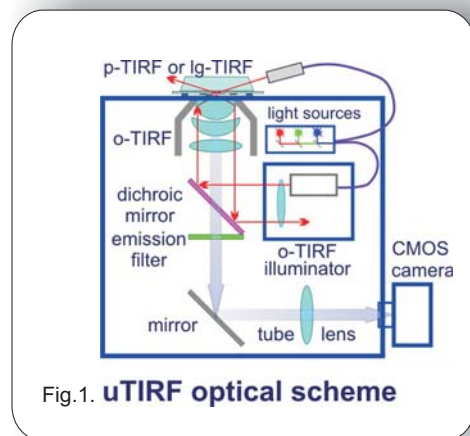


Fig.1. uTIRF optical scheme



Fig. 2. uTIRF with pTIRF installed, IgTIRF - upper right, and manual microarrayer.



Fig.3. iDiagnostics handheld cradle with cartridges.

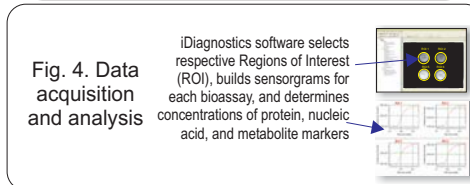


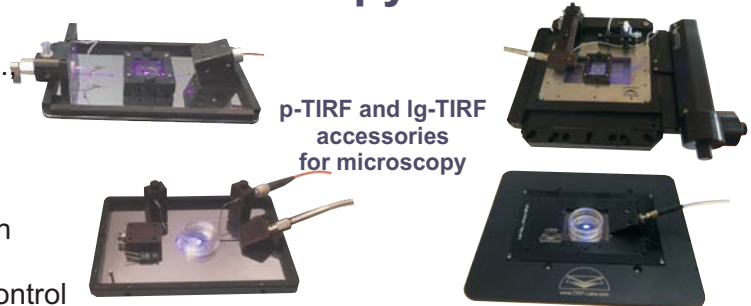
Fig. 4. Data acquisition and analysis

iDiagnostics software selects respective Regions of Interest (ROI), builds sensorgrams for each bioassay, and determines concentrations of protein, nucleic acid, and metabolite markers



Prism- and Lightguide-based TIRF Microscopy Accessories

- Single molecule detection
- Super-resolution microscopy: STED, PALM, STORM, ...
- Minimal stray light, crisp, high-contrast TIRF images
- Work with dry, water-, and oil-immersion objectives
- Use UV or visible excitation light 190-900 nm
- Use Petri-dish, open perfusion, or closed flow chamber
- Nested design - fits inside 96-well plate, K-frame, 4-inch round, or manual XY stages
- Optional temperature, dielectrophoresis, electric field control



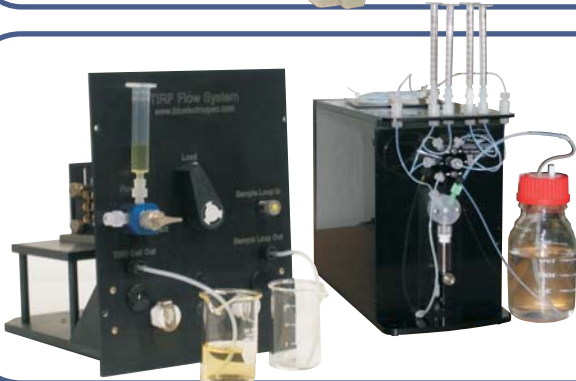
p-TIRF and Ig-TIRF accessories for microscopy

Turnkey Single Molecule Detection TIRF Microscopy Station



- Modular TIRFM stations include:**
- Fluorescence microscope
 - Ig-, p-, or/and o-TIRF microscopy flow systems
 - Low light EM CCD or sCMOS camera
 - Multi-color computer-controlled illuminator
 - Digital fluidics SmartFlow
 - Optional temperature and electric field control
 - Software for instrument control and data analysis

TIRF Accessories for Fluorometers



- **TIRF Accessory TA-1004** transforms a spectrofluorometer into a super-sensitive TIRF biosensor instrument
- Optional electrochemical, DEP and temperature control
- **SmartFlow** Fluidic System allows to run unattended TIRF experiments, measure sensograms to derive k_{on} and k_{off}
- Microfluidic system allows for handling nanoliter volumes

Single ion Channel Single Molecule Detection

fluorescence excitation

patch clamp pipette as light-guide

cell membrane

ion channel

pipette tip transmittance and excitation

pipette tip excitation only

1 micron

SC-SMD on microscope stage

Patch clamp technique combined with fluorescence single molecule detection

iDiagnostics

accurate handheld molecular diagnostics

Real-time TIRF microarrays:
 Parallel supersensitive detection of protein, nucleic acid, and metabolite biomarkers



We extended TIRF into the 3rd dimension and invented iDiagnostics
Now you can hold a hospital laboratory in the palm of your hand