

A Pulsed Live-Cell Quantum Microscope for Entangled Solid State and Biological Qubits

Javier Noé Ramos-Silva,^{1, a)} Sangjun Noh,^{1, a)} Minghao Jiang,² Parisa Aghaei,¹ Ayla Hazrathosseini,³ Jocelyn Leon,³ Philip Hemmer,⁴ and Peter J. Burke^{1, 5, 6, 7}

¹⁾*Department of Electrical Engineering and Computer Science, University of California, Irvine, Irvine, CA 92697, USA*

²⁾*Department of Physics and Astronomy, University of California, Irvine, Irvine, CA 92697, USA*

³⁾*Department of Physics, Texas A&M University, College Station, TX 77843-3128, USA*

⁴⁾*Department of Electrical and Computer Engineering, Texas A&M University, College Station, TX 77843-3128, USA*

⁵⁾*Department of Biomedical Engineering, University of California, Irvine, CA 92697, USA*

⁶⁾*Department of Chemical and Biomolecular Engineering, University of California, Irvine, CA 92697, USA*

⁷⁾*Department of Materials Science and Engineering, University of California, Irvine, CA 92697, USA*

(*Electronic mail: pburke@uci.edu)

(Dated: 7 July 2026)

Two revolutions in quantum sensing are converging on the same microscope stage. Biological qubits have emerged as genetically encoded, optically addressable quantum systems inside live cells. Solid state spin based qubits have been entangled and used as nanoscale correlator magnetometers, delivering sensitivity and spatial-resolution gains that single-spin probes cannot reach. Here we report a pulsed quantum microscope that enables simultaneous quantum state manipulation of both qubit technologies in live cells. The platform combines nanosecond-gated optical excitation at 450 nm and 520 nm, three-dimensional diffraction-limited addressing by galvo beam scanning and piezo objective focus, rapidly switched static and radio-frequency magnetic fields, single-photon timing with picosecond resolution, and microwave control for frequencies from DC to 5 GHz, including the 2.87 GHz resonance of solid state spins, the 500 MHz to 800 MHz resonance of radical pairs in biological qubits, and any future qubit resonance in the GHz range. Live cell imaging with simultaneous biological and solid state nanoparticle qubits in the same cell demonstrates the power of this technique for multiplexed quantum sensing. We anticipate this approach will open new opportunities for researchers to explore quantum sensing in live cells, and, ultimately, entanglement between a solid-state qubit and a protein-hosted spin qubit.

Keywords: Quantum sensing, MagLOV, NV center, biological qubit, quantum microscope

I. INTRODUCTION

Quantum sensors have moved from physics demonstrations to laboratory workhorses over the past decade, and the most consequential advances now sit at the interface between materials-physics qubits and biology. Two parallel lines of progress define the current frontier. On the biological side, spin-correlated radical pairs (SCRPs)^{1,2} and, more recently, genetically encoded fluorescent-protein spin qubits³⁻⁹ have established that a coherent spin degree of freedom can be embedded — and read out optically — directly inside a living cell. On the solid-state side, the negatively charged nitrogen-vacancy (NV) center in diamond has become the archetypal room-temperature optically addressable spin qubit, with optical initialization and readout enabling nanoscale magnetometry, thermometry, and electrometry¹⁰⁻¹².

Within the solid-state arm, the most important recent development is that entanglement has been converted from a

resource for quantum information into a resource for sensing. Superresolution imaging of 2 NVs for gradient magnetometry was demonstrated by one of us in 2010¹³, the same year entangled spin pairs were demonstrated by Neumann et al.¹⁴. Recently theory and experiments have been developed to exploit our initial findings for dramatically improved sensitivity. Rovny et al.¹⁵ prepared Bell states in a pair of NV centers separated by ~ 10 nm and showed that the signal-to-noise ratio for a correlated magnetic-field observable improves by up to $\sim 30\times$ over a non-interacting two-NV baseline, with a $\sqrt{2}$ quantum-correlation advantage and a tractable decoherence-penalty term. In a complementary experiment, Zhou and colleagues¹⁶ engineered a decoherence-free-subspace state, $(|\uparrow\uparrow\rangle + i|\downarrow\downarrow\rangle)/\sqrt{2}$, in a 5.2 nm NV pair and reported a $3.4\times$ sensitivity gain together with a $1.6\times$ spatial-resolution gain, applied to the detection of individual external electron spins. Le and co-workers¹⁷, working in the diffraction-limited regime, used covariance magnetometry between two spectrally resolved NV centers within the same optical spot to access noise correlations over the MHz-GHz band at $15 \text{ nT} \cdot \text{Hz}^{-1/4}$. These three papers, taken together, define the contemporary state of the art for NV-pair quantum sens-

^{a)}These authors contributed equally

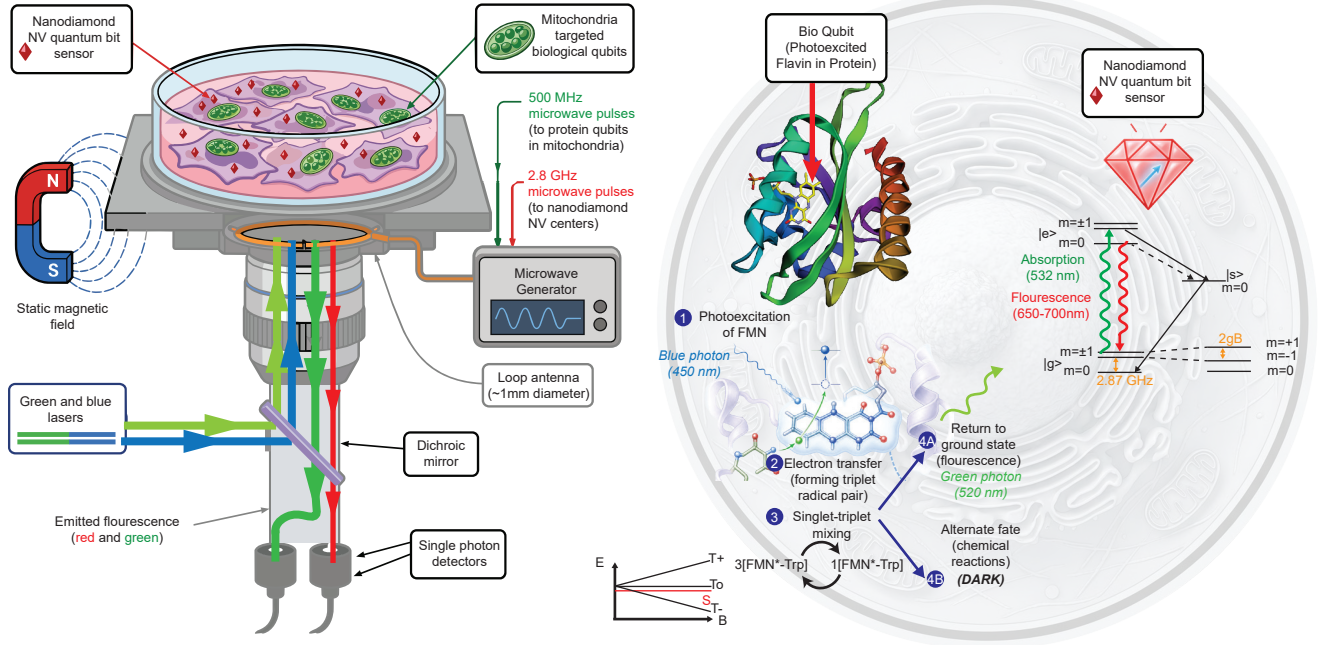


FIG. 1: Conceptual diagram of pulsed quantum microscope showing biological and solid state qubits in the same cell in one system. Full quantum state manipulation of both qubits is enabled by pulsed optics, single photon detectors, and pulsed microwave sources (left). Schematic of the energy levels for the two particular quantum bits demonstrated in this work: Biological quantum bit MagLOV and the associated Jablonski energy levels, showing the flavin absorption and emission events, the singlet-triplet mixing, and effect of magnetic field, as well as the well-known crystal structure of an NV defect in diamond, and the associated Jablonski energy levels (right). Generative AI was used to help in preparation of this figure. Figurelabs.ai (GPT Image 2 model) was used to generate an initial draft, with subsequent modifications by the authors in Adobe Illustrator.

ing. Although the demonstration has required luck since NV pairs are rare birds, advances in materials science and fabrication should enable entangled NV pairs to become a commodity in the next few years. Furthermore, this approach is not limited to NV centers in diamond, as extensive investment in the hundreds of millions of dollars to billions of dollars in spin based solid state qubits in other materials such as silicon is rapidly advancing the ability to entangle spin qubits with arbitrary control^{18–22}. Although this investment is meant for quantum computing, we anticipate it can be translated to quantum sensing in the future. Thus, the future prospects for entangled sensors seems very promising, and only the tip of the iceberg has been shown so far.

A conspicuous absence from this literature is biology. As in our original work¹³, the NV-pair experiments of references^{15–17} are performed on engineered diamond substrates – pillars, implantation-defined arrays, isotopically purified bulk – and none has been applied to, or even explicitly proposed for, living biological targets. We have recently taken a first step in this direction by demonstrating a noise-suppressed NV-pair gradiometer protocol designed with biological operation in mind²³ where the common-mode rejection of two closely-spaced NVs was exploited to null environmental magnetic noise while preserving sensitivity to local gradients of the kind produced by single biomolecules or cell-surface currents, the operating regime in which any future entangled NV-pair

biosensor must work. Additionally, we have demonstrated quantum sensing of magnetic fields with NV centers using an off-the-shelf super-resolution confocal microscope (Zeiss LSM 900 Airyscan) by measuring CW (continuous wave) optically detected magnetic resonance (ODMR)²⁴. The instrument described here is the optical front-end that makes such measurements compatible with inverted, live-cell microscopy in both CW and pulsed regimes.

The biological arm of the field has advanced in parallel. The foundational demonstration is that of Ikeya and Woodward²⁵, who imaged magnetic-field-sensitive endogenous auto-fluorescence in live HeLa cells under sub-25 mT static fields, extracted a MARY curve with $B_{1/2} = 18.0 \pm 0.5$ mT and a saturation magnetic-field effect of 3.7 %, and localized the signal to mitochondrial flavoprotein SCRPs. That experiment was performed in continuous-wave mode: steady-state photoexcitation combined with a low-frequency modulated static field. Continuous-wave detection has a well-known limitation – the observed MFE becomes strongly dependent on excitation intensity because the precursor and intermediate populations are forced into a non-universal steady state – and it offers no time resolution below the tens-of-ms modulation scale. The same group has since solved this problem with two companion techniques, pump-probe (PP) and pump-field-probe (PFP) fluorescence microscopy, in which a pair of nanosecond 450 nm pulses, optionally bracketing a

rapidly switched (< 30 ns) magnetic-field gate, samples the dark-state population and the instantaneous radical-pair lifetime down to ~ 50 ns²⁶. The Ikeya-Woodward PFP method is, to our knowledge, the only published route to time-resolved radical-pair lifetime / magnetic-field-effect (MFE) spectroscopy at cellular spatial scales, and its conceptual core – nanosecond-gated optics synchronized to a rapidly switched magnetic field – has guided the electronic architecture of the instrument reported here.

In parallel, directed evolution has produced genetically encoded proteins whose SCRPs are themselves the sensor. The MagLOV and MagLOV2 family, engineered from the AsLOV2 photoreceptor by C450A mutagenesis followed by iterative selection for large magnetic-field effects, generates a flavin-tryptophan SCRP whose singlet-triplet mixing is coherently driven in continuous-wave RYDMR by ~ 604 MHz radio-frequency fields, yielding RYDMR contrasts of order 50 % in mammalian cells and bacteria and has recently been extended to in-vivo RYDMR in *C. elegans*^{3,7,27}. Feder and colleagues⁴, working with enhanced yellow fluorescent protein, showed that the EYFP photoexcited triplet is a genuine microsecond-coherence spin qubit, with CPMG $T_2 = 16 \pm 2$ μ s, optically activated delayed-fluorescence readout at 912 nm, and coherent microwave control at zero field – inside live HEK293T cells and *E. coli* at room temperature. These two classes of biological qubit are complementary: MagLOV provides a tunable, high-contrast RYDMR line in the RF band, whereas EYFP provides a long-lived, microwave-addressable single-spin object. A pulsed microscope capable of handling both classes, and of doing so on the same optical bench as an NV experiment, does not currently exist in the published literature.

Our own group has recently demonstrated continuous-wave MagLOV RYDMR with super-resolution localization of mitochondrially targeted biological qubits in live mammalian cells⁵. Those experiments use the same inverted microscope body architecture that underlies the present work, in a commercial off the shelf turnkey confocal microscope, but operate in a CW regime and therefore inherit the steady-state limitations identified in references^{25,26}. The natural next step – and one of the principal motivations for the instrument reported here – is the transition from CW to pulsed operation: nanosecond-gated photoexcitation, rapidly switched RF and static fields, and time-resolved detection of the fluorescence response. The platform described in the present paper supplies exactly that capability, and does so in a geometry that is simultaneously compatible with NV-center spin control.

The broader significance of unifying these capabilities is that it opens, for the first time, a credible experimental path to entanglement across the solid-state-biological boundary. Two-qubit entanglement between NV centers has now been demonstrated^{15,16}; two-qubit entanglement between biological qubits – whether two MagLOV SCRPs in adjacent proteins or two EYFP triplets in neighbouring chromophores – has not been demonstrated but is not ruled out by any known physics. The experiment that would matter most, and that no existing platform can address, is the hybrid one: a single pulsed microscope capable of preparing, coherently manipu-

lating, and reading out an entangled state shared between a solid-state NV qubit in a surface-proximal nanodiamond and a biological qubit hosted in a protein in the adjacent cell. Such a hybrid-entangled quantum sensor would merge the long coherence and microwave addressability of the NV center with the genetic targetability and intracellular localization of a fluorescent-protein or MagLOV qubit, and would constitute, in our view, the true holy grail of the quantum interface with biology.

In this paper, we report the design, implementation, calibration, and validation of a pulsed quantum microscope that targets this regime as illustrated in Figure 1. Our specific contributions are four. First, we demonstrate a stationary-sample, galvo-galvo, inverted-microscope architecture that supports the green-excitation / red-collection / microwave-driven protocols required for NV-center ODMR and T_1 spectroscopy, while preserving an optical and sample geometry that can be extended to blue-excitation, nanosecond-gated, RF-driven biological-qubit measurements. Second, we benchmark NV performance on an implanted diamond plate mounted on the stationary Olympus IX71 stage, demonstrating single NV ODMR near 2.87 GHz and $T_1 \approx 1.3$ ms under ambient conditions, which establishes the NV baseline required for future NV-pair sensing protocols inspired by references^{15–17,23}. Third, we demonstrate biological fluorescence compatibility by imaging TMRE-stained and MagLOV-expressing HeLa cells after limited changes to the excitation and detection paths. Four, we demonstrate, for the first time, simultaneous live-cell imaging of biological and solid state qubits in the same cell. These results establish the microscope-level infrastructure needed to pursue future time-resolved RYDMR, NV-pair sensing, and solid-state-biological quantum-interface experiments.

II. SYSTEM DESIGN

A. Confocal Microscope Optical Setup: Imaging

In order to optimize the confocal microscope, it is important to be able to adjust the laser focus position relative to the microscope focus without using the microscope’s own adjustments. A $4 - f$ imaging system is well suited to this end. It works by imaging a conjugate plane — where all beam adjustments are made — onto the back aperture of the objective via the galvo mirrors. In this way, changes in angle or focus do not alter the laser beam size or position on the back aperture. The typical adjustments consist of mirror on a tilt plate for laser beam x, y angles which map to x, y position in the object plane, and a telescope to adjust convergence/divergence of the laser beam which maps to z position. The output lens of the telescope is as close as possible to the mirror, so that both are imaged onto the back aperture. The use of a $4 - f$ imaging system ensures that both laser beam position and curvature are imaged correctly.

The basic design of a galvo-galvo based confocal microscope involves four nested conjugate image planes, as shown in Figure 2. The first is defined by the microscope manufac-

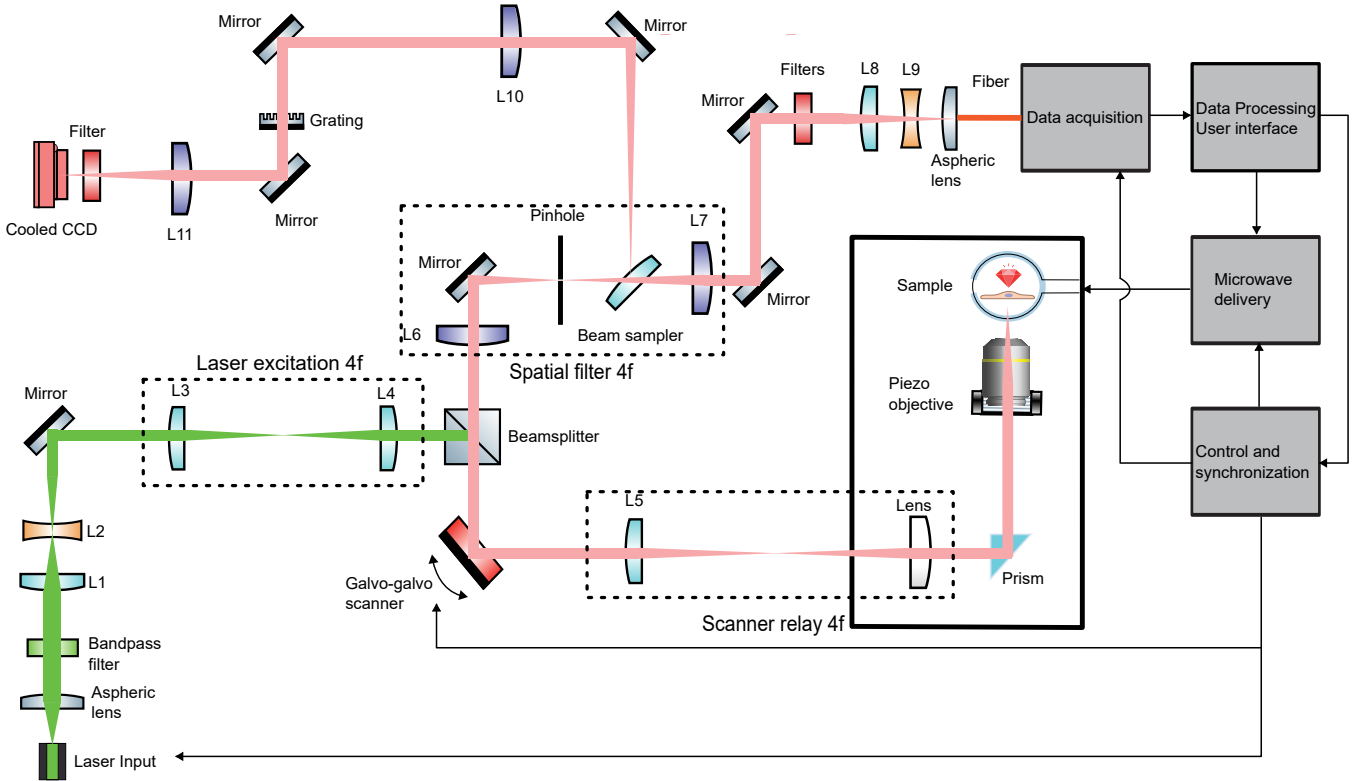


FIG. 2: Schematic of the pulsed quantum microscope platform. The optical path comprises a fiber-coupled 520 nm laser, four nested $4f$ relay systems, a galvo-galvo beam scanner, a confocal pinhole, and a single-photon detector (SPD). The microwave delivery routes a nanosecond-gated signal from the RF generator through a power amplifier and PCB loop antenna positioned beneath the sample. Data acquisition is synchronized with both the optical and microwave channels.

turer through the objective and internal field-lens combination. The second relays this image to a conjugate plane containing the confocal pinhole. The third is a nested imaging system that images the galvo mirrors onto the back aperture of the microscope objective, so that galvo scanning does not cause the laser beam to miss the objective pupil. In some implementations, the galvo may also be used with a matched $f - \theta$ lens, which further constrains the relay design. The fourth images a collimated laser beam, with a size matched to the objective back aperture, onto the galvo mirrors so that small alignment adjustments do not cause beam clipping at the scanner. Each of these imaging subsystems requires $x - y$ (tip-tilt) and z (focus) adjustment. In our implementation, each relay is based on a $4f$ design, which not only forms the required image plane but also preserves the appropriate beam curvature at that plane. Although there is some redundancy between the $x - y - z$ adjustments of the microscope and those of the pinhole, we find that this additional flexibility greatly simplifies both initial alignment and subsequent optimization.

The schematic implementation of this design in our system is shown in Figure 2. The confocal microscope platform is constructed on an Olympus IX71 inverted microscope. This microscope base is the most commonly used in live-cell imaging in biology, and serves as the mechanical platform for dual solid state and biological qubit imaging in this paper. The

setup combines a dual color 520 nm and 450 nm laser excitation path, a galvo-galvo based scanning module, and a single-photon detection path through three carefully aligned $4f$ imaging systems. Each $4f$ relay plays a distinct role in maintaining beam conjugation and minimizing aberrations across the excitation, scanning, and detection pathways. A detailed schematic and parts list are provided in Appendix A and Appendix J, respectively.

Excitation is provided by a single-mode fiber-coupled 520 nm laser for the solid state qubits, and a 450 nm fiber-coupled laser for the biological qubits. The detailed laser path is shown in Figure 3. The excitation light is collimated and guided into the beam-scanning module through the first $4f$ relay. This relay consists of two convex lenses and forms a beam waist at the plane of the galvo-galvo scanner (Thorlabs LSKGG4). The scanner comprises two orthogonally mounted galvanometric mirrors capable of independent angular deflection in the x and y directions. These mirrors introduce time-varying angular shifts into the excitation beam, which are then relayed to scan the beam laterally across the sample.

As shown in Figure 3, the two excitation beams are combined at the input of the first $4f$ relay using a dichroic mirror. The existing 520 nm path, which was aligned first for NV confocal imaging, passes through a bandpass filter before combination. The combined beam is then routed through

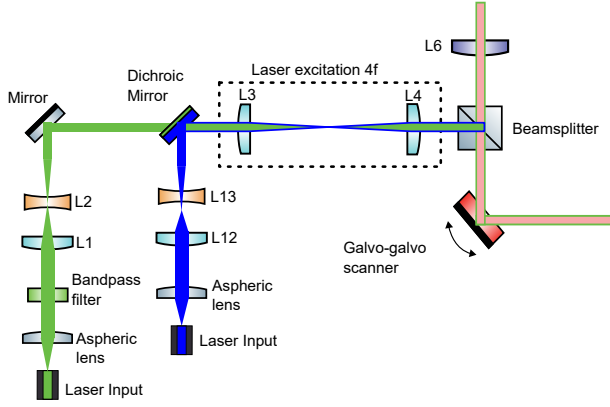


FIG. 3: Dual-excitation configuration at the laser input. A dichroic mirror combines the fiber-coupled 520 nm (NV) and 450 nm (MagLOV) excitation beams into a single collinear path routed through the shared $4f$ relay, galvo-galvo scanner, and microscope objective.

the same optics, galvo-galvo scanner, second $4f$ relay and objective. To convert the angular deflection from the galvo-galvo scanner into lateral displacement at the sample, a second $4f$ relay is used. This relay is composed of two more convex lenses and images the scanned beam onto the back aperture of the microscope objective. In this configuration, the angular deviation imparted by the galvo mirrors is transformed into an approximately linear lateral translation of the beam at the sample plane. Provided that the scanned beam adequately fills the objective aperture, this configuration supports diffraction-limited point excitation across the scan field. The focused beam is delivered to the sample through a $100\times$ high numerical aperture (NA) oil objective lens (Zeiss Alpha Plan-Apo 100x, NA 1.46). Fine axial focusing is performed by moving the objective with a piezoelectric z -stage (Thorlabs PFM450E), which provides closed-loop control over a $450\mu\text{m}$ travel range with 3 nm resolution. This piezo-based control is essential for axial sectioning and for optimizing signal collection from nanoscale NV emitters in the solid state or as nanodiamonds in live cells, and confocal live-cell imaging of biological qubits.

The fluorescence is collected through the same objective and descanned by the galvo mirrors, ensuring that the emission path is optically matched to the excitation path in reverse. The beam is then reflected by a beamsplitter, which separates the red-shifted NV emission from the excitation beam. In our implementation, we use an 10:90 beamsplitter to increase fluorescence throughput. Many reported NV microscope designs instead use a dichroic mirror at this stage, but we find that the beamsplitter-based approach significantly simplifies alignment while causing only a small reduction in detected signal. The emission is then focused through a pinhole located at a plane conjugate to the sample, for this purpose we used an achromatic doublet (Thorlabs AC254-150-AB) which is optimized to provide a nearly constant focal length across a broad bandwidth by minimizing the chromatic aberration of the lens. This spatial filter removes out-of-focus background

and enforces axial resolution by rejecting light that does not originate from the focal plane.

After the pinhole, the beam is relayed through a third $4f$ system, composed of two lenses, which images the pinhole plane onto the entrance of a multimode optical fiber. This fiber delivers the signal to a single-photon avalanche diode detector (SPD, Excelitas SPCM-AQRH-10-FC). A long-pass emission filter is placed before the fiber to further suppress residual excitation light. The TTL output pulses from the SPD are recorded by a Swabian Time Tagger 20, which performs high-resolution time-stamped photon counting for both raster-scanned confocal imaging and time-resolved fluorescence measurements.

As noted above, the use of three external $4f$ relay systems is critical for maintaining optical conjugation and alignment throughout the setup. The first $4f$ system ensures that the laser beam is properly imaged onto the galvo scanner, preserving both collimation and $x-y$ position to avoid clipping on the small (4 mm) galvo mirrors. The second $4f$ system projects galvo deflection onto the objective back aperture to achieve precise lateral beam scanning. The third $4f$ system ensures that the fluorescence signal exiting the pinhole is efficiently coupled to the detector with minimal aberration and signal loss. A spectroscopy arm is also incorporated through a related relay stage that includes a diffraction grating mirror.

The scanning field of view is defined by the galvo-galvo scanner scan range, typically operated below $\pm 2\text{V}$ on each axis. Calibration using a microscope resolution target yields a conversion factor of approximately $24\mu\text{m V}^{-1}$, corresponding to a maximum field of view of about $48\mu\text{m}$. With optimized alignment and stable excitation, the system is capable of sub-micron, diffraction limited spatial localization of solid state spin qubit and biological qubit fluorescence.

The system was assembled inside an isolated cabinet and on top of an anti-vibration table to reduce external optical noise and vibrations. Figure 4 shows a photograph of the completed bench top instrument, in which the galvo-galvo scanner, laser excitation path, and single-photon confocal detection module are mounted on an Olympus IX71 inverted microscope for both solid state and biological qubit imaging.

B. Confocal Microscope Optical Setup: Optical pulsing

Pulsed quantum state manipulation requires that excitation is delivered in short, well-defined bursts with high extinction ratio, synchronized to the MW fields and data acquisition. Two approaches were tested with our optical bench: free-space using acousto-optic modulators (AOMs), and electronic modulation of the fiber-coupled laser diode drive current. We implemented and benchmarked both during platform development; the relative merits, alignment complexity, and cost are discussed below.

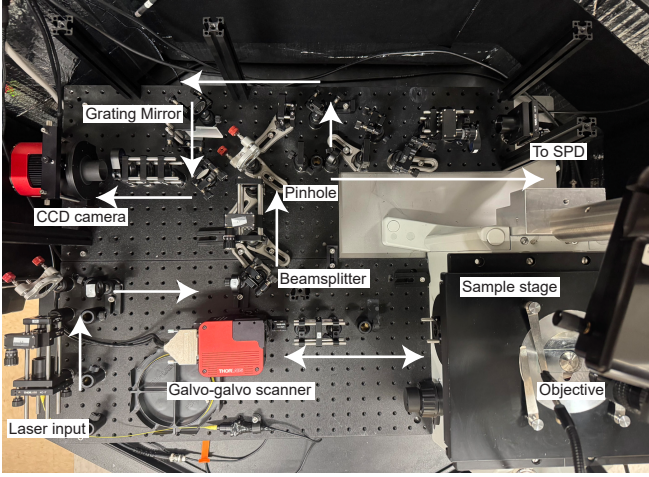


FIG. 4: Photograph of the pulsed quantum microscope showing the inverted Olympus IX71 microscope base (right) and assembled optical breadboard (left).

1. AOM based optical modulation

Most AOMs have an extinction ratio of 40 dB which, while sufficient for coherent quantum state manipulation such as Rabi oscillations, are not sufficient for T_1 lifetime measurements and, ultimately, entanglement, since the small "off" laser power can contaminate the quantum state distribution in sensitive coherence protocols. Some papers²⁸ are satisfied with a single AOM as it can enable some but not all quantum state manipulation. For more demanding quantum state manipulation, most researchers use two AOMs in series or add a double pass setup, giving around 80 dB of extinction ratio, which we have implemented here by combining a free space AOM (Isomet 1205C-2-790) and a fiber coupled AOM (Brimrose TEM-200-25-20-532-2FP), as shown in Figure 5. However, that gives extra complexity, requiring us an extra external optical board for the free space AOM section, and cost, since commercial AOMs are of order a few thousand dollars. We have found recently that commercial AOMs are globally out of stock or no longer manufactured, with months to years of lead time. Used AOMs have also been demonstrated in this work, but with mixed results, as high optical powers can damage the crystals giving visible burn marks in the crystal. One company promised 60 dB of extinction ratio in a single fiber coupled AOM, which we purchased for 5000 USD, but the product (even after a 6 months lead time) did not function properly, was not repaired for many months after sending back to the company, and is no longer offered commercially with that design. Finally, AOMs must be aligned properly in the optical system, and fiber coupled AOMs need care in the fiber mounting and fiber-free space conversion and are not broadband compatible which limits their use for multiple color configuration but it is enough for people working only with solid state qubits as NV-centers.

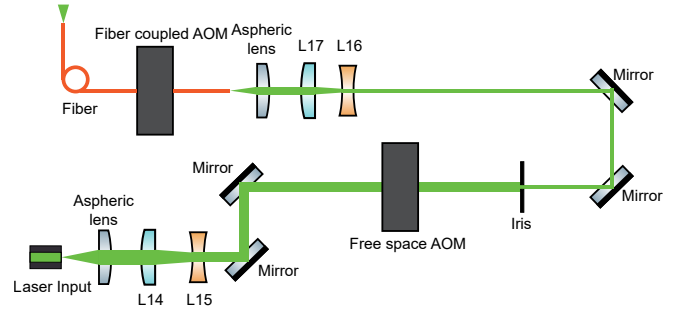


FIG. 5: Schematic of the dual AOM-based laser pulse configuration.

2. Electronic optical modulation

Most modern lasers function as solid state diodes, whose current can be modulated to modulate the optical power. The advantage is the simplicity without the need for additional optics to align and low cost. The disadvantage is the complexity of the current driver circuit for high speed, high current pulses. In addition, laser stability may be inferior when the drive current is not steady, due to thermal drift and transients. Nothing will be more steady than a laser that is turned on in the morning, warmed up, and allowed to stabilize before the run is started. However, we have found the electronic pulsed lasers are more than sufficient for the applications described in this paper. We demonstrate and provide to the community as part of this work an open source current driver that can be manufactured for about 20 USD by a commercial PCB foundry in about a week (including manufacturing and global delivery to anywhere in the world). The detailed design is provided in Appendix F.

C. Microwave System

The system is modular so that different RF subsystems can be swapped out. A general feature is the delivery of high power microwaves to the spin system connected to a pulse electronics setup, synchronized under computer control with the laser pulsing and photon detection setup.

1. Microwave delivery

Three different designs were benchmarked for microwave delivery at the sample for experiments done with this system. The first was a resonant antenna, designed specifically for NV center resonance of 2.87 GHz, based on the paper by Missonou²⁹. Figure 6a shows the geometry of the antenna and Figure 6b the $|S_{11}|$ parameter measured in the laboratory. The advantage of the resonant design is enhanced AC magnetic field at a given power level. The PCB antenna design was adapted from a previously reported layout optimized for quantum control experiments³⁰.

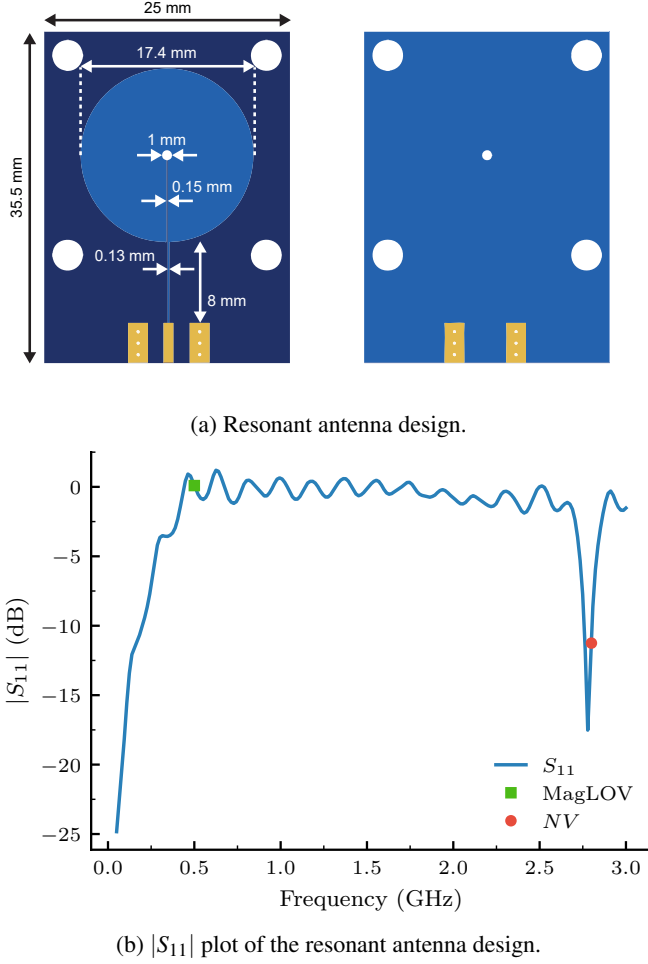


FIG. 6: Antenna design and $|S_{11}|$ plot of the resonant antenna.

The second antenna used in this work was previously reported by Yuan²⁸ and consist of a broadband 1 mm-loop transmission line. These design work properly for solid state qubits. However, they are not compatible with biological qubits. Therefore, a broadband transmission line compatible with petri dish for cell mounting was designed, which consisted of a 1 mm-loop in a 50Ω transmission line, the broadband $|S_{21}|$ flat response, as shown in Figure 7, makes the proposed design compatible with both solid state and biological qubit. This design is fully compatible with petri dishes 35 mm glass-bottom (No. 1.5, MatTek).

2. Solid state qubit RF system

Spin transitions in the NV centers are driven by microwave fields generated and routed through a custom RF chain terminating in a planar loop antenna, as shown in Figure 2³⁰. The chain begins with an RF signal generator (RIGOL DSG836), then that signal is fed into an I/Q Modulator (Texas Instruments TRF3705) where the TTL signal from

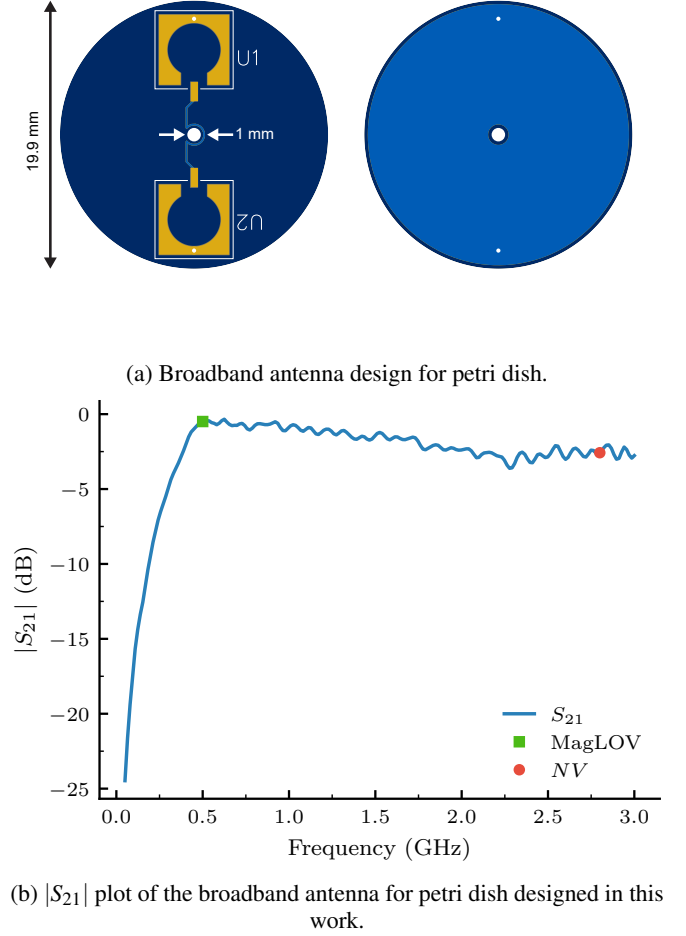


FIG. 7: Antenna design and $|S_{21}|$ plot of the broadband antenna for petri dish designed in this work.

Pulse Streamer enables time-resolved microwave pulse modulation with nanosecond-scale timing control.

The gated microwave signal is then amplified by a high-power broadband amplifier (Mini-Circuits ZHL-16W-43-S+), which delivers up to 16 W over the 1.8 GHz to 4.0 GHz range. To protect the amplifier from reflected power, a microwave circulator (Pasternack PE83CR005) is placed at its output. Then the amplified signal is passed through a step attenuator (Narda 4705B-99), which provides manual control of the microwave amplitude. The attenuator's output is then routed to a printed circuit board (PCB) loop antenna positioned directly beneath the sample.

The antenna is designed to generate an oscillating microwave magnetic field (B_1) that couples efficiently to the NV spin transition. The diamond sample is mounted immediately above the center of the loop, in proximity to the antenna surface while remaining optically accessible from above and electrically isolated from the microwave structure. This geometry provides strong local microwave coupling while preserving the optical working distance required for confocal excitation and fluorescence collection.

3. Biological qubit RF system

The biological qubit system also can undergo ODMR, as we have recently demonstrated in reference⁵. Since the ODMR frequency for MagLOV cells is around 500 MHz, the same setup is used, but the microwave amplifier needs to be replaced by one with the adequate bandwidth (Mini-Circuits ZHL-10M2G0005+) which in the addition to the petri dish compatible loop transmission line designed here allows us to drive the spin transitions of biological qubits.

4. Imaging and qubit quantum control

All the above use a CW generator, then a post-generator microwave I/Q modulator turns the RF on and off. The pulses fed to the switch are generated by the Swabian Pulse Streamer 8/2, which is also used to gate the photon detection and laser. An alternative is to use an FPGA based DAC, and synthesize the RF frequency directly in the digital domain. We have also implemented this solution for the NV centers, using an RFSoc 4x2 board from RealDigital/AMD³¹ which integrates a Zynq Ultrascale+ RFSoc XCZU48DR-2FFVG1517E FPGA. The 2 integrated DACs can synthesize up to 9 GB s^{-1} at 14 bits, so can create up to 5 GHz microwave signals under arbitrary control. An additional advantage is that the photon pulse counting is also completely handled by the RFSoc4x2 board. Finally, the cost is much lower, around 2000 USD for the RFSoc4x2 board, compared to around 18000 USD for Time Tagger, pulse streamer and signal generator solution.

Spin manipulation has been already demonstrated for NV centers elsewhere using QICK-DAWG³² (a fork of QICK³³). Additionally, we demonstrate here for the first time its use for confocal imaging as shown in Appendix D. As our antenna is broadband, using the FPGA solution enables arbitrary RF delivery to both quantum bits in live cells.

D. Solid state qubits

NV centers in diamond emit photoluminescence (PL) under green laser excitation. This emission includes a sharp zero-phonon line (ZPL) at 637 nm and a broad phonon sideband extending to approximately 800 nm³⁴. The ZPL corresponds to the direct optical transition between the ground and excited electronic states of the negatively charged NV center and serves as a key spectral signature for identifying NV PL³⁵.

Several types of diamond samples were used to evaluate the platform across different PL and spin-measurement conditions. For nanodiamond-based measurements, dispersions containing 70 nm nanodiamonds (Adamas, NDNV70nmHi10mL, ~ 100 NVs per particle), 40 nm nanodiamonds (Adamas, NDNV40nmHi1mL, ~ 10 NVs per particle) and 10 nm nanodiamonds (Adamas, NDNV10nmMd1mL, ~ 1 NV per particle) were prepared for confocal imaging and low-density emitter measurements. Equal volumes of the two dispersions were mixed in a 1:19 ratio and then further diluted in a 1 wt% aqueous

solution of polyvinyl alcohol (PVA) to improve dispersion uniformity and adhesion to the substrate. Prior to deposition, the mixture was vortexed and bath-sonicated for 10 min to promote homogeneous suspension. The resulting solution was deposited onto a quartz microscope slide and spin-coated at 3000 rpm for 20 s, followed by baking on a hot plate at 80 °C for 10 min. This procedure produced a well-separated layer of nanodiamonds suitable for confocal imaging and optical targeting of isolated emitters.

For ensemble NV measurements, 15 μm and 150 μm diamond powders (Adamas, MDNV15umHi30mg and MDNV150umHi30mg, NV concentration 3 ppm) were used. A thin layer of cyanoacrylate adhesive (Starbond) was applied to a glass coverslip, and a small amount of NV powder was drop-cast onto the surface. After drying, the coverslip was attached to the PCB antenna, providing stable sample mounting while maintaining compatibility with both optical excitation and microwave delivery. These nanodiamond and powder samples were used during initial platform alignment, confocal scanning calibration, PL-throughput optimization, and validation of the system's ability to locate and image NV emitters across a range of particle sizes. They were not used for the quantitative NV measurements reported below.

For single-NV measurements on a bulk diamond substrate, we used an implanted diamond plate from Institute for Quantum Optics - Universität Ulm. According to the implantation map for this sample, the substrate is an IIa electronic-grade single-crystal diamond (ELSC) implanted with $^{15}\text{N}^+$ ions at an implantation energy of 5 keV and subsequently annealed in ultra-high vacuum at 1000 °C for 3 h. The implanted side contains multiple circular implantation regions with a nominal diameter of 200 μm and implantation doses ranging from 1×10^9 to $1 \times 10^{14}\text{ cm}^{-2}$. Low-dose regions, particularly the $1 \times 10^9\text{ cm}^{-2}$ region, were used for isolating single emitters, while higher-dose regions provided brighter fluorescence for survey scans and alignment. Representative characterization conditions for this implanted sample included 300 μW excitation power in front of the objective, and a 650 nm long-pass filter for NV detection. No ND filter was required for the lower-dose regions, whereas ND filtering was necessary for the highest-dose regions because of the large photon count rates. For the confocal single-emitter localization, ODMR, and T_1 relaxation measurements presented in Section III, we used the implanted bulk diamond plate described before, which provides isolated emitters with well-defined implantation conditions suitable for controlled spin characterization.

E. Biological qubits

Directed evolution of the wild type AsLOV proteins has yielded the MagLOV2 family of proteins, whose fluorescence displays strong dependence on applied magnetic fields^{3,6,7,27,36,37}. The effect is attributed to a SCRPs formed after optical excitation, as sketched for MagLOV in Figure 1. Absorption of blue light promotes the flavin chromophore, after which photoinduced electron transfer generates a donor-

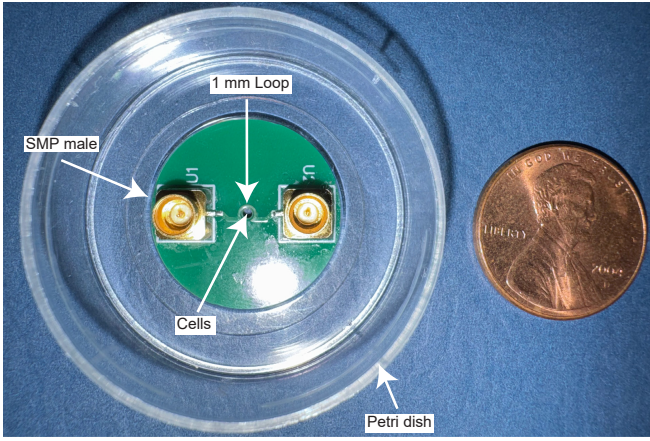


FIG. 8: Broadband loop transmission line mounted on a Petri dish. The antenna fits within the coverslip area of the Petri dish, where the cells are cultured.

acceptor radical pair. The pair may either separate or recombine to repopulate the emissive singlet state and produce fluorescence. Nuclear hyperfine interactions split the entangled radical-pair spin states into singlet and triplet manifolds that evolve at different rates in a static magnetic field, producing a magnetic-field-sensitive fluorescence yield. In MagLOV2, the non-covalently bound flavin mononucleotide (FMN) cofactor participates directly in this photocycle, with protein-side-chain donors within the AsLOV scaffold serving as the electron donor and FMN as the acceptor and primary emitter^{7,27}.

For live-cell experiments, we designed a mitochondria-localized derivative, mtMagLOV2, derived from the R10 Oxford MagLOV2 variant³ after ten rounds of directed evolution as described in⁵.

The fabricated loop transmission line prototype is shown in Figure 8, illustrating how the PCB is integrated with a standard Petri dish while fitting entirely within the coverslip area used for cell culture. Figure 9 shows the sample mounting for experiments with biological qubits using the designed loop transmission line compatible with petri dishes.

F. Software

The confocal scanning system is operated through a custom Python-based software suite designed to integrate the key hardware components of the platform, including the galvo-galvo scanner, DAQ (NI-USB 6453), single-photon detector (SPD), Time Tagger 20 for photon counting, Pulse Streamer 8/2 for spin control, real-time navigation camera, and piezo objective scanner. The software is built around a graphical user interface (GUI) implemented in the Napari visualization framework and supports real-time imaging, photon-count monitoring, and multiple scan modes. The code is organized in a modular structure so that different acquisition and imaging tasks can be executed within a single interface. The software is available through a public repository at [github.com/](https://github.com/burkelabuci/Single_NV_Scanning_Microscopy)

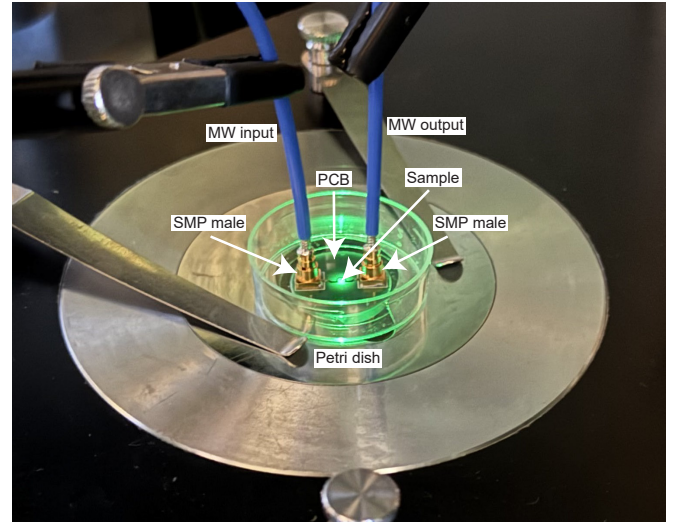


FIG. 9: Schematic diagram of the sample mounting for experiments with biological qubits using the designed loop transmission line compatible with petri dishes.

[burkelabuci/Single_NV_Scanning_Microscopy](https://github.com/burkelabuci/Single_NV_Scanning_Microscopy).

The GUI provides full-field scanning with live feedback as well as zoomed scans of user-selected regions of interest. Scan parameters such as range, resolution, dwell time, and settle time are set directly through the interface. For each acquisition, the software automatically saves the selected scan parameters together with the measured data in timestamped directories, improving reproducibility and simplifying later analysis.

To facilitate alignment and optimization, the software continuously displays real-time photon counts from the SPD in the main GUI. This feature assists in coarse sample positioning, optical alignment, and focal adjustment. For axial optimization, the software controls the piezo objective scanner (Thorlabs PFM450E) to perform automated z-sweeps and determine the focal plane with the highest photon count. The z-sweep range and step size are user-defined, with a typical operating range of $0\text{ }\mu\text{m}$ to $450\text{ }\mu\text{m}$ in $10\text{ }\mu\text{m}$ steps. In addition to two-dimensional raster scans, the software also supports single-axis scans along either x or y , which are useful for beam alignment and for generating position-dependent PL profiles during optimization.

The software further supports live and single-frame image acquisition from a camera. Camera operation is handled in a separate thread so that image streaming does not interrupt the confocal acquisition workflow. In practice, the cameras are used primarily for sample navigation, coarse positioning, and emitter pre-localization before photon-count-based scans are performed.

For each confocal scan, the software generates a raster grid based on the user-defined scan parameters and converts this grid into the corresponding voltage sequence for the galvo-galvo scanner controller through the DAQ. For small scan areas, the use of a high-resolution DAQ is advantageous. The sample is then scanned in a row-by-row raster pattern. Be-

cause the galvo mirrors must reposition at the beginning of each new scan line, an additional settle time is introduced at the start of each row to suppress image artifacts associated with rapid beam motion. At each scan point, the software records the PL signal using the Time Tagger for a defined bin-width. In the present implementation, the resulting counts are normalized to counts per second for display and analysis.

To improve scan fidelity and acquisition speed, photon counting was synchronized to the raster scan through a hardware-based strategy using the Time Tagger CountBetweenMarkers function together with pre-generated x and y axis scan waveforms loaded into the DAQ buffer. In this configuration, the DAQ clock was routed to the Time Tagger to provide hardware-level synchronization between galvo position and photon-count acquisition. This approach substantially reduced timing jitter, eliminated imaging artifacts observed in earlier software-synchronized implementation, and reduced the minimum time per pixel to $\approx 100\mu\text{s}$, limited primarily by the stepping speed of the galvo-galvo scanner.

The software interface also includes galvo positioning, allowing the user to return to any selected point within the scan field for real-time optimization or detailed follow-up measurements. This functionality is particularly useful when identifying isolated bright spots and revisiting them after higher-resolution scans. An autoscale function is implemented to facilitate scans over different spatial ranges. This feature relies on prior calibration of the galvo scan voltage to the real-space sample coordinates. A calibration microscope slide was used for this purpose. This value is incorporated into the software to enable automatic pixel-to-micron conversion during imaging and analysis.

III. EXPERIMENTAL DEMONSTRATIONS

A. Confocal Microscopy

High-resolution confocal imaging was performed using the proposed confocal microscope design. The excitation laser was scanned across the sample by the galvo mirrors, while the resulting PL was collected through the high NA oil-immersion objective, see Section II A. The emitted signal was spatially filtered by a $100\mu\text{m}$ pinhole and detected with a single-photon avalanche diode, enabling background rejection and high-sensitivity photon-count-based imaging.

Figure 10 shows a representative confocal single NV scan acquired from a low-dose implanted region of the bulk diamond plate. This bulk diamond sample was used to localize isolated emitters for follow-up optical and spin measurements. The confocal image reveals multiple spatially separated bright spots within an approximately $100\mu\text{m}^2$ field of view.

The lateral resolution of the confocal microscope was determined from PL intensity profiles of isolated single NV centers, which behave as point emitters much smaller than the diffraction limit, from Figure 10. Line profiles were extracted perpendicular to the scan axis using Fiji/ImageJ³⁸, and the FWHM was evaluated directly from the profile data as shown

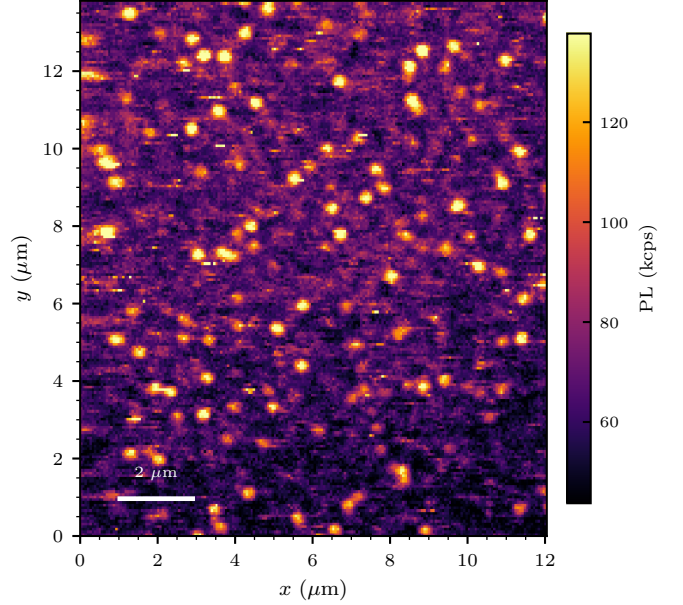


FIG. 10: (a) Confocal PL image of a low-dose implanted region ($1\text{e}9\text{ }^{15}\text{N}^+\text{cm}^{-2}$) in the diamond plate. Hot pixels were corrected by median filtering (3×3 kernel).

in Figure 11 to extract the resolution.

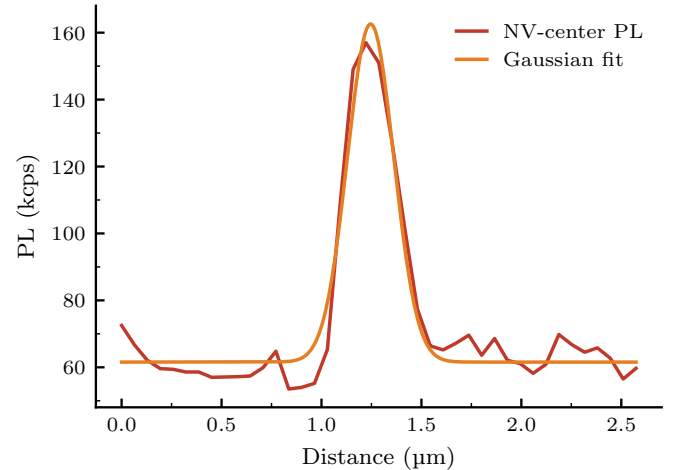


FIG. 11: Lateral resolution of the confocal microscope determined from PL intensity profiles of sub-diffraction Single NV spots from Figure 10.

The half-maximum intensity was calculated as:

$$I_{1/2} = I_{\text{bg}} + \frac{I_{\text{max}} - I_{\text{bg}}}{2}, \quad (1)$$

with $I_{\text{max}} = 1.6 \times 10^5$ counts and $I_{\text{bg}} = 6.0 \times 10^4$ counts, giving $I_{1/2} = 1.1 \times 10^5$ counts. The lateral FWHM was measured to be approximately 300 nm, in good agreement with the Abbe diffraction limit for 520 nm excitation.

$$d = \frac{\lambda}{2NA}, \quad (2)$$

The emission spectrum measured at the marked position is shown in Figure 12. The observed spectrum is consistent with NV-related PL, with the NV^- and NV^0 zero-phonon-line reference positions indicated for comparison. It supports identification of the marked spot as an isolated NV center suitable for subsequent ODMR and T_1 measurements.

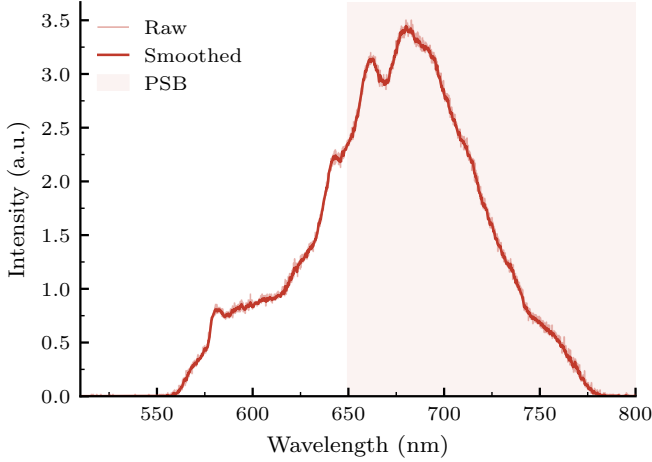


FIG. 12: Single NV emission spectrum.

To obtain these data, the sample was first coarsely positioned using the mechanical stage and navigation camera, followed by galvo-based survey and zoom scans to identify regions of interest. Fine optimization was then performed through piezo-assisted z focusing and real-time photon-count monitoring in the software interface. This workflow enabled efficient localization and verification of candidate emitters in the implanted diamond plate and provided the spatial starting point for the spin-resonance measurements presented below.

These results demonstrate that the platform can resolve isolated emitters in a bulk implanted diamond sample. In the present system, confocal imaging serves both as a characterization tool and as an essential targeting step for spectroscopy and spin-resonance experiments on selected emitters.

B. Optically Detected Magnetic Resonance

Continuous-wave optically detected magnetic resonance (CW ODMR) measurements were performed on the selected NV center. During the measurement, the emitter was continuously illuminated with the 520 nm laser while a continuous microwave drive was applied through the PCB antenna. The microwave frequency was swept across the resonance range, and the PL signal was recorded by the single-photon detector. To reduce the effect of slow laser-power drift, the ODMR spectrum was analyzed using the normalized signal-to-reference ratio.

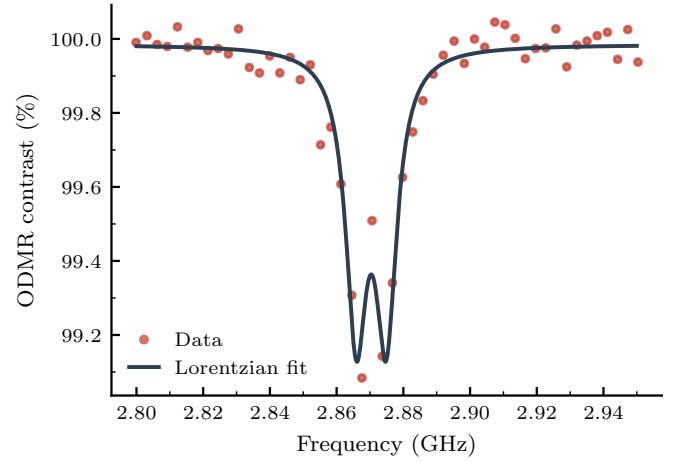


FIG. 13: Single NV CW ODMR spectrum. A clear resonance dip is observed near 2.87 GHz.

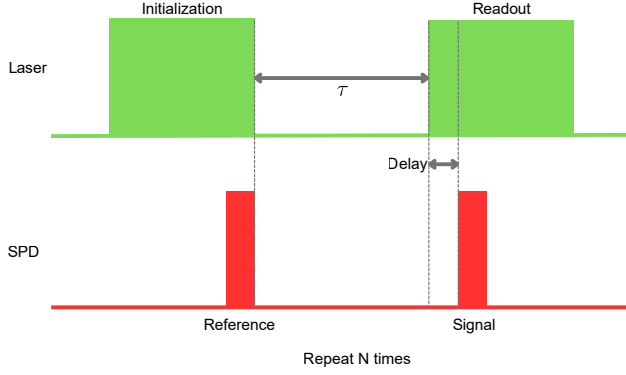
Figure 13 shows a representative CW ODMR spectrum acquired from the selected single NV center in the implanted diamond sample. A clear PL dip is observed near 2.87 GHz, consistent with the ground-state zero-field spin resonance of an NV center. The measured ODMR contrast was approximately 1%, and reproducible spectra were obtained across repeated measurements. The limited contrast is attributed to insufficient microwave power delivered to the sample, a consequence of the current mechanical mounting geometry. An improved sample stage is under development to enhance microwave delivery and ensure full compatibility with bulk crystals, nanoparticles, and biological specimens.

This result demonstrates that the platform supports microwave-driven spin readout following confocal localization. In the present workflow, CW ODMR served as the first spin-resonance measurement on the selected center and provided the basis for the subsequent T_1 measurement.

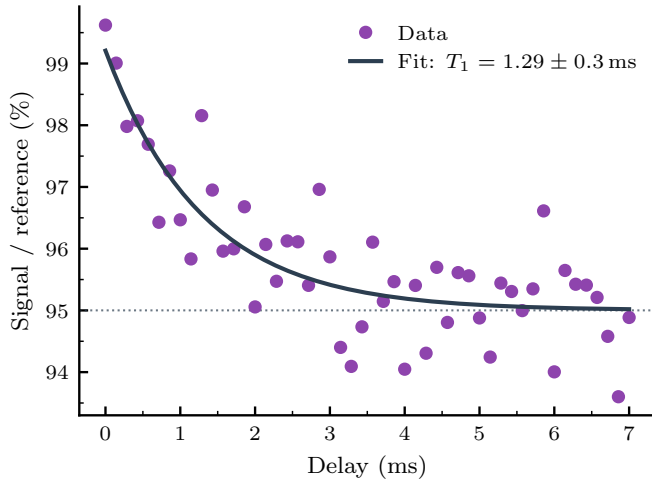
C. Spin Relaxation

Spin relaxation measurements were performed on the NV center selected from the confocal scan. The pulse sequence used for this measurement is shown in Figure 14(a). A green laser pulse was first applied to initialize the NV spin state, and photon counts collected near the end of this pulse were used as the reference signal. After a variable dark interval τ , a second laser pulse was applied for readout. The photon-counting window for the readout signal was opened after a delay of 1.5 μ s, and the ratio of the readout signal to the reference signal was used for normalization in order to reduce sensitivity to slow laser-power fluctuations.

Figure 14(b) shows the measured relaxation curve obtained by varying the delay time τ . As τ increases, the normalized PL approaches its equilibrium value, consistent with longitudinal spin relaxation under ambient conditions. The data were fitted with an exponential decay function, yielding a relaxation time of $T_1 \approx 1.3$ ms.



(a) Pulse sequence used for T_1 measurement. The first laser pulse initializes the NV spin state and provides the reference window, while the second pulse is used for readout after a variable delay τ .



(b) Measured T_1 relaxation curve plotted as signal/reference versus delay time, together with a single-exponential fit yielding $T_1 \approx 1.3$ ms.

FIG. 14: Spin relaxation measurements on the selected NV center in the implanted diamond sample.

This result demonstrates that the platform supports time-resolved spin measurements in addition to confocal imaging and ODMR. In the present workflow, the T_1 measurements were performed after confocal localization and emitter selection, showing that the microscope platform can be used not only to identify NV centers but also to carry out pulsed spin-relaxation measurements on a selected center.

D. Live cell imaging

The inverted, stationary-sample confocal architecture was designed to support both solid-state quantum measurements and biological fluorescence samples. To validate this biological imaging channel, we performed fluorescence imaging of

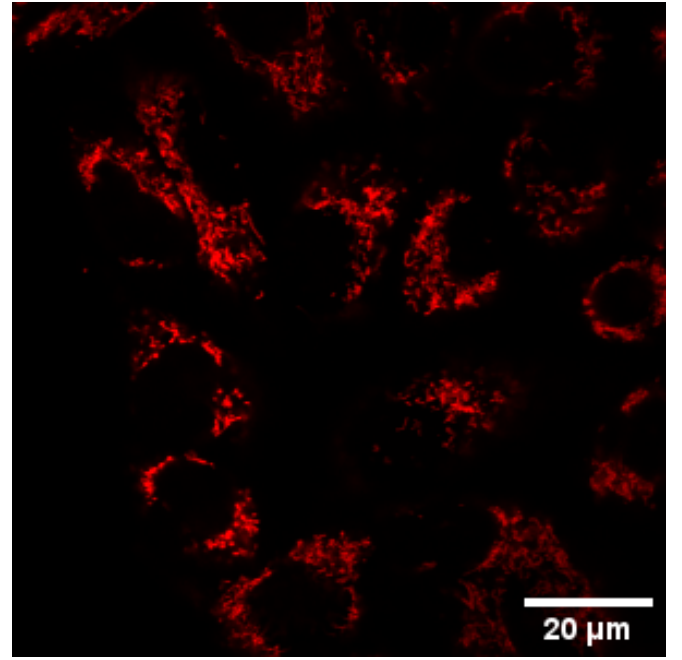


FIG. 15: Confocal image of TMRE-stained HeLa cells acquired using the Olympus IX71-based confocal platform.

HeLa cells. In this configuration, the sample remains fixed on the microscope stage while the excitation beam is raster-scanned by the galvo-galvo scanner.

Representative confocal images of TMRE-stained HeLa cells are shown in Figure 15. TMRE is a voltage dye, only live cells will sustain a mitochondrial membrane voltage. The images resolve the cellular morphology and fluorescence distribution across the sample, demonstrating that the platform can accommodate cell-based fluorescence imaging in addition to nanodiamond and bulk-diamond measurements. The inverted microscope geometry is particularly useful for biological samples because it is compatible with coverslips, culture dishes, and microfluidic sample formats.

E. Biological qubit interrogation

We next applied the platform to HeLa cells expressing MagLOV, a genetically encoded magneto-sensitive fluorescent protein relevant to biological quantum sensing. For this measurement, the excitation source was a Thorlabs LP450-SF25 laser, and the detection filter on the SPD side was replaced with a Thorlabs FBH520-40 bandpass filter to match the MagLOV fluorescence band. The remaining imaging workflow followed the same procedure used for single-NV confocal imaging.

Figure 16 shows a confocal image of the MagLOV-HeLa cells using the proposed Olympus IX71-based confocal platform. This result demonstrates that the same confocal imaging workflow used for NV measurements can resolve MagLOV fluorescence patterns at the cellular scale; the relevant figure of merit for the present platform is single-photon sensi-

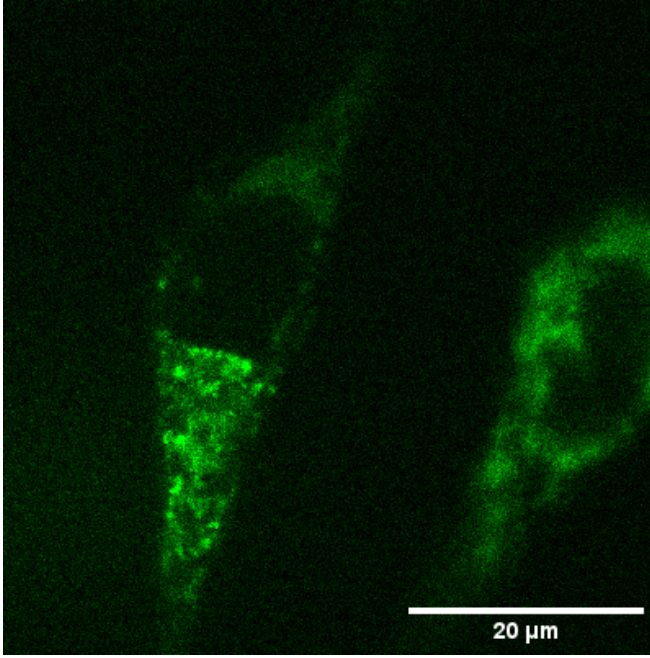


FIG. 16: Confocal image of MagLOV-expressing HeLa cells acquired using the proposed Olympus IX71-based confocal platform.

tivity and galvo-scan compatibility with the NV-center spin-control workflow, both of which are confirmed here. This result demonstrates that the optical framework developed for NV-center quantum experiments can be reconfigured for genetically encoded biological spin systems through only limited changes to the excitation and detection paths.

Together, these cell-imaging results establish the optical compatibility of the platform with biological fluorescence samples and support its extension toward MagLOV-based ODMR/RydMR measurements, biological spin-dynamics studies, and future solid-state–biological quantum sensing interfaces.

F. Biological and solid state Qubit Interrogation

In order to demonstrate simultaneous integration of live cells and solid state quantum sensors, we cultured MagLOV HeLa cells in the presence of nanodiamonds. The nanodiamonds were taken up by endocytosis. In Figure 17 we show a simultaneous image of nanodiamonds and MagLOV HeLa cells. This demonstrates the first example of live-cell simultaneous solid-state and biological qubits in the same system.

IV. CONCLUSION

We have designed, implemented, and initially validated a pulsed quantum microscope built around a commercial Olympus IX71 inverted body that combines solid-state NV-center spin measurements with an optical channel compatible with

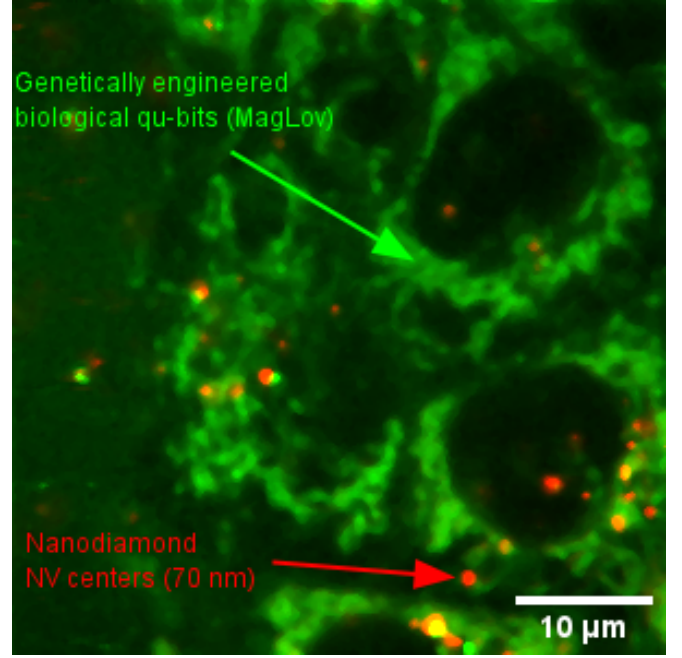


FIG. 17: Confocal image of MagLOV HeLa cells with NV solution added to the culture medium.

genetically encoded biological spin systems within a single stationary-sample, galvo-galvo architecture. On the solid-state side, confocal localization of single NV centers in implanted diamond, combined with CW ODMR at 2.87 GHz and a pulsed spin-relaxation measurement yielding $T_1 \approx 1.3$ ms, this confirms that the platform operates within the sensitivity and coherence envelope established by recent NV-pair quantum sensing experiments^{15–17}. On the biological side, confocal imaging of TMRE-stained HeLa cells and MagLOV HeLa cells – a genetically encoded magneto-sensitive fluorescent protein – demonstrates that the same optical framework supports spatially resolved readout of biological spin systems through only limited reconfiguration of the excitation and detection paths. The coexistence of these two channels on one bench is the central experimental result of this work.

The architectural significance of this coexistence goes beyond convenience. The stationary-sample geometry, in which lateral beam scanning is performed entirely by the galvo-galvo scanner while the sample remains mechanically fixed, is not merely a practical alternative to sample-scanning implementations: it is a prerequisite for the class of experiments that motivates this instrument. Entangled NV-pair sensing^{15,16}, noise-suppressed gradiometry in biologically relevant field environments²³, and time-resolved RydMR at cellular spatial scales²⁶ all require that microelectrodes, RF coils, magnetic-field delivery structures, and microfluidic or cell-culture interfaces remain undisturbed during acquisition. The present platform satisfies this constraint while simultaneously providing nanosecond-gated optical excitation at 450 nm and 520 nm, rapidly switched static and radio-frequency magnetic fields, single-photon timing with picosecond resolution, and microwave control.

The natural trajectory of this work follows three converging lines. The first is the extension of the NV channel from single-spin to two-spin operation: the Bell-state and decoherence-free-subspace protocols of references^{15,16} require no hardware additions beyond a second implantation-defined NV site within the confocal volume, and the T_1 benchmark reported here confirms that the platform's coherence budget is compatible with those protocols. The second is the transition of the biological channel from continuous-wave to pulsed operation: replacing steady-state photoexcitation with the pump-probe and pump-field-probe nanosecond pulse sequences introduced by Ikeya and Woodward²⁶ will unlock time-resolved RYDMR on MagLOV^{7,27} and EYFP⁴ targets with sub-100 ns temporal resolution, removing the steady-state limitation that constrains all current CW biological-qubit experiments including our own⁵. The third and most consequential, line is the hybrid regime: a single acquisition in which a solid-state NV qubit in a surface-proximal nanodiamond and a genetically encoded biological qubit in an adjacent cell share a common optical volume and are addressed by the same pulsed sequence. Demonstrating entanglement across this solid-state-to-biological boundary – merging the long coherence and microwave addressability of the NV center with the genetic targetability and intracellular localisation of a fluorescent-protein qubit – remains, in our view, the true holy grail of quantum bioimaging, and the instrument reported here is architecturally positioned to pursue it.

ACKNOWLEDGMENTS

This work was supported by the Air Force Office of Scientific Research (AFOSR) under awards FA9550-23-1-0061 and FA9550-23-1-0436, and also in part by the National Science Foundation (NSF) under Award 2153425, and the Noyce Foundation. Any opinions, findings, and conclusions or recommendations expressed in this material are those of the author(s) and do not necessarily reflect the views of the United States Air Force. We thank Prof. Dr. Fedor Jelezko and Jens Fuhrmann from Institute for Quantum Optics - Universität Ulm for providing the ion-implanted NV center samples.

DATA AVAILABILITY STATEMENT

The data that support the findings of this study are available from the corresponding author upon reasonable request.

REFERENCES

- P. J. Hore and H. Mouritsen, "The radical-pair mechanism of magnetoreception," *Annual Review of Biophysics* **45**, 299–344 (2016).
- U. E. Steiner and T. Ulrich, "Magnetic field effects in chemical kinetics and related phenomena," *Chem. Rev.* **89**, 51 (1989).
- G. Abrahams, A. Štuhlec, V. Spreng, R. Henry, I. Kempf, J. James, K. Sechkar, S. Stacey, V. Trelles-Fernandez, L. M. Antill, C. R. Timmel, J. J. Miller, M. Ingaramo, A. G. York, J.-P. Tetienne, and H. Steel, "Quantum spin resonance in engineered proteins for multimodal sensing," *Nature* **2026** 649:8099 **649**, 1172–1179 (2026).
- J. S. Feder, B. S. Soloway, S. Verma, Z. Z. Geng, S. Wang, B. B. Kifle, E. G. Riendeau, Y. Tsaturyan, L. R. Weiss, M. Xie, J. Huang, A. Esser-Kahn, L. Gagliardi, D. D. Awschalom, and P. C. Maurer, "A fluorescent-protein spin qubit," *Nature* **2025** 645:8079 **645**, 73–79 (2025).
- P. Aghaei, S. Noh, J. N. Ramos-Silva, M. Jiang, P.-L. Chen, Y. Chen, F. Goodarzinia, R. Usselman, P. Hemmer, D. C. Wallace, and P. J. Burke, "A quantum interface with mitochondrial bioenergetics," *BioRxiv* [Preprint] (2026).
- M. Ingaramo, R. F. Hayward, J. R. Lazzari-Dean, and A. G. York, "Magnetic control of gfp-like fluorescent proteins," in *Quantum Effects and Measurement Techniques in Biology and Biophotonics* (SPIE, 2024) p. PC128630U.
- S. C. Burd, N. Bagheri, A. F. Condon, M. Ingaramo, S. Mondal, D. P. Dowlatshahi, J. A. Summers, S. Mukherjee, A. G. York, S. Wakatsuki, S. G. Boxer, and M. Kasevich, "Magnetic resonance control of spin-correlated radical pair dynamics in vivo," *Nature* **2026** 651:8107 **651**, 940–945 (2026).
- K. M. Xiang, H. Lampson, R. F. Hayward, A. G. York, M. Ingaramo, and A. E. Cohen, "Mechanism of giant magnetic field effect in a red fluorescent protein," *Journal of the American Chemical Society* **147**, 18088–18099 (2025).
- B. L. Ross, A. Lodesani, and C. D. Aiello, "The magnetic field-dependent fluorescence of maglov2 in live bacterial cells is consistent with the radical pair mechanism," *bioRxiv*, 2026–02 (2026).
- M. W. Doherty, N. B. Manson, P. Delaney, F. Jelezko, J. Wrachtrup, and L. C. Hollenberg, "The nitrogen-vacancy colour centre in diamond," *Physics Reports* **528**, 1–45 (2013).
- R. Schirhagl, K. Chang, M. Loretz, and C. L. Degen, "Nitrogen-vacancy centers in diamond: Nanoscale sensors for physics and biology," *Annual Review of Physical Chemistry* **65**, 83–105 (2014).
- C. L. Degen, F. Reinhard, and P. Cappellaro, "Quantum sensing," *Reviews of Modern Physics* **89**, 035002 (2017).
- C. Shin, C. Kim, R. Kolesov, G. Balasubramanian, F. Jelezko, J. Wrachtrup, and P. R. Hemmer, "Sub-optical resolution of single spins using magnetic resonance imaging at room temperature in diamond," *Journal of Luminescence* **130**, 1635–1645 (2010).
- P. Neumann, R. Kolesov, B. Naydenov, J. Beck, F. Rempp, M. Steiner, V. Jacques, G. Balasubramanian, M. Markham, D. Twitchen, *et al.*, "Quantum register based on coupled electron spins in a room-temperature solid," *Nature Physics* **6**, 249–253 (2010).
- J. Rovny, S. Kolkowitz, and N. P. de Leon, "Multi-qubit nanoscale sensing with entanglement as a resource," *Nature* **2025** 647:8091 **647**, 876–882 (2025).
- X. Zhou, M. Wang, X. Ye, H. Sun, Y. Guo, S. Han, Z. Chai, W. Ji, K. Xia, F. Shi, Y. Wang, and J. Du, "Entanglement-enhanced nanoscale single-spin sensing," *Nature* **2025** 647:8091 **647**, 883–888 (2025).
- X. H. Le, P. E. Dolgirev, P. Put, E. L. Peterson, A. Pillai, A. A. Zibrov, E. Demler, H. Park, and M. D. Lukin, "Wideband covariance magnetometry below the diffraction limit," *Physical Review Letters* **135**, 170803 (2025).
- N. Dumoulin Stuyck, A. Saraiva, W. Gilbert, J. Cifuentes Pardo, R. Li, C. C. Escott, K. De Greve, S. Voinigescu, D. J. Reilly, and A. S. Dzurak, "Cmos compatibility of semiconductor spin qubits," *Nature Reviews Electrical Engineering*, 1–16 (2026).
- M. De Michielis, E. Ferraro, E. Prati, L. Hutin, B. Bertrand, E. Charbon, D. J. Ibberson, and M. Fernando Gonzalez-Zalba, "Silicon spin qubits from laboratory to industry," *Journal of Physics D: Applied Physics* **56**, 363001 (2023).
- G. Hu, W. W. Huang, R. Cai, L. Wang, C. H. Yang, G. Cao, X. Xue, P. Huang, and Y. He, "Single-electron spin qubits in silicon for quantum computing," *Intelligent Computing* **4**, 0115 (2025).
- W. Gilbert, T. Tantt, W. H. Lim, M. Feng, J. Y. Huang, J. D. Cifuentes, S. Serrano, P. Y. Mai, R. C. Leon, C. C. Escott, *et al.*, "On-demand electrical control of spin qubits," *Nature Nanotechnology* **18**, 131–136 (2023).
- H. Edlbauer, J. Wang, A. S.-E. Huq, I. Thorvaldson, M. T. Jones, S. H. Misha, W. J. Pappas, C. M. Moehle, Y.-L. Hsueh, H. Bornemann, *et al.*, "An 11-qubit atom processor in silicon," *Nature* **648**, 569–575 (2025).
- M. H. Alkahtani, Y. A. Alzahrani, A. Hazrathosheini, M. Caldarella, J. Leon, P. J. Burke, and P. R. Hemmer, "Noise-suppressed quantum sensing using

- nv pair gradiometer,” *Fluctuation and Noise Letters*, 2650046 (2026).
- ²⁴S. Noh, M. Jiang, J. Leon, P. Hemmer, and P. J. Burke, “Quantum sensing with an off-the-shelf super resolution microscope,” in *Quantum Effects and Measurement Techniques in Biology and Biophotonics II*, Vol. 13340 (SPIE, 2025) pp. 44–49.
 - ²⁵N. Ikeya and J. R. Woodward, “Cellular autofluorescence is magnetic field sensitive,” *Proceedings of the National Academy of Sciences* **118**, e2018043118 (2021).
 - ²⁶N. Ikeya and J. R. Woodward, “A fluorescence microscopy platform for time-resolved studies of spin-correlated radical pairs in biological systems,” *Journal of the American Chemical Society* **148**, 13798–13810 (2026).
 - ²⁷L. M. Antill, J. Baidoo, and L. Gerhards, “Revealing properties for enhanced quantum sensing in engineered proteins,” *bioRxiv*, 2026.02.27.708212 (2026).
 - ²⁸Z. Yuan, S. Mukherjee, J. D. Thompson, N. P. de Leon, A. Gardill, and S. Kolkowitz, “An instructional lab apparatus for quantum experiments with single nitrogen-vacancy centers in diamond,” *American Journal of Physics* **92**, 892–900 (2024).
 - ²⁹D. Misonou, K. Sasaki, S. Ishizu, Y. Monnai, K. M. Itoh, and E. Abe, “Construction and operation of a tabletop system for nanoscale magnetometry with single nitrogen-vacancy centers in diamond,” *AIP Advances* **10** (2020), 10.1063/1.5128716.
 - ³⁰G. Mariani, A. Umemoto, and S. Nomura, “A home-made portable device based on arduino uno for pulsed magnetic resonance of nv centers in diamond,” *AIP Advances* **12** (2022), 10.1063/5.0089161.
 - ³¹Advanced Micro Devices, Inc., “RFSoc 4x2 Kit,” <https://www.amd.com/en/corporate/university-program/aup-boards/rfsoc4x2.html> (2023), accessed: 20 June 2026.
 - ³²E. G. Riendeau, L. Basso, J. J. Mah, R. Cong, M. Sadi, J. Henshaw, K. Azizur-Rahman, A. Jones, G. Joshi, M. P. Lilly, L. Andrew A. Mounce, Basso, J. J. Mah, R. Cong, M. A. Sadi, J. Henshaw, K. M. Azizur-Rahman, A. Jones, G. Joshi, M. P. Lilly, and A. A. Mounce, “Quantum Instrumentation Control Kit — Defect Arbitrary Waveform Generator (QICK-DAWG): A Quantum Sensing Control Framework for Quantum Defects,” (2023), arXiv:2311.18253 [quant-ph].
 - ³³L. Stefanazzi, K. Treptow, N. Wilcer, C. Stoughton, C. Bradford, S. Uemura, S. Zorzetti, S. Montella, G. Cancelo, S. Sussman, A. Houck, S. Saxena, H. Arnaldi, A. Agrawal, H. Zhang, C. Ding, and D. I. Schuster, “The qick (quantum instrumentation control kit): Readout and control for qubits and detectors,” *Review of Scientific Instruments* **93**, 44709 (2022).
 - ³⁴A. Alkauskas, B. B. Buckley, D. D. Awschalom, and C. G. V. D. Walle, “First-principles theory of the luminescence lineshape for the triplet transition in diamond nv centres,” *New Journal of Physics* **16**, 073026 (2014).
 - ³⁵V. K. Sewani, H. H. Vallabhapurapu, Y. Yang, H. R. Firgau, C. Adambukulam, B. C. Johnson, J. J. Pla, and A. Laucht, “Coherent control of nv centers in diamond in a quantum teaching lab,” *American Journal of Physics* **88**, 1156–1169 (2020).
 - ³⁶A. Flores-Ibarra, R. N. Maia, B. Olsasz, J. R. Church, G. Gotthard, I. Schapiro, J. Heberle, and P. Nogly, “Light-oxygen-voltage (lov)-sensing domains: activation mechanism and optogenetic stimulation,” *Journal of molecular biology* **436**, 168356 (2024).
 - ³⁷C. Kim, S. R. Yun, S. J. Lee, S. O. Kim, H. Lee, J. Choi, J. G. Kim, T. W. Kim, S. You, I. Kosheleva, *et al.*, “Structural dynamics of protein-protein association involved in the light-induced transition of *avena sativa* lov2 protein,” *Nature Communications* **15**, 6991 (2024).
 - ³⁸J. Schindelin *et al.*, “Fiji: an open-source platform for biological-image analysis,” *Nature Methods* **9**, 676–682 (2012).
 - ³⁹H. Zhang, C. Belvin, W. Li, J. Wang, J. Wainwright, R. Berg, and J. Bridger, “Little bits of diamond: Optically detected magnetic resonance of nitrogen-vacancy centers,” *American Journal of Physics* **86**, 225–236 (2018).
 - ⁴⁰P. J. Burke, “qlaser: Open-Hardware Pulsed Current Source for Laser Drivers of Quantum Bits,” <https://github.com/burkelabuci/qlaser> (2024), accessed: 20 June 2026.

Appendix A: Alignment Procedure

The alignment procedure consists of three sequential stages: (1) alignment of the microscope, galvo-galvo scanner, and pinhole; (2) alignment of the excitation laser and collection path to the pinhole; and (3) alignment of the pinhole output to the single-photon detector (SPD).

1. Microscope to pinhole alignment

The purpose of this step is to align the output of the camera port of the Olympus IX71 microscope to galvo-galvo scanner. To make the alignment easier, an additional red laser was used. The laser propagates from the microscope objective toward the pinhole through the galvo-galvo scanner.

Careful alignment at this stage is critical because even a small angular error can significantly complicate subsequent alignment procedures. When the red laser enters the microscope objective, its image can be observed at the microscope camera port. The alignment beam is then directed toward the galvo-galvo scanner output port by using an alignment plate and physically moving either the galvo-galvo scanner or the microscope position. The beam position is then verified after the beamsplitter to ensure that the optical axis is properly directed toward the pinhole.

2. Excitation laser to pinhole alignment

There are two goals for this step: (1) to align the excitation laser onto the sample plane and (2) to ensure that the reflected or emitted light from the sample is efficiently coupled through the pinhole.

The alignment begins by steering the green excitation laser through the galvo-galvo scanner using the kinematic mirror in the excitation path and the beamsplitter. A sheet of white paper placed between the scanner and beamsplitter is used to compare the positions of the red alignment laser and the green excitation laser. The green beam is adjusted until it is concentric with the red beam. Once both beams overlap at this location, the remaining optical path is typically aligned automatically because both beams share the same optical axis.

After the beam has been aligned through the microscope, the lenses forming the excitation-path $4f$ relay are installed. The position of each lens is adjusted such that the beam passes through its center. The separation between the lenses must remain equal to the designed $4f$ distance to preserve proper imaging conditions.

The collection path is then optimized using a calibration microscope slide. A calibration target with a minimum feature size of 0.01 mm is placed on the microscope stage, and the objective focus is adjusted until the grid pattern is sharply resolved. A white paper screen positioned above the slide can be used to visualize the illuminated region.

Fine adjustments of the focus are performed until the laser spot is confined to a single grid feature. The sample position is then adjusted in the $x-y$ plane so that a grid line passes

through the center of the laser spot. Additional z -axis adjustments are made until the illuminated region appears dark, indicating that the excitation beam is reflected by the calibration grid rather than transmitted through it. As the sample is translated across the grid pattern, alternating bright and dark regions should be observed.

Once this condition is achieved, a power meter is placed after the pinhole. The kinematic mirror immediately before the pinhole and the axial position of the collection lens are adjusted to maximize the measured optical power. This procedure ensures that the reflected light from the sample is properly focused through the pinhole.

3. Pinhole to the SPD alignment

The final stage consists of coupling the fluorescence signal exiting the pinhole into the SPD. Ideally, the beam is relayed to the detector through the fluorescence-path $4f$ system. In practice, additional optics may be introduced to improve coupling efficiency by reducing the beam diameter and matching it to the acceptance conditions of the SPD fiber collimator.

For alignment, the optical fiber is temporarily disconnected from the SPD and connected to a calibrated optical power meter. The kinematic mirrors located after the pinhole are adjusted to optimize the beam position, while the axial positions of the coupling lenses are adjusted to match the beam size to the fiber input.

The alignment is considered optimized when the optical power coupled into the multimode fiber is maximized. In our setup, coupling efficiencies of approximately 80 % are routinely achieved. After optimization, the optical fiber is reconnected to the SPD for fluorescence measurements.

Appendix B: Standard Confocal Imaging Procedure

The following procedure was consistently used for image acquisition with the Olympus-based confocal microscope platform. This protocol ensures optimal PL signal collection from NV centers embedded in diamond particles.

First, the sample is mounted on the stage of the Olympus IX71 inverted microscope. The z -axis position is manually adjusted while monitoring real-time photon counts to bring the sample into approximate focus. Once the photon counts have been optimized, a broad scan is initiated. This generates an initial low-resolution image of the entire field of view.

From the acquired image, a bright spot of interest is selected, and a zoom scan is performed in the software to refine the spatial region. This zooming and scanning step can be repeated multiple times until a clearly defined region is identified. After determining the desired target area, the laser beam is manually positioned at the center of the selected bright spot, and the autofocus routine is executed. This triggers the piezo objective (z -axis) to perform a fine scan and automatically select the focal plane with the highest photon count.

To further maximize the signal, the user manually fine-tunes the microscope's mechanical z -stage and iteratively ad-

justs the x/y positions of the kinematic mirrors of the excitation and collection paths. These minimal adjustments are repeated until the photon counts are optimized.

Finally, a high-resolution scan is performed over the region of interest to obtain a diffraction-limited image suitable for subsequent analysis.

Appendix C: Troubleshooting

1. SPD saturation

When the SPD becomes saturated distinct horizontal artifacts appear in the scan image as shown in Figure C1. These artifacts manifest as bright horizontal lines across the image, corresponding to rows where the photon count exceeded the detector's linear response range. Saturation can also be confirmed by a visual alarm implemented in the real time acquisition window in the GUI software. The issue was mitigated by introducing neutral density filters as needed in front of the detector. For low-emission samples no ND filters were required due to significantly reduced PL intensity.

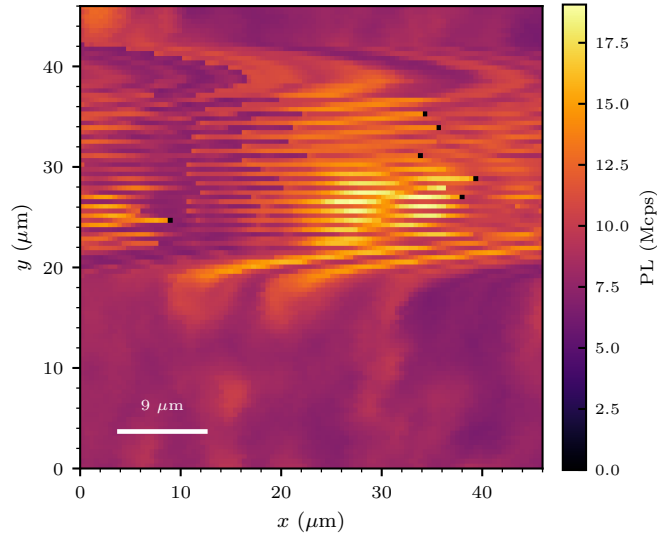


FIG. C1: Confocal imaging artifacts when SPD saturation occurs.

2. Image edge artifacts

Blurring and non-uniform brightness near the left edges of scan images were consistently observed as observed in Figure C2. These artifacts were traced to rapid galvo mirror movements without sufficient stabilization time. Adding a short delay before each scan line effectively suppressed these edge artifacts and was implemented in the software. The delay time depends on the technical details of the galvo-galvo scanner.

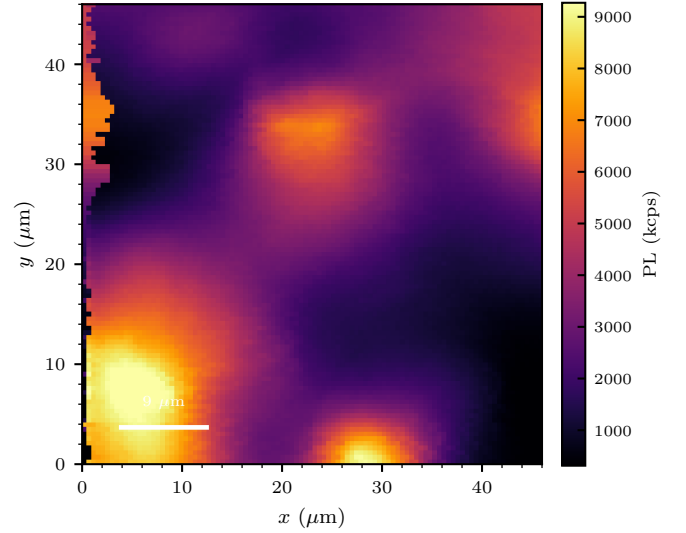


FIG. C2: Artifacts at the left of the image caused by fast scans.

Appendix D: FPGA-based (Xilinx RFSoc 4x2 board) data acquisition and experiment control

As an alternative to the proposed hardware configuration, the system was also tested using the Xilinx Radio Frequency System-on-Chip (RFSoc 4x2) board to acquire data during the imaging process and to control quantum sensing experiments via the QICK-DAWG framework³². A branch called `feature/RFSoc4x2` is available on our GitHub repository.

The primary advantage of the RFSoc 4x2 board is a simpler hardware configuration: the microwave pulsed signal is synthesized directly on the board, in the same manner as the read-out signal is read and analyzed in real-time. Pulse sequence programs and data analysis are provided through the QICK-DAWG python interface, which supports the collection and characterization of PL intensity, ODMR, T_1 relaxation time, Rabi oscillations, Ramsey, and Hahn echo sequences.

However, in its current version the QICK-DAWG framework is not optimized for confocal imaging, resulting in longer acquisition times compared with the Time Tagger approach. For a 100×100 pixel image, the acquisition time using the RFSoc 4x2 board was ≈ 180 s, whereas the Time Tagger required only ≈ 20 s. An example image acquired using the RFSoc 4x2 is shown in Figure D1.

A further limitation of the RFSoc 4x2 and QICK-DAWG combination arises in quantum experiments with a single NV center, where the signal intensity is inherently low. In the current implementation, the maximum integration time per acquisition window is limited to $213 \mu\text{s}$, a constraint imposed by the finite bit-depth of the tProc internal counters in the QICK firmware (base firmware of QICK-DAWG)³³. This restriction has a significant impact on the signal-to-noise ratio (SNR) of single-NV measurements, which are fundamentally limited by shot noise.

In photon-counting experiments, the shot noise arises from

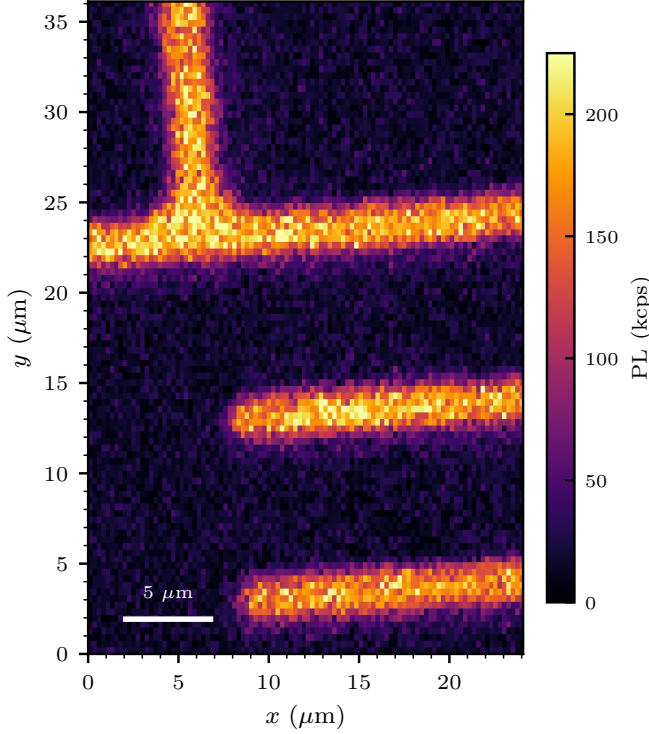


FIG. D1: Confocal image acquired using the RFSoc4x2 FPGA board.

the Poissonian statistics of photon detection events. For N detected photons, the shot noise standard deviation is $\sigma = \sqrt{N}$ yielding a contrast-weighted SNR that scales as

$$\text{SNR} = \frac{C\sqrt{MN}}{\sqrt{1+C}} \quad (\text{D1})$$

where C is the measurement contrast, N is the number of photons collected per repetition, and M is the total number of repetitions. In our confocal system, using a $100\mu\text{m}$ pin-hole, a typical single NV center yields a photon count rate of 10kcps to 20kcps in the bright state. Within the maximum integration window of $213\mu\text{s}$, this corresponds to only $N = 2 - 4$ photons per acquisition window. For a typical Single-NV ODMR contrast of $C \approx 0.03 - 0.05$, achieving a target $\text{SNR} \geq 10$ from Eq. D1 requires the number of repetitions to satisfy

$$M \geq \frac{10^2(1+C)}{C^2N} \approx 10^4 \text{ to } 6 \times 10^4 \quad (\text{D2})$$

Substantially increasing the total acquisition time and limiting the practical throughput of the system for single-NV quantum sensing protocols such as ODMR, T_1 , Rabi, Ramsey, and Hahn echo sequences.

Despite these current limitations, active development is underway to optimize the QICK-DAWG framework for confocal imaging and single-NV quantum sensing experiments using

the RFSoc 4x2 board. Efforts are focused on achieving imaging speeds comparable to the Time Tagger approach. Full integration of the RFSoc 4x2 into our confocal platform is particularly compelling because it consolidates microwave pulse synthesis, readout, and data acquisition into a single board, substantially reducing hardware complexity relative to the configuration described in Figure 2. Furthermore, the significantly lower cost of the RFSoc 4x2 board compared to dedicated hardware lowers the barrier to entry for laboratories seeking to implement NV-based quantum sensing and confocal imaging without access to expensive instrumentation, broadening the accessibility of these techniques to the wider research community.

Appendix E: Live cell imaging using Zeiss LSM 900 super resolution confocal microscope

In order to have a comparison point for our Olympus-based confocal microscope for MagLOV cells with NVs, we used a Zeiss LSM 900 confocal microscope equipped with an Airyscan detector. A $63\times$ oil-immersion objective with a NA of 1.4. to image MagLOV cells fluorescence. The sample was excited using a 488 nm laser at 1 % laser power, and emission was collected between 500 nm to 550 nm. The results are shown in Figure E2 and the sample was the same as the one reported in this work, see Figure 17.

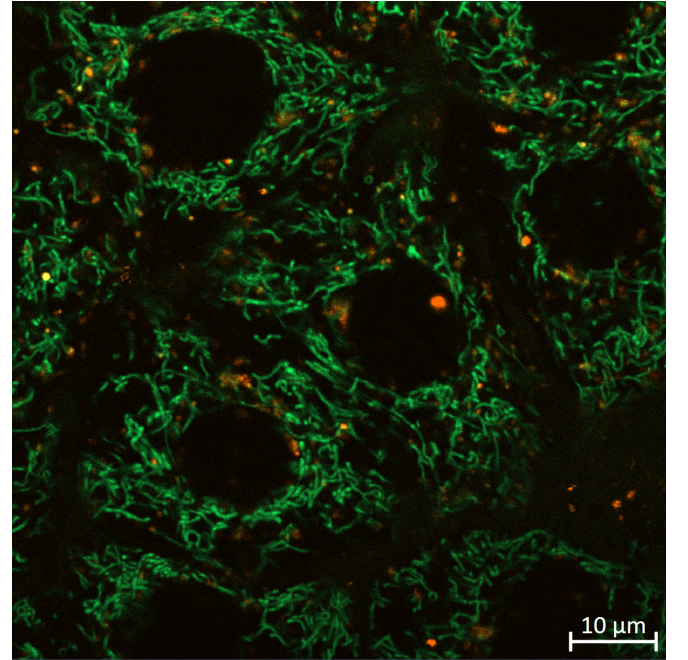


FIG. E2: Dual-color imaging using Zeiss LSM 900 super resolution confocal microscope.

Appendix F: Laser diode current driver

As an alternative to acousto-optic modulators (AOMs) for laser power modulation, we developed an electronic laser pulse driver capable of directly modulating the current of a laser diode. The design is based on the work of Sewani et al.³⁵, but incorporates several important improvements. First, the biasing network was redesigned to minimize transient ringing during pulse operation. Second, the entire circuit was migrated to a commercially manufacturable printed circuit board (PCB) design using EasyEDA and released as an open-source hardware project on GitHub (<https://github.com/burkelabuci/qlaser>).

The development of this driver was motivated by the limited availability and long procurement times of AOMs, which are commonly used to generate the laser pulses required for quantum-state initialization and readout. To avoid this bottleneck, we investigated direct current modulation of laser diodes, such as the Thorlabs LP520-SF15A operating at 520 nm.

The resulting pulse driver, shown in Figures F3 and F4, provides a low-cost and readily accessible alternative to an AOM. The complete hardware design is open source and available at <https://github.com/burkelabuci/qlaser>. While the original design reported by Sewani et al.³⁵ required manual assembly and hand soldering of individual components, our implementation was specifically optimized for commercial PCB assembly. All components, including connectors, can be factory assembled, allowing the board to be delivered fully populated and ready for use without additional fabrication effort.

In addition to simplifying manufacturing, the PCB layout incorporates several high-speed design improvements based on the authors' expertise in RF and microwave electronics. These modifications improve pulse integrity and facilitate reliable nanosecond-scale laser modulation. The total fabrication cost is approximately \$100 for a batch of five assembled boards, making the solution substantially less expensive than a typical AOM system, which can cost several thousand dollars. Consequently, this design provides an attractive alternative for laboratories that either cannot justify the cost of an AOM or face long procurement delays. The circuit schematic is shown in Figure F6.

In order to benchmark the rise time, we measured the photocurrent from a pulsed laser, shown in Figure F7. The bias was adjusted down to 6 V from the initial design of 9 V, which significantly reduced the ringing, as seen in the figure. The rise time of the photocurrent (10-90%) was 25 ns. This was sufficient to observe Rabi oscillations in ensembles of NV centers as shown in Figure F5.

Appendix G: Green Excitation Options: 532 nm Coherent Laser vs 520 nm Thorlabs Diode

The excitation parts list (Table J2) offers two green-laser configurations suited to different cost, pulsing, and dual-color requirements. Option 1 is built around a fiber-coupled Coherent Sapphire FP laser (532 nm, 120 mW, CW). Option 2 re-

places this source with a single-mode fiber-coupled Thorlabs LP520-SF15A laser diode (520 nm, 15 mW), pulsed directly by the qlaser current driver described in Appendix F. Both options share the same galvo-scanned 4f relay and detection path described in Section II. The principal tradeoffs are summarized in Table J1.

The first option is the conventional choice for pulsed NV experiments: a high-power CW laser gated by one or two fiber-coupled AOMs in series provides high extinction, nanosecond switching. In practice, however, AOMs presented significant obstacles during platform development. A fiber-coupled 532 nm AOM (G&H 1200AFP-AD-1.0) carried an eight-month lead time and was found to be non-functional upon delivery due to a design defect identified only after an extended warranty investigation. A subsequent dual-AOM free-space configuration using available laboratory units proved unreliable, with only one of three units performing within specification. The resulting optical path required a dedicated auxiliary table and introduced alignment complexity incompatible with the dual-color requirement of the biological imaging channel, since fiber-coupled AOMs optimized for 532 nm cannot be repurposed for 450 nm excitation.

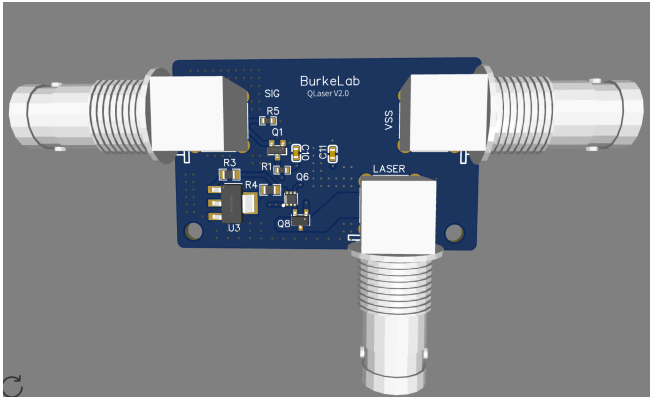
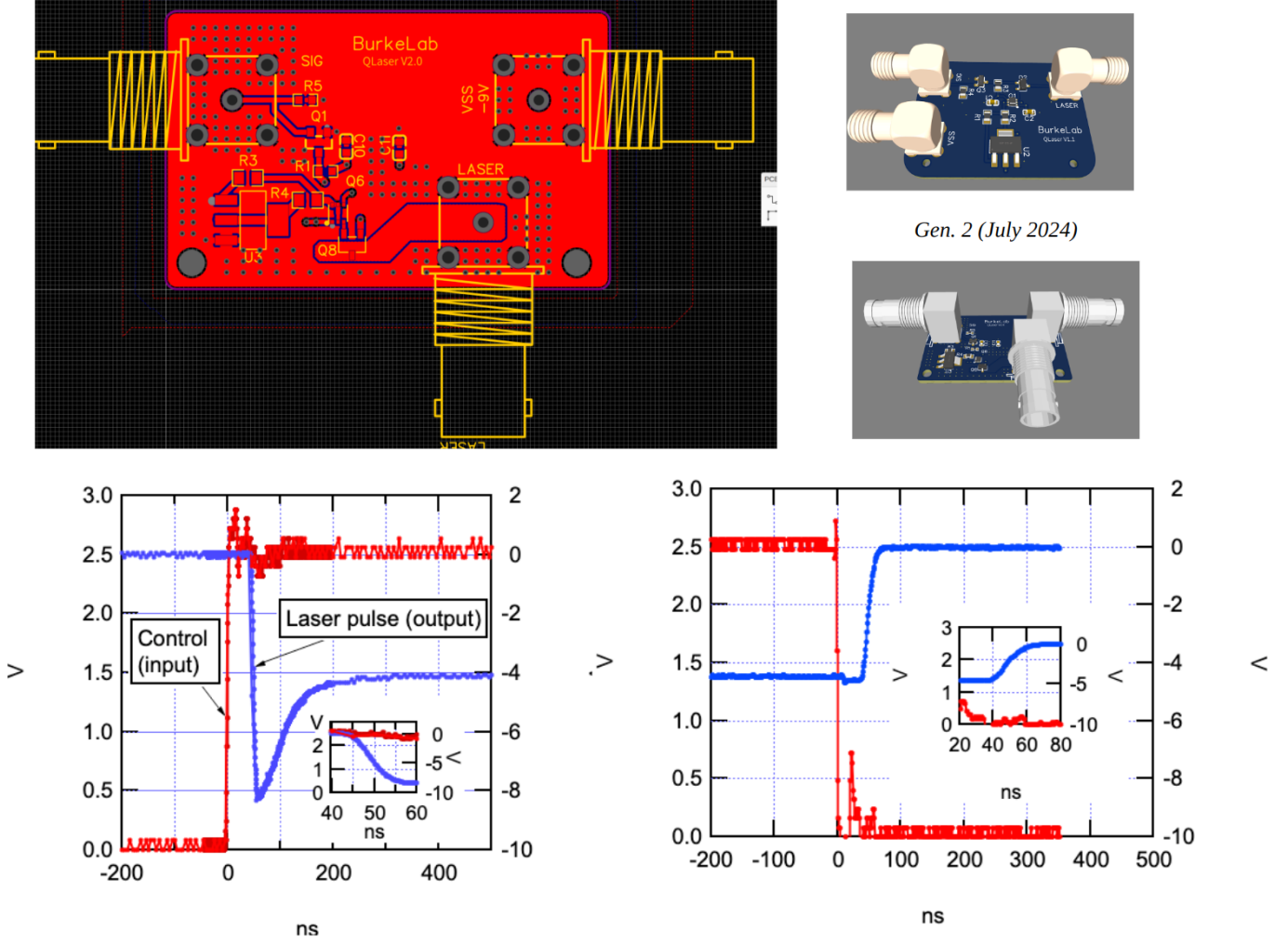
These constraints motivated the adoption of the second approach: direct current modulation of a fiber-pigtailed laser diode using the open-hardware qlaser driver (Appendix F). This approach achieves a 25 ns rise time (10-90%), which is sufficient for ODMR, Rabi, Ramsey, and T_1 protocols on, as demonstrated during previous experiments using ensemble NVs. The 520 nm diode is also compatible with the dual-color configuration: the same dichroic combiner that merges the 450 nm MagLOV channel accepts 520 nm without modification, enabling both excitation wavelengths to share the full 4f relay and galvo-scanning path.

Appendix H: Prior Art and Comparison

Our instrument sits at the intersection of two distinct literatures – open / teaching-lab NV microscopy and time-resolved fluorescence microscopy of biological radical pairs – and differs from every previously published platform in each. In this section, we summarize the relevant prior art and make those differences explicit.

1. Solid-state NV-center microscopes

Four previously reported platforms define the current landscape of NV-center instrumentation that is, at least in principle, reproducible outside a dedicated quantum-optics group. Zhang and co-workers³⁹ described a minimal, continuous-wave ODMR platform based on a Mini-Circuits voltage-controlled oscillator and a simple photodiode readout, with total build cost between ~\$1k (magnetometer variant) and ~\$3k (fluorescence-microscope variant); ensemble nanodiamond was used, and no pulsed experiments were performed. Sewani and co-workers³⁵ extended the teaching-lab concept



to ensemble pulsed NV quantum sensing, delivering the complete CW-ODMR $\rightarrow$ Rabi $\rightarrow$ Hahn-echo $\rightarrow$ CPMG cascade at a build cost of $\sim$ \$18.6k on $\langle 111 \rangle$ HPHT diamond, with a PCB loop-gap-resonator microwave antenna. Misonou and co-workers²⁹ presented a compact tabletop single-NV magnetometer explicitly designed to be reproducible “by nonspecialists”, reporting Rabi, Ramsey ($T_2^* = 0.50\mu\text{s}$), Hahn echo ($T_2 = 364\mu\text{s}$), and single-ensemble proton NMR ($T_2 = 16.1\mu\text{s}$, $d_{\text{NV}} = 6.26\text{nm}$) on a Keio/RIKEN-built instrument. Yuan, de Leon and co-workers²⁸ reported an instructional single-NV confocal microscope at a build cost of \$68k–\$100k, demonstrating $g^{(2)}(0) < 0.5$ single-NV verification, 38 ns Rabi, ^{15}N hyperfine Ramsey, and Hahn-echo T_2 at 23 G external magnetic field. Taken together, references^{28,29,35,39} form a clean cost/capability ladder from $\sim$ \$1k CW-only ODMR to $\sim$ \$100k single-NV coherent control.

Our platform differs from each of references^{28,29,35,39} in five concrete respects, none of which is merely a cost optimization.

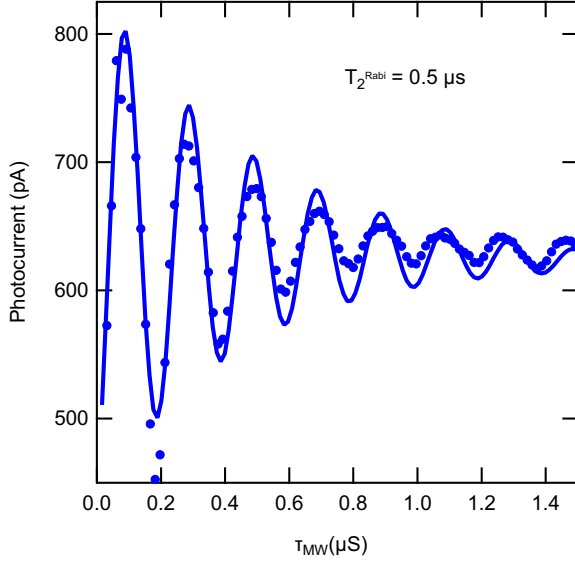


FIG. F5: Rabi oscillations on ensembles of NV centers using electrically pulsed laser source.

- **Commercial inverted microscope body with stationary-sample geometry.** References^{28,29,35,39} are all built on cage-optics frames; the sample is moved on a stage (Misonou) or the objective on a piezo with a fixed diamond (Yuan). We instead build on an Olympus IX71 inverted microscope body, which is the standard chassis of mammalian cell biology, and scan the excitation beam by a galvo-galvo pair through $4f$ relays while holding the sample stationary. This decouples the biological sample environment from the scanning mechanism and makes the instrument immediately compatible with live-cell culture dishes, microfluidic manifolds, patch-clamp electrodes, and objective-side optical tweezers.
- **Two complementary control-electronics paths, benchmarked on one optical bench.** References^{28,29} use a Tektronix AWG7102 + Anritsu MG3700A combination ($\gg$ \$20k) or an equivalent arbitrary-waveform / vector-signal-generator pair, and references³⁵ relies on a SpinCore PulseBlaster ESR Pro 250 plus an external lock-in. We evaluate two alternatives on the same optical train – an open-source RFSoc stack running QUICK-DAWG³² on a Real Digital RFSoc4x2 board ($\approx$ \$2k), and a dedicated hardware time tagger – and show that they occupy two distinct, complementary operating regimes of NV sensing. This comparative benchmarking is itself a contribution of the present work and is expanded in the subsection below.
- **Open-hardware pulsed laser driver (qlaser) in place of fiber-coupled acousto-optic modulators.** High-extinction optical gating for pulsed NV experiments is conventionally implemented with fiber-coupled AOMs

such as the Gooch & Housego Fiber-Q used in reference²⁹. These modules cost of order \$10k each, two are typically required in series to achieve a combined extinction ratio sufficient for single-NV T_1 measurements, and, at the time of writing, lead times of six months or more are routine due to persistent supply-chain constraints. We replace the AOM pair with qlaser⁴⁰, an open-hardware pulsed current source we have released under the GNU GPL v3 license at github.com/burkelabuci/qlaser. The board is a modified derivative of the pulsed laser driver described in Sewani et al.³⁵; five fully assembled boards can be ordered from a standard prototyping fabricator for $\sim$ \$100 total, i.e. of order \$10–\$20 per drive channel. Direct current-mode pulsing of the laser diode yields nanosecond rise and fall times sufficient for ODMR, Rabi, Ramsey and T_1 protocols without any moving optical element.

- **Demonstrated compatibility with live, respiring mitochondria.** None of the NV microscopes in references^{28,29,35,39} has been applied to a living biological sample. The combination of a stationary-sample inverted geometry, and a dual-excitation optical path at 520 nm (NV) and 450 nm (flavin / MagLOV / EYFP) makes the present instrument the first NV confocal platform whose optics, mechanics and electronics have been jointly validated for imaging vital, metabolically active mitochondria. The live-mitochondrial imaging demonstrated in Section III is, to our knowledge, the first of its kind on a single-NV-capable microscope.
- **A single optical path designed to connect solid-state and biological qubit protocols.** References^{28,29,35,39} are monochromatic NV instruments. Our $4f$ -relay architecture combines a 520 nm NV excitation path with a reconfigurable 450 nm blue-excitation path for flavin / MagLOV / EYFP-class biological fluorescence measurements, collects emission through the same confocal detection framework, and gates detection with a single-photon timing module. In the present work, the experimentally demonstrated spin measurements are NV ODMR and T_1 relaxation. The same optical, timing, microwave, and RF architecture is designed to support future Rabi, Ramsey, T_2 , and time-resolved RYDMR measurements, but those experiments are not claimed as completed results here.

2. Biological-qubit fluorescence microscopes

The instrumentation most directly comparable on the biological-qubit side is the fluorescence-microscopy platform of Ikeya and Woodward^{25,26}. Their 2021 PNAS instrument²⁵ is a custom, upright fluorescence microscope based on a $100\times/\text{NA } 1.49$ oil objective, a 450 nm CW diode laser at $\sim 0.5 \text{ kW cm}^{-2}$, an sCMOS camera, and a GMW 5204 vector electromagnet. It established that endogenous HeLa cell autofluorescence responds to static magnetic fields up to 25 mT with a triplet-born flavin SCRIP signature ($B_{1/2} = 18.0 \pm 0.5$

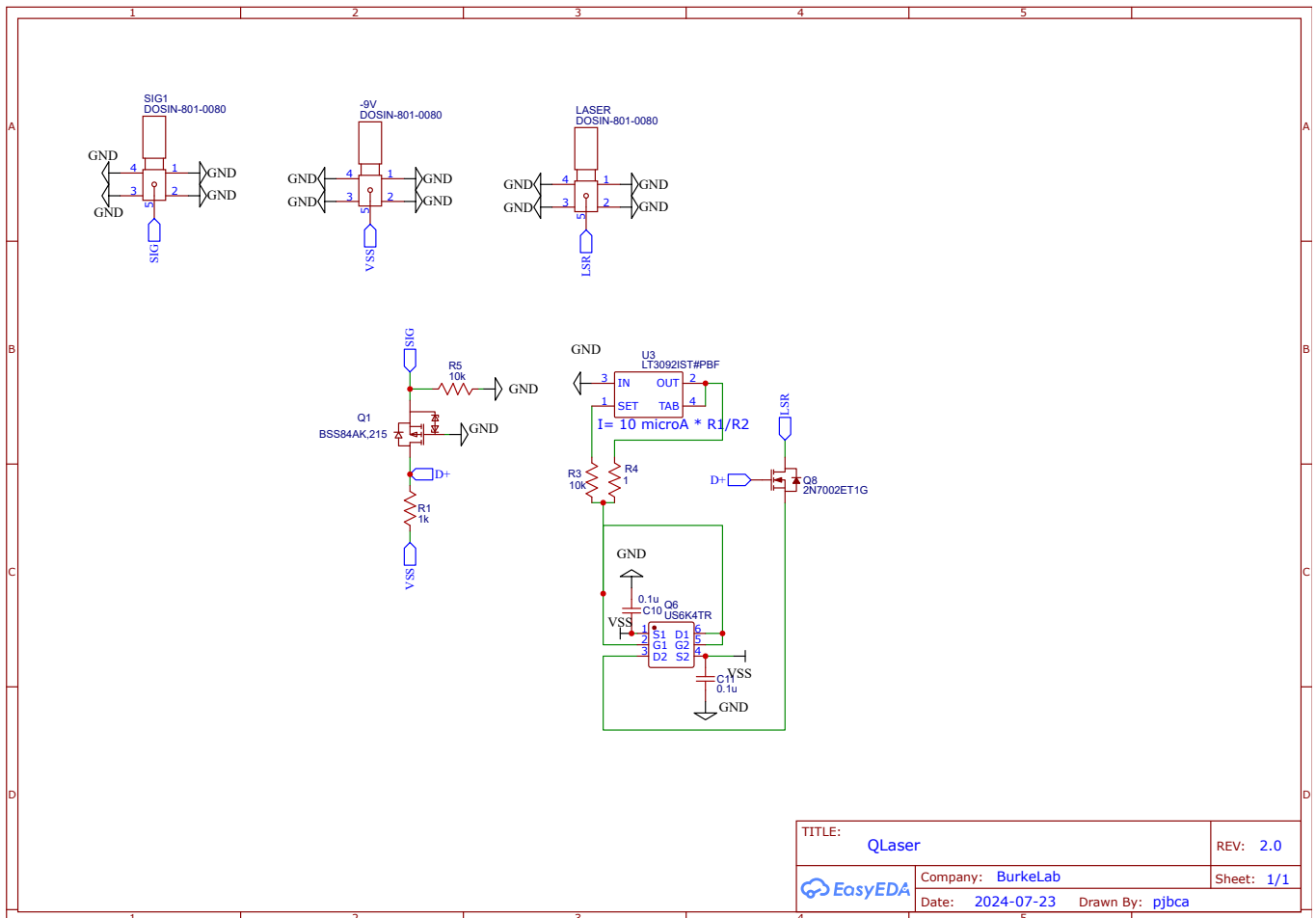


FIG. F6: Laser driver circuit schematic.

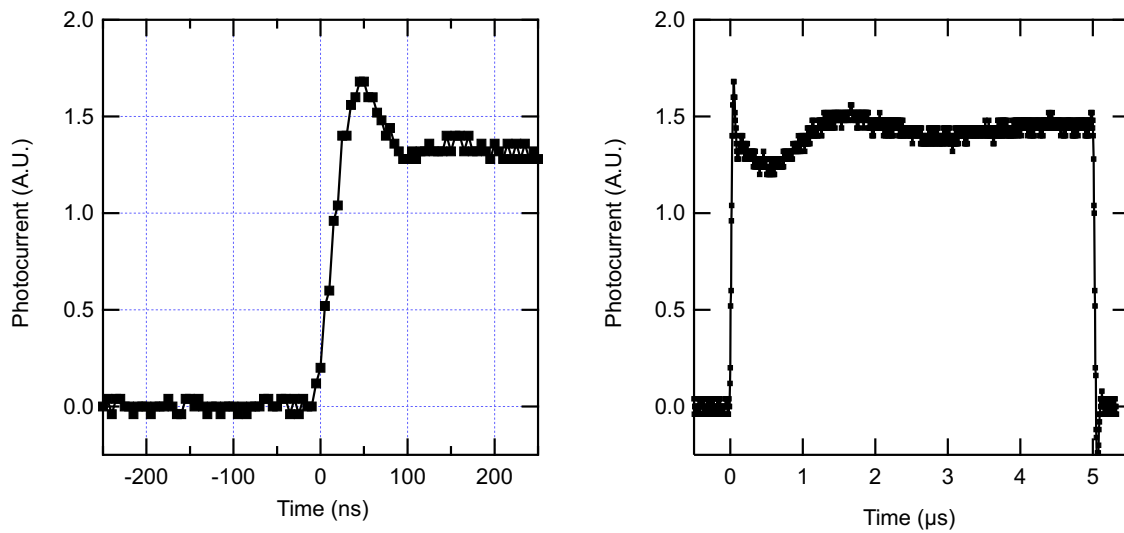


FIG. F7: Photocurrent vs. time, showing 25 ns rise time for electrically driven laser.

mT, saturation MFE 3.7%). Their 2026 JACS platform²⁶ adds two identical 450 nm nanosecond pulsed lasers combined on a multimode fiber, a rapidly switched magnetic field coil with < 30 ns rise/fall time, and LabVIEW / Raspberry-Pi-Pico orchestration, and introduces the pump–probe (PP) and pump–field–probe (PFP) techniques, resolving radical-pair lifetimes down to ~50 ns in subcellular (~1–4 pL) volumes.

Our instrument differs from references^{25,26} in four concrete respects.

- **Microwave and radio-frequency spin control on the biological-qubit channel.** References^{25,26} measure magnetic-field effects only: a static or a rapidly switched magnetic field alters the SCRP singlet–triplet branching, and the fluorescence change is recorded. They do not deliver a coherent microwave or RF drive to the biological qubit and therefore cannot perform RY-DMR, Rabi or Ramsey experiments on the SCRP itself. Our platform integrates a 0.5 GHz to 1.0 GHz RF drive channel, matched to the ~604 MHz S–T splitting of engineered MagLOV2 SCRPs³, together with the 2.87 GHz NV microwave chain. We can therefore perform true pulsed RYDMR and, in principle, coherent S–T control on biological qubits – something not possible on references^{25,26}.
- **Solid-state qubit capability on the same bench.** References^{25,26} have no NV channel: no green excitation, no 2.87 GHz microwave resonator, no pulsed-ODMR readout. The biological-qubit and solid-state-qubit communities have so far run on separate instruments. The present platform is, to our knowledge, the first reported microscope that natively supports both protocols at the same diffraction-limited spot, making it the first plausible experimental venue for hybrid NV–biological-qubit measurements.
- **Inverted, stationary-sample geometry for live-cell imaging.** References^{25,26} are upright and use a 100×/NA 1.49 oil objective pressed against a cover-glass from above. Live-cell culture chambers, incubation inserts and patch-clamp manipulators are designed for inverted microscopes; our Olympus IX71 chassis and stationary-sample galvo-galvo geometry therefore enables classes of experiment that are not accessible to the Woodward platform.
- **Super-resolution compatible optics and sub-diffraction quantum sensing.** The Woodward PFP instrument reports an ~8 μm detection spot and therefore averages over tens of mitochondria. Our galvo-galvo 4f-relay optics support single-point confocal addressing at the diffraction limit and, in combination with the super-resolution MagLOV ODMR approach we have recently demonstrated in live mammalian cells in CW mode⁵, provides a natural route to super-resolved, time-resolved RYDMR on mitochondrion-targeted biological qubits. The pulsed, RF-addressable variant of reference⁵ is one of the explicit motivations for the present instrument.

Appendix I: Coordinated experimental program with the Ikeya–Woodward platform

The most productive relationship between the Ikeya–Woodward platform and the present Olympus-based pulsed quantum microscope is not competitive but sequential. We propose, and our platform is designed to support, the following coordinated program.

- (i) Replicate the IW26 validation suite, including FMN, FAD, FMN/TrpH, and FAD/TrpH at pH 2.3, on our platform using PP and PFP pulse programs implemented on QUICK-DAWG³² and qlaser⁴⁰. A successful cross-validation would place the two instruments on a common methodological footing.
- (ii) Extend from incoherent magnetic-field gating to coherent RF drive at ~ 604 MHz on MagLOV2²³ and related engineered flavoproteins. This is the extension IW26 explicitly identifies as future work, and our RF chain is designed for this operating regime.
- (iii) Introduce NV magnetometry as an in-situ calibration method for the rapidly switched magnetic field. In this mode, the transient field profile measured by a surface-proximal NV center can be compared directly with the IW26 Hall-sensor calibration.
- (iv) Demonstrate pulsed RYDMR on mitochondrially targeted biological qubits, following the super-resolution continuous-wave approach of our earlier work⁵, while using the NV channel to benchmark the local magnetic-field environment in the same optical voxel.
- (v) Ultimately, prepare and measure an entangled state shared between an NV center in a surface-proximal nanodiamond and a biological qubit in the adjacent cell. This experiment is accessible, in principle, only on the class of platform reported here, but it is best understood as the endpoint of the IW26 → MagLOV → entangled-sensor trajectory that IW26 itself begins.

1. Summary of novelty

The novelty of the present work is therefore not a single new completed measurement, but the integration of microscope-level capabilities that have each existed separately in the literature and whose combination opens a new experimental regime.

- (i) We unify, in one inverted-microscope body, the green / 2.87 GHz NV channel of references^{28,29,35}, the optical geometry required for blue / nanosecond-pulsed biological-qubit excitation in references^{25,26}, and the RF-frequency range relevant to MagLOV-class biological qubits in references^{3,7,27}. In the present work, the experimentally demonstrated NV spin measurements are ODMR and T_1 relaxation. The same architecture is designed to support future T_2 , Ramsey, Hahn-echo, and

time-resolved RYDMR measurements after the corresponding pulse-sequence validation.

- (ii) We present and benchmark two complementary control-electronics paths on the same optical bench: an open-source RFSoc stack running QICK-DAWG³² on a Real Digital RFSoc4x2 board for multi-NV ensemble operation, and a dedicated hardware time tagger for single-NV operation.
- (iii) We adopt a stationary-sample galvo-galvo geometry on a standard Olympus IX71 body, eliminating sample-motion artifacts associated with sample-scanning implementations in references^{28,29,35} and making the instrument natively compatible with inverted biological-imaging formats.
- (iv) We demonstrate biological fluorescence compatibility on a single-NV-capable optical platform by imaging TMRE-stained and MagLOV-expressing HeLa cells.

This result does not yet constitute biological-qubit RYDMR, but it establishes that the same microscope framework used for NV confocal imaging, ODMR, and T_1 measurement can be reconfigured for genetically encoded biological fluorescence samples through limited optical changes.

- (v) The architecture is furthermore a natural optical front end for the noise-suppressed NV-pair gradiometer protocol we have recently proposed for biological operation²³. The present data establish the required confocal imaging, ODMR, T_1 readout, stationary-sample scanning, and biological fluorescence compatibility, while two-qubit entanglement-enhanced sensing remains a future experimental target inspired by references^{15–17}.

Appendix J: Parts List

Here we organized parts list with different options. You can build your setup according to your budget

TABLE J1: Comparison of green excitation options.

Property	Option 1 (Coherent Sapphire)	Option 2 (Thorlabs LP520)
Wavelength	532 nm	520 nm
Output power	120 mW	15 mW
Fiber coupling	Single-mode, factory	Single-mode, factory
Pulsing method	External AOM required	Direct current modulation (qlaser)
Rise/fall time	< 10 ns (AOM-limited)	~25 ns (qlaser-limited)
Dual-color compatibility	No (AOM is 532 nm-specific)	Yes (dichroic combination at input)
Approximate cost	~\$5k (laser) + ~\$10k (AOM)	~\$800 (laser) + ~\$20 (driver board)

TABLE J2: List of parts for the 4f system of the laser excitation setup.

Item	Vendor	Model	Description	Qty
Laser Option 1:				
Green Laser	Coherent	Sapphire FP	532 nm, 120 mW, fiber-coupled	1
Laser Option 2:				
Green Laser Diode	Thorlabs	LP520-SF15A	520 nm, 15 mW, SM Fiber-Pigtailed Laser Diode, FC/PC	1
Blue Laser Diode	Thorlabs	LPF450-SF25	450 nm, 25 mW, SM Fiber-Pigtailed Laser Diode, FC/PC	1
AOM 1:				
AOM	ISOMET	1205C-2-790	2 mm aperture, 3.63mm/ μ s acoustic velocity	1
AOM Driver	ISOMET	222R-2-81	50 Ω , 80 MHz, 1.6 W	1
AOM 2:				
AOM+Driver	Brimrose	TEM-200-25-20-532-2FP	Fiber coupled, 532 nm, 200 MHz, 1W	1
Other Optics:				
Fiber Adaptor Plate	Thorlabs	SM1FCA2	FC/APC Fiber Adaptor	3
Mounted Aspheric Lens	Thorlabs	C280TMD-A	Collimate light exiting fiber/diode	3
Lens Cell Adaptor	Thorlabs	S1TM09	Aspheric Lens holder	3
Kinematic Mount	Thorlabs	KC1-T	For aspheric lens/fiber adapter plate	3
Lens Tube	Thorlabs	SM1L10	For aspheric lens focus/lens hood	3
30 mm Cage Plate	Thorlabs	CP45	Removable cage plate	3
SM1 Cage Plate	Thorlabs	CP33	For cage rods and mounting lenses	3
Cage Assembly Rod	Thorlabs	R4-P4	4" Long, $\varnothing$ 6 mm, 4 Pack	3
Z-axis Translation Mount	Thorlabs	SM1ZA	Fine tuning distance between lenses	3
Plano-Convex Lens	Thorlabs	LA1251-AB	f=100 mm	5
Plano-Concave Lens	Thorlabs	LC4513-AB	f=-7.5 cm	3
Bandpass Filter	Thorlabs	FLH532-4	532 nm bandpass filter	1
Beamsplitter	Thorlabs	BSN10	$\varnothing$ 1", 10:90, 400-700 nm	1
Kinematic Mount	Thorlabs	KM100	For reflection mirror	6
Centering Plate	Thorlabs	KCP1	For kinetic mount	6
Centering Plate	Thorlabs	KCP1	For kinetic mount	6
Lens Mount	Thorlabs	LMR1	Fixed lens mount	4
Iris	Thorlabs	CP20D	For alignment	1
Iris	Thorlabs	ID25	For alignment	3
Optical Post	Thorlabs	TR1	1.0" post	4
Optical Post	Thorlabs	TR1.5	1.5" post	22
Post Holder	Thorlabs	PH1.0	1" holder	4
Post Holder	Thorlabs	PH1.5	1.5" holder	22
Clamping Fork	Thorlabs	CF125	For beamsplitter	8
Pedestal Base Adapter	Thorlabs	BE1	For beamsplitter	8
Reflection Mirror	Thorlabs	PF10-03-P01	$\varnothing$ 1" Protected Silver Mirror	5
Spacer	Thorlabs	RSxM	Height adjustment	as needed

TABLE J3: List of parts for the 4f system of the fluorescence collection setup.

Item	Vendor	Model	Description	Qty
Achromatic Doublet Lens	Thorlabs	AC254-150-AB	f=150 mm	1
Plano-Convex Lens	Thorlabs	LA1251-AB	f=100 mm, N-BK7	2
Z-axis Translation Mount	Thorlabs	SM1ZA	Fine tuning between lenses	1
Pinhole	Thorlabs	P50K1	50 μ m, SM1-threaded, mounted	1
Longpass Filter	Thorlabs	FELH0600	600 nm cut-on	1
ND Filter Wheel	Thorlabs	FW1AND	5 ND filters included	1
Fiber Adaptor Plate	Thorlabs	SM1FC2	FC/PC Fiber Adaptor, For SPD	1
Mounted Aspheric Lens	Thorlabs	C280TMD-A	Collimate light exiting fiber/diode	1
Lens Cell Adaptor	Thorlabs	S1TM09	Aspheric Lens holder	1
Kinematic Mount	Thorlabs	KC1-T	For aspheric lens/fiber adapter plate	1
Lens Tube	Thorlabs	SM1L10	For aspheric lens focus/lens hood	1
Fiber Patch Cable	Thorlabs	M42L01	FC/PC multimode fiber	1
SPD	Excelitas	SPCM-AQRH-10-FC	1500 cps dark, 22 ns dead time	1
Kinematic Mount	Thorlabs	KM100	For mirrors	3
Reflection Mirror	Thorlabs	PF10-03-P01	Ø1" Protected Silver Mirror	3
Centering Plate	Thorlabs	KCP1	For kinetic mount	3
Optical Post	Thorlabs	TR1.5	1.5" post	13
Post Holder	Thorlabs	PH1.5	1.5" holder	13
Cage Cube	Thorlabs	CM1-DCH	For beamsplitter mounting	1
SM1 Cage Plate	Thorlabs	CP33	For cage rods	3
Cage Assembly Rod	Thorlabs	R4-P4	4" Long, Ø6 mm, 4 Pack	1
Spacer	Thorlabs	RSxM	Height adjustment	as needed

TABLE J4: List of parts for the spectroscopy setup.

Item	Vendor	Model	Description	Qty
Beam sampler	Thorlabs	BSF10-A	1" beam sampler	1
Kinematic Mount	Thorlabs	KM100C	For beamsplitter plate	1
Diffraction Grating	Thorlabs	GT50-03	Visible transmission grating	1
Plano-Convex Lens	Thorlabs	LA1251-AB	f=100 mm	2
Monochrome CCD	Player One	Poseidon-M Pro (IMX571)	Cooled camera for spectra	1
Kinematic Mount	Thorlabs	KM100	For mirrors	3
Reflection Mirror	Thorlabs	PF10-03-P01	Ø1" Protected Silver Mirror	3
Centering Plate	Thorlabs	KCP1	For kinetic mount	3
Optical Post	Thorlabs	TR1.5	1.5" post	6
Post Holder	Thorlabs	PH1.5	1.5" holder	6
Spacer	Thorlabs	RSxM	Height adjustment	as needed

TABLE J5: List of parts for the 4f system of the nanoscale positioning in the experiment.

Item	Vendor	Model	Description	Qty
Galvo-Galvo Scanner	Thorlabs	LSKGG4	XY scan, $\pm 15^\circ$, 20 μ rad resolution	1
Kinematic Mount	Thorlabs	KM100	For mirrors	2
Reflection Mirror	Thorlabs	PF10-03-P01	Ø1" Protected Silver Mirror	2
Centering Plate	Thorlabs	KCP1	For kinetic mount	2
SM1 Cage Plate	Thorlabs	CP33	For cage rods and mounting lenses	3
Cage Assembly Rod	Thorlabs	R4-P4	4" Long, Ø6 mm, 4 Pack	1
Plano-Convex Lens	Thorlabs	LA1251-AB	f=100 mm	1
Iris	Thorlabs	CP20D	For alignment	2
Objective Piezo	Thorlabs	PFM450E	Z scan, 3 nm resolution, 450 μ m range	1
Microscope Objective	Olympus	UPLSAPO40X2	40x/0.95, WD 0.18 mm	1

TABLE J6: List of parts for fluorescence imaging and laser monitoring.

Item	Vendor	Model	Description	Qty
Microscope objective	Zeiss	420792-9800-000	Alpha Plan-Apo 100x/1.46Oil DIC M27	1
Notch Filter	Thorlabs	NF514-17	514 nm, FWHM = 17 nm	2
CMOS Camera	Thorlabs	CS165CU	Sample Imaging	1
Power Meter	Thorlabs	PM100D	100 pW - 200 W range	1
Power Sensor	Thorlabs	S120C	400-1100 nm, 50 nW - 50 mW	1
Reflection Mirror	Thorlabs	PF10-03-P01	Ø1" Protected Silver Mirror	as needed
Optical Post	Thorlabs	TR1.5	1.5" optical post	as needed
Post Holder	Thorlabs	PH1.5	1.5" holder	as needed
Kinematic Mount	Thorlabs	KM100	For mirror/collimator	as needed

TABLE J7: List of electronics used in the experiment.

Item	Vendor	Model	Description	Qty
Pulse Streamer	Swabian	PS 8.2	8 digital/2 analog outputs	1
Time Tagger	Swabian	Time Tagger 20	8 inputs, 34 ps jitter	1
DAQ	NI	USB-6453	32 AI channels, 20-bit	1
Microwave Amplifier	Mini-Circuits	ZHL-16W-43-S+	1.8-4 GHz, 16 W	1
IQ Modulator	TI	TRF37t05EVM	Microwave drive/modulation	1
Circulator	Mini-Circuits	PE83CR005	2-4 GHz	1
Directional Coupler	Mini-Circuits	ZDFC-20-33-S+	20-3000 MHz, 20.5 dB	1
Step Attenuator	—	—	1 dB - 10 dB steps	1
Spectrum Analyzer	Rigol	DSA832E	9 kHz-3.2 GHz	1
High Power Terminator	Fairview	ST6S-10	SMA terminator, used on circulator port	1
Oscilloscope	Rigol	DS1054	4 ch, 50 MHz, 1 GSa/s	1
MW Cables	Thorlabs	SMMxx	50 Ω , DC-26 GHz	several
SMA 50 Ω Terminator	—	—	For port termination/impedance matching	several
SMA-F / SMA-M Adapters	—	—	Gender adapters for SMA connectors	several