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There's Plenty of Room at the Bottom: Opportunites and Challenges for Microrobotics

Arianna Menciassi, PhD Scuola Superiore Sant'Anna, Pisa (Italy)

Roma, February 26th 2024



Robots are more and more outside the cages...



Dustbot, robot for urban hygiene (SSSA)

MATE, wearable robot for motion augmentation (IUVO-COMAU) they can be wearable ...

and ...

... robots can work inside the body for chronic and acute interventions











ROBOTS NAVIGATING THE BODY



ROBOTS RESIDING IN THE BODY









Health and Robotics ...

No longer science fiction, robotics has emerged as a leading alternative for many healthcare applications



Dr. Daniel Kraft - "What's next in healthcare?"

Daniel Kraft is a physician-scientist, inventor and innovator. He is chair of the Medicine track for Singularity University and Executive Director for FutureMed, a program which explores convergent, exponentially developing technologies and their potential in biomedicine and healthcare

Worrell Infographic, vol. 1, no. 3, figure 01, 2015







ROBOTS ENTERING THE BODY WITH THEIR



Robotics and Minimally Invasive surgery





Catherine Mohr - Intuitive Surgical

Mazor's New Renaissance Robotic Spinal Surgery System

Robotic technologies to make surgery more accurate and less operator depending, to reach unreachable areas of the body without scars...





The beginning: **Industrial Robotics meets Clinical Imaging**

Robotic Assisted Surgery – RAS

Computer Integrated Surgery - CIS

IEEE TRANSACTIONS ON BIOMEDICAL ENGINEERING, VOL. 35, NO. 2, FEBRUARY

Large Scanning Apertury

Flexibility

A Robot with Improved Absolute Positioning Accuracy for CT Guided Stereotactic Brain Surgery

YIK SAN KWOH, MEMBER, IEEE, JOAHIN HOU, EDMOND A. JONCKHEERE, SENIOR MEMBER, IEEE, AND SAMAD HAYATI



Image

Acquisition

Planning in Virtual Environment



Computer-assisted surgery

Abstract-In this paper, we describe how a Unimation Puma 200 robot, properly interfaced with a CT scanner and with a probe guide mounted at its end effector, can be used for CT-guided brain tumor biopsies. Once the target is identified on the CT picture, a simple command allows the robot to move to a position such that the end effector probe guide points towards the target. This results in a procedure faster than one with a manually adjustable frame. But probably the most important advantage, as we show in this paper, is the improved accuracy that can be reached by proper calibration of the robot.

Y. S. Kwoh, CT Research, Department of Radiology, Memorial Medical Center, Long Beach, CA, USA

Medical Images make possible the typical CAD-CAM process ... in surgery





CAM (Computer Aided Manufacturing) toolchain





The framework today









INTUITIVE



Two exercises in the last two years: The first one...

SCIENCE ROBOTICS | REVIEW

MEDICAL ROBOTS

A decade retrospective of medical robotics research from 2010 to 2020

Pierre E. Dupont¹*, Bradley J. Nelson², Michael Goldfarb³, Blake Hannaford⁴, Arianna Menciassi⁵, Marcia K. O'Malley⁶, Nabil Simaan³, Pietro Valdastri⁷, Guang-Zhong Yang⁸

Robotics is a forward-looking discipline. Attention is focused on identifying the next grand challenges. In an applied field such as medical robotics, however, it is important to plan the future based on a clear understanding of what the research community has recently accomplished and where this work stands with respect to clinical needs and commercialization. This Review article identifies and analyzes the eight key research themes in medical robotics over the past decade. These thematic areas were identified using search criteria that identified the most highly cited papers of the decade. Our goal for this Review article is to provide an accessible way for readers to quickly appreciate some of the most exciting accomplishments in medical robotics over the past decade; for this reason, we have focused only on a small number of seminal papers in each thematic area. We hope that this article serves to foster an entrepreneurial spirit in researchers to reduce the widening gap between research and translation.



Two exercises in the last two years: The second one...

Proceedings of EEE

SPECIAL ISSUE

Surgical Robotics and Computer-Integrated Interventional Medicine



STATISTICAL PROCESS IMPROVEMENT



SPECIAL ISSUE

SURGICAL ROBOTICS AND COMPUTER-INTEGRATED INTERVENTIONAL MEDICINE

Edited by R. H. Taylor, N. Simaan, A. Menciassi, and G.-Z. Yang

835 Robot-Assisted Minimally Invasive Surgery—Surgical Robotics in the Data Age

By T. Haidegger, S. Speidel, D. Stoyanov, and R. M. Satava

|INVITED PAPER| This article summarizes the state of the art in robot-assisted minimally invasive surgery and provides an overview of key emerging technologies associated with next-generation systems.

847 Continuum Robots for Medical Interventions

By P. E. Dupont, N. Simaan, H. Choset, and C. Rucker

|INVITED PAPER| This article provides a unified summary of the state of the art of continuum robot architectures with respect to design for specific clinical applications.

871 Soft Robot-Assisted Minimally Invasive Surgery and Interventions: Advances and Outlook

By K.-W. Kwok, H. Wurdemann, A. Arezzo, A. Menciassi, and K. Althoefer

|INVITED PAPER| This article provides an in-depth overview of recent progress in soft robotics for surgery and outlines remaining challenges in the development of soft robotics technologies for in-body operation, such as materials selection, tunable stiffness, soft design paradigms, and control issues.

893 Robotic Assistance for Intraocular Microsurgery: Challenges and Perspectives

By I. i. Iordachita, M. D. de Smet, G. Naus, M. Mitsuishi, and C. N. Riviere

INVITED PAPER | This article analyzes the advances in retinal robotic microsurgery, its current drawbacks and limitations, as well as the possible new directions to expand retinal microsurgery to techniques currently beyond human boundaries or infeasible without robotics.

909 Advancement of Flexible Robot Technologies for Endoluminal Surgeries

By J. Kim, M. de Mathelin, K. Ikuta, and D.-S. Kwon

|INVITED PAPER| This article covers the key technical issues in flexible surgical robotics, such as manipulator design, modeling, and control, and it introduces emerging flexible technologies organized according to their target application in the endoluminal surgical field.

932 Image-Guided Interventional Robotics: Lost in Translation? By G. Fichtinger, J. Troccaz, and T. Haidegger

|INVITED PAPER| This article provides an overview of the state of the art in image-guide surgical systems, together with a discussion of key issues for system developers in the translation of scientific research to clinical application.

951 Robot-Assisted Medical Imaging: A Review

By S. E. Salcudean, H. Moradi, D. G. Black, and N. Navab

|INVITED PAPER| This article provides an overview of the current state of the art and potential research directions for robotic imaging systems, with special emphasis on instances in which the accurate placement and trajectory control of the imaging system using a robot are of paramount importance.

968 State of the Art and Future Opportunities in MRI-Guided Robot-Assisted Surgery and Interventions

By H. Su, K.-W. Kwok, K. Cleary, I. Iordachita, M. C. Cavusoglu, J. P. Desai, and G. S. Fischer

|INVITED PAPER| This article describes challenges and history of robotic systems operating in an MRI environment, and outlines promising clinical applications and associated state-of-the-art MRI-compatible robotic systems and technology.

993 Concepts and Trends in Autonomy for Robot-Assisted Surgery By P. Fiorini, K. Y. Goldberg, Y. Liu, and R. H. Taylor

|INVITED PAPER| This article provides a unified summary of the state of the art of the continuum robot architectures with respect to design for specific clinical applications and illustrates these themes with examples from current research.

1012 Haptic Feedback and Force-Based Teleoperation in Surgical Robotics

By R. V. Patel, S. F. Atashzar, and M. Tavakoli

|INVITED PAPER| This article examines key challenges associated with the application of haptic feedback and force-based teleoperation for surgical robots, such as instrumentation, fidelity, stability, and force-reflection modalities.

1028 Magnetically Actuated Medical Robots: An in vivo Perspective

By B. J. Nelson, S. Gervasoni, P. W. Y. Chiu, L. Zhang, and A. Zemmar

INVITED PAPER | This article describes magnetically guided medical robots, both tethered and untethered, working at different scales and it analyses the *in vivo* translation with increased control and safety.



Main challenges and main needs

- Being targeted, i.e. helping the surgeon to reach «unreachable» areas – targeted therapy
- Bringing dexterity inside the body with minimal access and high performance actuators
- Being safe in interaction
- Moving towards scarless operations





The problem: reducing the invasiveness, augmenting the dexterity at the distal part





The problem: reducing the invasiveness, augmenting the dexterity in the distal part



The problem: reducing the invasiveness, augmenting the dexterity in the distal part



University of Nebraska, USA





Tension Propagation Analysis of Novel Robotized Surgical Platform for Transumbilical Single-Port Access Surgery

> Jongwon Lee, Yongjae Kim, Se-gon Roh, Jiyoung Kim, Younbaek Lee, Jeonghun Kim, Byungjune Choi, and Kyoungsik Roh

Samsung Advanced Institute of Technology, Samsung Electronics Co., South Korea







Wrist

Single port modular surgery: how to deploy many degrees of freedom through a small hole





Multiarticulated platform for Minimally Invasive Aortic Valve Replacement

ValveTech: a Novel Robotic Approach for Minimally Invasive Aortic Valve Replacement

Izadyar Tamadon¹ , Virginia Mamone², Yu Huan¹, Sara Condino², Claudio Quaglia¹, Vincenzo Ferrari², Mauro Ferrari², Arianna Menciassi¹

Demo Movie



¹The BioRobotics Institute, Scuola Superiore Sant'Anna, Pontedera, Pisa, Italy. ²EndoCAS Center for Computer-Assisted Surgery, Università di Pisa, Italy.



and second, the frame



The easiest navigation environment without incisions - Miniature robots navigating in the GI tract...



ROBOTS NAVIGATING THE BODY



Smart flexible endoscopes





An endoscope with biomimetic locomotion (2000-2010)



ww.endotics.co

Endatics



Endoscopes and surgical tools without tails... the trend to capsule-like robots



Cathete

Flexibility of traditional wired devices limits access to some target areas (i.e. limitation to targeted therapy)

ht atrium of hear



Small diameter and remote districts can be reached **only by wireless or softly tethered devices**







The idea of bringing therapeutic and advanced diagnostic solutions where they are needed: the endoscopic capsule

Available wireless capsules: visual investigation of normally not explored areas



G. Iddan and P. Swain. History and development of capsule endoscopy. *Gastrointestinal Endoscopy.* 14: 1-9 (2004)



Active/teleoperated locomotion for giving "legs" to advanced diagnostic and therapeutic solutions





ACTIVE capsules with on-board PROPULSION











The EU VECTOR Project Korean IMC Project







Bottleneck for active on-board propulsion





A legged capsule incorporating stateof-art batteries could only walk for less than 30 minutes along the GI tract



Which solutions for a real scarless intervention and limiting actuation/powering problems at distal level?



Which solutions for a real scarless intervention and limiting actuation/powering problems at distal level?

Torque



William Gilbert, 1600

De Magnete, Magneticisque Corporibus, et de Magno Magnete Tellure (On the Magnet and Magnetic Bodies, and on That Great Magnet the Earth)

Magnetic endoscopic capsules, magnetic retraction systems, magnetic catheters, magnetic particles for drug delivery and drug targeting... $\vec{\tau} = V\vec{M} \times \vec{B}$ Force





The easiest navigation environment - Miniature robots navigating in the GI tract...









...wireless magnetic dragging is not for free: localization issues open!





M. Salerno et al., «A discrete-time localization method for capsule endoscopy based on on-board magnetic sensing», Measurement Science and Technology 23 (1), 2011

and also Jake Abbott, Pietro Valdastri, etc...







ROBOTS NAVIGATING THE BODY





JOURNAL OF MICROELECTROMECHANICAL SYSTEMS VOL. 1, NO. 1, MARCH 1992

There's Plenty of Room at the Bottom

Richard P. Feynman

I imagine experimental physicists must often look with envy at men like Kamerlingh Onnes, who discovered a field like low temperature, which seems to be bottomless and in which one can go down and down. Such a man is then a leader and has some temporary monopoly in a scientific adventure. Percy Bridgman, in designing a way to obtain-higher pressures, opened up another new field and was able to move into it and to lead us all along. The development of ever higher vacuum was a continuing development of the same kind.

I would like to describe a field, in which little has been done, but in which an enormous amount can be done in principle. This field is not quite the same as the others in that it will not tell us much of fundamental physics (in the dots on the fine half-tone reproductions in the Encyclopaedia. This, when you demagnify it by 25 000 times, is still 80 angstroms in diameter—32 atoms across, in an ordinary metal. In other words, one of those dots still would contain in its area 1000 atoms. So, each dot can easily be adjusted in size as required by the photoengraving, and there is no question that there is enough room on the head of a pin to put all of the Encyclopaedia Britannica.

Furthermore, it can be read if it is so written. Let's imagine that it is written in raised letters of metal; that is, where the black is in the Encyclopaedia, we have raised letters of metal that are actually 1/25~000 of their ordinary size. How would we read it?

If we had compathing written in such a way we could



Main Chalengesten... - For catheters: flexibility, maneuverabisityaanstertip contro - For microrobots: control,

biocompatibility, tracking (+ many others!) NAVION, ETH



Microparticles in the blood flow (Fantastic Voya\ge)



3D Printing of Small-Scale Soft Robots with Programmable Magnetization



HMD Ansari et al., Adavanced Functional Materials, 2023

3D Printing of Small-Scale Soft Robots with Programmable Magnetization



3D Printing of Small-Scale Soft Robots with Programmable Magnetization

Mohammad Hasan Dad Ansari, Veronica Iacovacci, Stefano Pane, Mouloud Ourak, Gianni Borghesan, Izadyar Tamadon, Emmanuel Vander Poorten, Arianna Menciassi



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HMD Ansari et al., Adavanced Functional Materials, 2023



Magnetic catheters with programmable magnetization

- Distributed magnetic particles miniaturization
- Magnetic anisotropy programmed bending patient-specific
- 3D printing different materials, sizes, shapes
 versatile





Ansari et. al. Actuators, 2023

Robot-assisted guidance of the magnetic catheter



Robot assisted catheterization

- Automatic insertion using catheter driver
- EPM path defined using several waypoints
- EPM moved along the path based on visual feedback of the operator
- Success rate = 83.3% (5/6)





Main Challenges:

- For catheters: flexibility, maneuverability and tip control.
- For microrobots: control, biocompatibility, tracking (+ many others!)

- Space constraints
- Need for higher spatial resolution
- Need for higher temporal resolution
- Need to shift from microscope-based lab settings to tissue-compliant imaging modalities
- Lower contrast mismatch between microrobots and tissue



Microparticles in the boold flow (Fantastic Voya\ge)



Let's focus on challenges for microrobots in the vasculature



(Magnetic) Microrobots for in vivo applications – Open Challenges



A helical micromotor helps an immotile but healthy bovine sperm cell get to an egg in culture.

Medical microbots need better imaging and control

Mariana Medina-Sánchez and Oliver G. Schmidt set priorities for more realistic tests of tiny machines that could be used to diagnose and treat conditions. So far, most microbot experiments have been done *in vitro* under conditions very different from those in the human body. Many devices rely on toxic fuels, such as hydrogen peroxide. They are simple to steer in a Petri dish, but harder to control in biological fluids full of proteins and cells, and through the body's complex channels and cavities.

To enter clinical trials, microbots must clear two major hurdles. First, researchers need to be able to see and control them operating inside the body — current imaging techniques have insufficient resolution and sensitivity. <u>Second, the</u> vehicles need to be biocompatible and be removed or stabilized after use. Achieving both aims would set the stage for further improvements — in steering and mobility, materials and capabilities.

We call on microrobotics researchers, materials scientists and bioimaging and medical specialists to work together to solve these problems. And regulatory agencies need to put in place directives for testing therapeutics that are based on microbots.



Micro & Nanorobotics towards in vivo applications: challenges



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Micro & Nanorobotics towards in vivo applications: challenges



Micro & Nanorobotics towards in vivo applications: challenges



5729.

(Magnetic) Microrobots for in vivo applications – Open Challenges

Development of fully degradable structures



Loaded drugs and magnetic particles are biodistributed in the body after degradation

X. Wang et al., 2018.H. Ceylan et al., 2019.

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HW Huang et al., *Nat. Comm.*, 2016. Arianna Menciassi, ROBOVIS 2024



Intravascular magnetic catheter to retrieve micro and nanoagents from the bloodstream lacovacci, V., et al. Adv. Sci. 2018

lacovacci, V., et al. ICRA 2019

MODULAR CATHETER STRUCTURE

TARGET: ORGANS FEATURED BY TERMINAL CIRCULATION (e.g. LIVER, KIDNEY, PANCREAS)

TIP o face these issues, we developed BALLOON in intravascular magnetic microcatheter able to capture unaccumulated anoparticles, thus preventing side effects MAGNETIC MODULE DEPLETION MODULE SELDINGER **GUIDEWIRE CHANNEL** VENOUS OUTLET PERMANENT MAGNETS SPACERS PARTICLE **CONNECTION** FLOW **CANALISED IN** ELEMENT **INFLATED BALLOON** THE CATHETER nciassi, ROBOVIS 2024

Magnetic module design – FEM modeling

- SPHERICAL PARTICLE
- POINT DIPOLE APPROXIMATION
- LAMINAR FLOW IN A CHANNEL $v_p = v + \zeta f(H)(H \cdot \nabla)H$

$$\zeta = \frac{\mu_0(1+\chi_f)}{6\pi\eta_f} \frac{V_{mag}}{r_h}$$

$$f(H) = \begin{cases} \frac{3(\chi_p - \chi_f)}{(\chi_p - \chi_f) + 3(1 + \chi_f)} \\ \frac{M_{sp}}{H} \end{cases}$$

 $\frac{MODELING}{\frac{M_{sp}}{H}} > \frac{3(\chi_p - \chi_f)}{(\chi_p - \chi_f) + 3(1 + \chi_f)}$ $\frac{M_{sp}}{H} \le \frac{3(\chi_p - \chi_f)}{(\chi_p - \chi_f) + 3(1 + \chi_f)}$

PARTICLE IN A FLUIDIC AND MAGNETIC FIELD SIMPLIFIED



- CATHETER DIAMETER (12 F, 15 F)
- MAGNET NUMBER, GROUPING
- PARTICLES DIMENSION





Iacovacci, V., et al. Adv. Sci. 2018 Iacovacci, V., et al. ICRA 2019

Experimental validation



Iacovacci, V., et al. Adv. Sci. 2018 Iacovacci, V., et al. ICRA 2019



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Microrobots imaging – state of the art



Microrobots US imaging – examples



B-MODE IMAGING OF SUB-MILLIMETRIC ROBOTS



Li et a 2019 IN HIGHLY ECHOGENIC MEDIA



Sanchez et al., IEEE BioRob, 2014



Yu et al., Nat. Comm, 2019



Microrobots US imaging – examples



Motion-based analysis can help OPPORTUNITIES OFFERED BY DOPPLER IMAGING



Microrobots US imaging – examples



Selective motion filtering can improve tracking stability and precision ACOUSTIC PHASE ANALYSIS (APA)



Microrobots US imaging – APA



Different microrobot locomotions induce different phase feedback







Experimental setup for real-time imaging and tracking





Vibrations detection algorithm



1. Fourier analysis Analyze the frequency components of the acoustic phase

2. Frequency filtering Isolate vibrations at microrobot frequency

3. Phase filtering Isolate vibrations in-phase with the magnetic field

4. Overlap with B-mode



Tracking soft vibrating microrobots in tissues





Rotations detection algorithm



1. Temporal analysis Analyze the differential phase in the time domain

2. Block-matching

Cross-correlate the mean differential phase with a rotation template

3. Derive microrobot features

Localize the maximum in the cross-correlation map and identify size of motion diagram

4. Overlap with B-mode



Tracking rotating microrobots in vascular phantom





Use APA feedback for control

Microrobot states

2 time [s]

US waves



Pane, S., et al., IEEE- Transactions on Robotics, (2022).





Microrobot blind localization

Visual feedback





Closed-loop control performances





At a glance comparison between APA and Doppler





Static background conditions



Color Doppler

US-APA + B-mode

APL Bioengineering

Disturbance: fluid flow motion

Disturbance: tissue motion

Disturbance: fluid flow in bifurcation



S Pane, M Zhang, V Iacovacci, L Zhang, A Menciassi, *APL Bioengineering*, (2022).





Motion detection by US imaging: Could more traditional solutions work?

APA requires specific US imaging devices (need for RF data) and low temporal performances (few Hz)

Difficult translation

There exist other motion detection strategies with higher output rates and based on standard Bmode One option is OPTICAL FLOW





Comparison between APA and Optical Flow

The comparison between the two techniques was carried out considering:

- Different microrobot dimensions (from 1200 to 250 μm in diameter)
- Different locomotion patterns (rolling and vibration from 5 to 1 Hz)
 - Different environmental conditions (vascular and tissue-like)





Comparison between APA and Optical Flow

Optical Flow consistently achieved submillimetric tracking accuracies in all tested conditions (error < 0.6 *body length* for rolling ~1 *body length* for vibration)

Spatial performances are comparable to US-APA with no need for RF data

Major increase in output rate from 1-2 Hz up to 40 Hz





Conclusions

- Many challenges are still ahead, but motivations are strong!
- It is time to think about more advanced paradigms for microrobotic control, application, human interaction...
- Open research platforms and published datasets are key for developing / testing new methodologies
- Identifying the correct balance between AI methods and physical modelling in the miniature/micro domain



Surgical Robotics and Allied **Technologies** Area



Thank you for your attention!

THE BIOROBOTICS INSTITUTE

> Sant'Anna School of Advanced Studies – Pisa







