A wearable cardiac ultrasound imager

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Hongjie Hu^{1,15}, Hao Huang^{1,15}, Mohan Li^{2,15}, Xiaoxiang Gao^{1,15}, Lu Yin¹, Ruixiang Qi³, Ray S. Wu¹, Xiangjun Chen⁴, Yuxiang Ma^{1,5}, Keren Shi^{4,6}, Chenghai Li⁷, Timothy M. Maus⁸, Brady Huang⁹, Chengchangfeng Lu², Muyang Lin¹, Sai Zhou⁴, Zhiyuan Lou¹, Yue Gu^{4,10}, Yimu Chen¹, Yusheng Lei^{1,11}, Xinyu Wang¹, Ruotao Wang¹, Wentong Yue¹, Xinyi Yang⁴, Yizhou Bian¹, Jing Mu⁴, Geonho Park¹, Shu Xiang¹², Shengqiang Cai^{4,7}, Paul W. Corey¹³, Joseph Wang^{1,4} & Sheng Xu^{1,2,4,9,14}

Continuous imaging of cardiac functions is highly desirable for the assessment of long-term cardiovascular health, detection of acute cardiac dysfunction and clinical management of critically ill or surgical patients¹⁻⁴. However, conventional non-invasive approaches to image the cardiac function cannot provide continuous measurements owing to device bulkiness⁵⁻¹¹, and existing wearable cardiac devices can only capture signals on the skin¹²⁻¹⁶. Here we report a wearable ultrasonic device for continuous, real-time and direct cardiac function assessment. We introduce innovations in device design and material fabrication that improve the mechanical coupling between the device and human skin, allowing the left ventricle to be examined from different views during motion. We also develop a deep learning model that automatically extracts the left ventricular volume from the continuous image recording, yielding waveforms of key cardiac performance indices such as stroke volume, cardiac output and ejection fraction. This technology enables dynamic wearable monitoring of cardiac performance with substantially improved accuracy in various environments.

The device features piezoelectric transducer arrays, liquid metal composite electrodes and triblock copolymer encapsulation, as shown by the exploded schematics (Fig. 1a, left, Extended Data Fig. 1 and Supplementary Discussion 3). The device is built on styrene–ethylene– butylene–styrene (SEBS). To provide a comprehensive view of the heart, standard clinical practice is to image it in two orthogonal orientations by rotating the ultrasound probe¹⁷. To eliminate the need for manual rotation, we designed the device with an orthogonal configuration (Fig. 1a, right and Supplementary Videos 1 and 2). Each transducer element consisted of an anisotropic 1-3 piezoelectric composite and a silver-epoxy-based backing layer^{18,19}. To balance the penetration depth and spatial resolution, we chose a centre resonant frequency of 3 MHz for deep tissue imaging¹⁹ (Supplementary Fig. 1). The array pitch was 0.4 mm (that is, 0.78 ultrasonic wavelengths), which enhances lateral resolutions and reduces grating lobes²⁰.

To individually address each element in such a compact array, we made high-density multilayered stretchable electrodes based on a composite of eutectic gallium–indium liquid metal and SEBS²¹. The composite is highly conductive and easy to pattern (Fig. 1b,c, Supplementary Figs. 2–4 and Methods). Lap shear measurements show that the interfacial bonding strength is about 250 kPa between the transducer element and the SEBS substrate, and about 236 kPa between the transducer element and the composite electrode (Fig. 1d and

Supplementary Fig. 5), which are both stronger than typical commercial adhesives²² (Supplementary Table 2). The resulting electrode has a thickness of only about 8 µm (Supplementary Figs. 6 and 7). Electromagnetic shielding, also made of the composite, can mitigate the interference of ambient electromagnetic waves, which reduces the noise in the ultrasound radiofrequency signals and enhances the image quality²³ (Supplementary Fig. 8 and Supplementary Discussion 4). The device has excellent electromechanical properties, as determined by its high electromechanical coupling coefficient, low dielectric loss, wide bandwidth and negligible crosstalk (Supplementary Fig. 1 and Methods). The entire device has a low Young's modulus of 921 kPa, comparable with the human skin modulus²⁴ (Supplementary Fig. 9). The device exhibits a high stretchability of up to approximately 110% (Fig. 1e and Supplementary Fig. 10) and can withstand various deformations (Fig. 1f). Considering that the typical strain on the human skin is within 20% (ref.¹⁹), these mechanical properties allow the wearable imager to maintain intimate contact with the skin over a large area, which is challenging for rigid ultrasound devices²⁵.

Imaging strategies and characterizations

We evaluated the quality of the generated images based on the five most crucial metrics for anatomical imaging: spatial resolutions (axial,

¹Department of Nanoengineering, University of California San Diego, La Jolla, CA, USA. ²Department of Electrical and Computer Engineering, University of California San Diego, La Jolla, CA, USA. ³Department of Computer Science and Engineering, University of California San Diego, La Jolla, CA, USA. ⁴Materials Science and Engineering Program, University of California San Diego, La Jolla, CA, USA. ⁴Materials Science and Engineering Program, University of California San Diego, La Jolla, CA, USA. ⁴Materials Science and Engineering Program, University of California, San Diego, La Jolla, CA, USA. ⁵Department of Mechanical Engineering, Massachusetts Institute of Technology, Cambridge, MA, USA. ⁶Materials Science and Engineering Program, University of California, Riverside, CA, USA. ⁷Department of Mechanical and Aerospace Engineering, University of California San Diego, La Jolla, CA, USA. ⁷Department of Anesthesiology, University of California, San Diego Health Sulpizio Cardiovascular Center, La Jolla, CA, USA. ⁹Department of Radiology, School of Medicine, University of California San Diego, La Jolla, CA, USA. ¹⁰Department of Neurosurgery, Yale University, New Haven, CT, USA. ¹¹Department of Engineering, Stanford University, Stanford, CA, USA. ¹²Softsonics, Inc., San Diego, CA, USA. ¹³Department of Bioengineering, University of California San Diego, La Jolla, CA, USA. ¹⁵These authors contributed equally: Hongjie Hu, Hao Huang, Mohan Li, Xiaoxiang Gao. ⁶³E-mail: shengxu@ucsd.edu



Fig. 1 | **Design and characterization of the wearable cardiac imager. a**, Schematics showing the exploded view of the wearable imager, with key components labelled (left) and its working principle (right). **b**, Resistance of the liquid metal composite electrode as a function of uniaxial tensile strain. The electrode can be stretched up to about 750% without failure. The *y* axis is the relative resistance defined as R/R_0 , in which R_0 and R are the measured resistances at 0% strain and a given strain, respectively. The inset is a scanning electron micrograph of the liquid metal composite electrodes with a width as small as about 30 µm. Scale bar, 50 µm. **c**, Cycling performance of the electrode

lateral and elevational), signal-to-noise ratio, location accuracies (axial and lateral), dynamic range and contrast-to-noise-ratio²⁶.

The transmit beamforming strategy is critical for image quality. Therefore, we compared three distinct strategies: plane-wave, monofocus and wide-beam compounding. Phantoms containing monofilament wires were used for this comparison (Supplementary Fig. 11, between 0% and 100% uniaxial tensile strain, showing the robustness of the electrode. The inset shows the zoomed-in features of the graph during cyclic stretching and relaxing of the electrode. **d**, Lap shear strength of the bonding between transducer elements and SEBS or liquid metal composite electrode. Data are mean and s.d. from n = 3 tests. The inset is a schematic setup of the lap shear test. **e**, Finite element analysis of the entire device under 110% biaxial stretching. **f**, Optical images showing the mechanical compliance of the wearable imager when bent on a developable surface, wrapped around a non-developable surface, poked and twisted. Scale bars, 5 mm.

position 1). Among the three strategies, the wide-beam compounding implements a sequence of divergent acoustic waves with a series of transmission angles, and the generated images of each transmission are coherently combined to create a compounding image, which has the best quality with an expanded sonographic window²⁷ (Fig. 2a,b and Supplementary Figs. 12–14). We also used a receive beamforming



Fig. 2 | **B-mode imaging strategies and characterizations. a**, Imaging results on wire (100 μm in diameter) phantoms using different transmit beamforming strategies. The first three columns show the images through plane-wave, monofocus and wide-beam compounding at different depths, respectively. The fourth column shows the imaging resolution of wide-beam compounding in the elevational direction. The bottom row shows images of laterally distributed wires by the wide-beam compounding, from which the lateral accuracy and spatial resolutions at different lateral distances from the central axis can be obtained. **b**, Signal-to-noise ratios as a function of the imaging depth under

different transmission strategies. **c**, Simulated acoustic fields of the wide-beam compounding, with enhanced acoustic field across the entire insonation area. **d**, Elevational, lateral and axial resolutions of the device using wide-beam compounding at different depths. **e**, Lateral and axial resolutions of the device using wide-beam compounding with different lateral distances from the central axis. Data in **d** and **e** are mean and s.d. from five tests (*n* = 5). **f**, Imaging inclusions with different contrasts to the matrix. On the basis of these B-mode images, the dynamic range (**g**) and contrast-to-noise ratio (**h**) of the device can be quantified.

strategy to further improve the image quality (Supplementary Fig. 15 and Methods). The wide-beam compounding achieves a synthetic focusing effect and, therefore, a high acoustic intensity across the entire insonation area (Fig. 2c and Supplementary Fig. 13), which leads to the best signal-to-noise ratio and spatial resolutions (Fig. 2a, third column, Fig. 2b and Supplementary Fig. 12).

To quantify the device spatial resolutions using the wide-beam compounding strategy, we measured full widths at half maximum from the point spread function curves²⁸ extracted from the images (Fig. 2a, third and fourth columns and the bottom row and Supplementary Fig. 11, positions 1 and 2). As the depth increases, the elevational resolution deteriorates (Fig. 2d) because the beam becomes more divergent in the elevational direction. Therefore, we integrated six small elements into a long element (Extended Data Fig. 1) to offer better acoustic beam convergence and elevational resolution. The lateral resolution deteriorates only slightly with depth (Fig. 2d) owing to the process of receive beamforming (Methods). The axial resolution remains almost constant with depth (Fig. 2d) because it depends only on the frequency and bandwidth of the transducer array. Similarly, at the same depth, the axial resolution remains consistent with different lateral distances from the central axis of the device, whereas the lateral resolution is the best at the centre, where there is a high overlap of acoustic beams after compounding (Fig. 2e and Methods).

Another critical metric for imaging is the location accuracies. The agreements between the imaging results and the ground truths (the red dots in Fig. 2a) in the axial and lateral directions are 96.01% and 95.90%, respectively, indicating excellent location accuracies (Methods).

Finally, we evaluated the dynamic range and contrast-to-noise ratio of the device using the wide-beam compounding strategy. Phantoms containing cylindrical inclusions with different acoustic impedances

were used for the evaluation (Supplementary Fig. 11, position 3). A high acoustic impedance mismatch results in images with high contrast (Fig. 2f). We extracted the average grey values of the inclusion images and performed a linear regression²⁹, and determined the dynamic range to be 63.2 dB (Fig. 2g, Supplementary Fig. 16 and Methods), which is well above the 60-dB threshold typically used in medical diagnosis³⁰.

We selected two regions of interest, one inside and the other outside each inclusion area, to derive the contrast-to-noise ratios³¹, which range from 0.63 to 2.07 (Fig. 2h and Methods). A higher inclusion contrast leads to a higher contrast-to-noise ratio of the image. The inclusions with the lowest contrast (+3 dB or -3 dB) can be clearly visualized, demonstrating the outstanding sensitivity of this device²⁰. The performance of the wearable imager is comparable with that of the commercial device (Supplementary Figs. 17 and 18, Extended Data Table 1 and Supplementary Discussion 5).

Echocardiography from several views

Echocardiography is commonly used to examine the structural integrity and blood-delivery capabilities of the heart. Uniquely for soft devices, the contours of the human chest cause a non-planar distribution of the transducer elements, which leads to phase distortion and therefore image artefacts³². We used a three-dimensional scanner to collect the chest curvature to compensate for element position shifts within the wearable imager and thus correct phase distortion during transmit and receive beamforming (Supplementary Fig. 19, Extended Data Fig. 2 and Supplementary Discussion 6).

We compared the performance of the wearable device with a commercial device in four primary views of echocardiography, in which critical cardiac features can be identified (Extended Data Fig. 3). Figure 3a shows the schematics and corresponding B-mode images of these four views, including apical four-chamber view, apical two-chamber view, parasternal long-axis view and parasternal short-axis view. The difference between the results from the wearable and commercial devices is negligible. The parasternal short-axis view is particularly useful for evaluating the contractile function of the myocardium based on its motion in the radial direction and its relative thickening, as both are easily seen from this view. During contraction and relaxation, healthy myocardium undergoes strain and the wall thickness changes accordingly: thickening during contraction and thinning during relaxation. The strength of the left ventricle's contractile function can be directly reflected on the ultrasound image through the magnitude of the myocardial strain. Abnormalities in the contractile function, such as akinesia, can be indicative of ischaemic heart disease and myocardial infarction³³.

To better localize the specific segment of the left ventricular wall that is potentially pathological, the 17-segment model can be adopted as in standard clinical practice³³ (Fig. 3b). We took the basal, mid-cavity and apical slices of the parasternal short-axis view from the left ventricular wall, and divided them into segments according to the model. Each segment is linked to a certain coronary artery, allowing ischaemia in the coronary arteries to be localized on the basis of akinesia in the corresponding myocardial segment³³. We then recorded the displacement waveforms of the myocardium boundaries (Fig. 3c and Supplementary Discussion 6). The two peaks in each cardiac cycle in the displacement curves correspond to the two inflows into the left ventricle during diastole. The wall displacements, as measured in the basal, mid-cavity and apical views, become sequentially smaller owing to the decreasing radius of the myocardium along the conical shape of the left ventricle.

Motion-mode (M-mode) images track activities over time in a one-dimensional target region^{34,35}. We extracted M-mode images from parasternal long-axis view B-mode images (Fig. 3d). Primary targets include the left ventricular chamber, the septum and the mitral/aortic valves. In M-mode, structural information, such as the myocardial thickness and the left ventricular diameter, can be tracked according to the distances between the boundaries of features. Valvular functions, for

example, their opening and closing velocities, can be evaluated on the basis of the distance between the leaflet and septal wall (Supplementary Discussion 1). Moreover, we can correlate the mechanical activities in the M-mode images with the electrical activities in the electrocardiogram measured simultaneously during different phases in a cardiac cycle (Fig. 3d and Supplementary Discussions 1 and 6).

Monitoring during motion

Stress echocardiography assesses cardiac responses to stress induced by exercise or pharmacologic agents, which may include new or worsened ischaemia presenting as wall-motion abnormalities, and is crucial in the diagnosis of coronary artery diseases³⁶. Furthermore, subjects with heart failure may sometimes seem asymptomatic at rest, as the heart sacrifices its efficiency to maintain the same cardiac output^{37,38}. Thus, by pushing the heart towards its limits during exercise, the lack of efficiency becomes apparent. However, in current procedures, ultrasound images are obtained only before and after exercise. With the cumbersome apparatus, it is impossible to acquire data during exercise, which may contain invaluable real-time insights when new abnormalities initiate³⁹ (Supplementary Discussion 7). Also, because images are traditionally obtained after exercise, a quick recovery can mask any transient pathologic response during stress and lead to false-negative $examinations^{40}. In addition, the end point for terminating the exercise$ is subjective, which may result in suboptimal testing.

The wearable ultrasonic patch is ideal for overcoming these challenges. The device can be attached to the chest with minimal constraint to the movement of the subject, providing a continuous recording of cardiac activities before, during and after exercise with negligible motion artefacts (Extended Data Fig. 4). This not only captures the real-time responses during the test but also offers objective data to standardize the end point and enhances patient safety throughout the test (Supplementary Discussion 7). We used liquidus silicone as the couplant to achieve images of stable quality instead of water-based ultrasound gels that evaporate over time (Supplementary Figs. 20 and 21 and Supplementary Discussion 8). We observed no skin irritation or allergy after 24 h of continuous wear (Supplementary Fig. 22). The heart rate of the subject remained stable with a constant device temperature of about 32 °C after the device continuously worked for 1 h (Supplementary Fig. 23). In addition, one device was tested on different subjects (Supplementary Fig. 24). The reproducible results indicate the stable and reliable performance of the wearable imager.

We performed stress echocardiography to demonstrate the performance of the device during exercise (Supplementary Discussion 7). The device was attached to the subject for continuous recording along the parasternal long axis during the entire process, which consisted of three main stages (Fig. 4a). In the rest stage, the subject laid in the supine position. In the exercise stage, the subject exercised on a stationary bike with several intervals until a possible maximal heart rate was reached. In the recovery stage, the subject was placed in the supine position again. The device demonstrated uninterrupted tracking of the left ventricular activities, including the corresponding M-mode echocardiography and synchronized heart-rate waveform (Fig. 4b,c, Extended Data Fig. 5 and Supplementary Video 3). We examined a representative section of each testing stage and extracted the left ventricular internal diameter end systole (LVIDs) and left ventricular internal diameter end diastole (LVIDd) (Fig. 4d). The LVIDs and LVIDd of the subject remained stable during the rest stage (Fig. 4e). In the exercise stage, the interventricular septum and left ventricular posterior wall of the subject moved closer to the skin surface, with the latter moving more than the former, resulting in a decrease in LVIDs and LVIDd. In the recovery stage, the LVIDs and LVIDd slowly returned to normal. The variation in fractional shortening, a measure of the cardiac muscular contractility, reflects the changing demand for blood supply in different stages of stress echocardiography (Fig. 4e). Particularly, section 4 in Fig. 4b includes periods of exercise



Fig. 3 | Echocardiography in several standard views. a, Schematics and B-mode images of cardiac anatomies from the wearable and commercial imagers. The wearable imager was placed in the parasternal position for imaging in the parasternal long-axis and short-axis views and relocated at the apical position for imaging in the apical four-chamber and two-chamber views. b, 17-segment model representation of the left ventricular wall. Each of the concentrically nested rings that make up the circular plot represents the parasternal short-axis view of the myocardial wall from a different level of the left ventricle. c, B-mode images of the left ventricle in basal, mid-cavity and apical views (top row) and corresponding typical displacement for segments 3, 10 and 14, respectively (bottom row). The physical regions of the left ventricular wall represented by each segment of the 17-segment model have been labelled on the corresponding short-axis views. The peaks are marked with red dots. d, M-mode images (upper left) extracted from parasternal long-axis view and corresponding electrocardiogram signals (lower left). A zoomed-in plot shows the different phases of a representative cardiac cycle (right). Primary events include diastole and opening of the mitral valve during the P-wave of the electrocardiogram, opening of the aortic valve and systole during the QRS complex and closure of the aortic valve during the T-wave. AC, atrial contraction; AMVL, anterior mitral valve leaflet; C.I., commercial imager; ERF, early rapid filling; Ej., ejection; IVCT, isovolumetric contraction time; IVRT, isovolumetric relaxation time; IVS, interventricular septum; LA, left atrium; LV, left ventricular internal diameter end diastole; LVIDs, left ventricular internal diameter end systole; LVOT, left ventricular outflow tract; LVPW, left ventricular posterior wall; MV, mitral valve; RA, right atrium; RV, right ventricle; TV, tricuspid valve; W.I., wearable imager.



Fig. 4 | **Monitoring during motion. a**, Three stages of stress echocardiography. In the rest stage, the subject laid supine for around 4 min. In the exercise stage, the subject rode a stationary bike for about 15 min, with intervals for rest. In the recovery stage, the subject laid supine again for about 10 min. The wearable imager was attached to the chest of the subject throughout the entire test, even during the transitions between the stages. **b**, Continuous M-mode echocardiography extracted from the parasternal long-axis-view B-mode images of the entire process. Key features of the interventricular septum and left ventricular posterior wall are identified. The stages of rest, exercise (with intervals of rest) and recovery are labelled. **c**, Variations in the heart rate extracted from the M-mode echocardiography. **d**, Zoomed-in images of sections 1 (rest), 2 (exercise) and 3 (recovery) (dashed boxes) in **b**. **e**, Left ventricular internal diameter end diastole (LVIDd) and left ventricular internal diameter end systole (LVIDs) waveforms of the three different sections of the recording and corresponding average fractional shortenings. **f**, Zoomed-in images of section 4 (dashed box) during exercise with intervals of rest in **b**. In the first interval, the subject took a rhythmic deep breath six times, whereas during exercise, there seems to be no obvious signs of a deep breath, probably because the subject switched from diaphragmatic (rest) to thoracic (exercise) breathing, which is shallower and usually takes less time.

and intervals for rest, when patterns of a deep breath can also be seen from the left ventricular posterior wall motions (Fig. 4f).

Automatic image processing

Cardiovascular diseases are often associated with changes in the pumping capabilities of the heart, which could be measured by stroke volume, cardiac output and ejection fraction. Non-invasive, continuous monitoring of these indices are valuable for the early detection and surveillance of cardiovascular conditions (Supplementary Discussion 9). Critical information embodied in these waveforms may help precisely determine potential risk factors and track the health state⁴¹ (Supplementary Discussion 10). On the other hand, processing of the unprecedented image data streams, if done manually, can be overwhelming for clinicians, which potentially introduces interobserver variability or even errors⁴².

Automatic image processing can overcome the challenges. We applied a deep learning neural network to extract key information (for example, the left ventricular volume in apical four-chamber view) from the continuous stream of images (Fig. 5a, Supplementary Fig. 25 and Supplementary Discussion 11). We evaluated different types of deep learning models⁴³ through the output images and waveforms of the left ventricular volume (Extended Data Figs. 6 and 7, Supplementary Table 3 and Supplementary Video 4). The FCN-32 model outperforms others based on qualitative and quantitative analyses (Supplementary Fig. 26, Supplementary Tables 4 and 5 and Supplementary Discussion 11). We also applied data augmentation to expand the dataset and improve the performance (Supplementary Fig. 27 and Supplementary Discussion 11).



Fig. 5 | **Automatic image processing by deep learning. a**, Schematic workflow. Pre-processed images are used to train the FCN-32 model. The trained model accepts the unprocessed images and predicts the left ventricular (LV) volume, based on which stroke volume, cardiac output and ejection fraction are derived. b, Left ventricular volume waveform generated by the FCN-32 model from both the wearable imager (W.I.) and the commercial imager (C.I.) (left). Critical features are labelled in one detailed cardiac cycle (right). c, Bland–Altman analysis of the average of (*x* axis) and the difference between (*y* axis) the model generated and manually labelled left ventricular volumes for the wearable (black) and commercial (red) imagers. Dashed lines indicate the 95% confidence interval and about 95% of the data points are within the interval for both imagers. Solid lines indicate mean differences. d, Comparing the stroke volume, cardiac output and ejection fraction extracted from results by the wearable and commercial imagers. Data are mean and s.d. from twelve cardiac cycles (n = 12). **e**, The model-generated left ventricular volume waveform in the recovery stage. **f**, Three representative sections of the recording from the initial, middle and end stages of **e**. End-systolic volume (ESV), end-diastolic volume (EDV), stroke volume and ejection fraction (**g**) and cardiac output and heart rate waveforms (**h**) derived from the left ventricular volume waveform. The end-systolic volume and end-diastolic volume gradually recover to the normal range in the end section. The stroke volume increases from about 60 ml to about 70 ml. The ejection fraction decreases from about 80% to about 60%. The cardiac output decreases from about 11 l min⁻¹ to about 9 l min⁻¹, indicating that the decreasing heart rate from about 175 bpm to about 130 bpm overshadowed the increasing stroke volume. AS, atrial systole; IC, isovolumetric contraction; IR, isovolumetric relaxation; RI, rapid inflow.

The output left ventricular volumes for the wearable and commercial imagers show similar waveform morphologies (Fig. 5b, left). From the waveforms, corresponding phases of a cardiac cycle can be identified (Fig. 5b, right and Extended Data Fig. 8). Bland–Altman analysis gives a quantitative comparison between the model-generated and manually labelled left ventricular volumes, indicating a stable and reliable performance of the FCN-32 model⁴⁴ (Fig. 5c and Supplementary Discussion 11). The mean differences in the left ventricular volume are both approximately –1.5 ml, which is acceptable for standard medical diagnosis⁴⁵. We then derived stroke volume, cardiac output and ejection fraction from the left ventricular volume waveforms. No marked difference is observed in the averages or standard deviations between the two devices (Fig. 5d). The results verified the comparable performance of the wearable imager to the commercial imager.

The left ventricular volume is constantly changing and generally follows a steady-state pattern at rest in healthy subjects. Therefore, stroke volume, cardiac output and ejection fraction also tend to remain constant. However, cardiac pathologies or ordinary daily activities such as exercise may dynamically change those indices. To validate the performance of the wearable imager under dynamic situations, we extracted the left ventricular volume from recordings in the recovery stage of stress echocardiography (Fig. 5e). The dimensions of the left ventricle cannot be accurately determined when the images are collected in the standing position, owing to anatomical limitations of the human body (Supplementary Fig. 28 and Supplementary Discussion 9). Owing to the deep breathing after exercise, the heart was sometimes blocked by the lungs in the image. We used an image-imputation algorithm to complement the blocked part (Supplementary Fig. 29 and Supplementary Discussion 11). The acquired waveform shows an increasing trend in the left ventricular volume. Figure 5f illustrates three representative sections of the recording taken from the beginning, middle and end of the recovery stage. In the initial section, the diastasis stage is barely noticeable because of the high heart rate. In the middle section, the diastasis stage becomes visible. In the end section, the heart rate decreases notably. The end-diastolic and end-systolic volumes are increasing, because the slowing heartbeat during recovery allows more time for blood to fill the left ventricle⁴⁶ (Fig. 5g). The stroke volume gradually increases, indicating that the end-diastolic volume increases slightly faster than the end-systolic volume (Fig. 5g). The ejection fraction decreases as heart contraction decreases during the recovery (Fig. 5g). The cardiac output reduces, indicating a larger influence brought about by the decreasing heart rate than the increasing stroke volume (Supplementary Discussion 9).

Discussion

Echocardiography is crucial in the diagnosis of cardiac diseases, but the current implementation in clinics is cumbersome and limits its application in continuous monitoring. Emerging technologies based on wearable rigid modules²⁵ or flexible patches⁴⁷ lack one or more of the ideal properties of wearable ultrasound technologies (Extended Data Table 2). In this work, we provided uninterrupted frame-by-frame acquisitions of cardiac images even when the subject was undertaking intensive exercise. In addition, the wearable imager with deep learning gave actionable information by automatically and continuously outputting curves of critical cardiac metrics, such as myocardial displacement, stroke volume, ejection fraction and cardiac output, which are highly desirable in critical care, cardiovascular disease management and sports science⁴⁸. This capability is unprecedented in conventional clinical practice⁹ and the non-invasiveness can extend potential benefits to the outpatient and athletic populations.

The implications of this technology go far beyond imaging the heart, as it can be generalized to image other deep tissues, such as the inferior vena cava, abdominal aorta, spine and liver (Supplementary Fig. 30). For example, as demonstrated in an ultrasound-guided biopsy procedure on a cyst phantom (Supplementary Fig. 31), the two orthogonal imaging sections present the entire biopsy process simultaneously, freeing up one hand of the operator (Supplementary Video 5). The uniquely enabling capability of this technology forgoes the need for an operator to constantly hold the device.

Other future efforts could ensue by further improving spatial resolutions (Supplementary Fig. 32). A three-dimensional scanner can only provide the curvature of a static human chest. To accommodate the dynamic chest curvature, advanced imaging algorithms need to be developed to compensate for the phase distortion and thus improve spatial resolutions. In addition, the wearable imager is connected to the back-end system for data processing by means of a flexible cable (Supplementary Fig. 33) and future work needs to focus on system miniaturization and integration. Besides, the FCN-32 neural network can only be applied to subjects in the training dataset at present. Its generalizability could potentially be improved by expanding the training dataset or optimizing the network with few-shot-learning⁴⁹ or reinforcement-learning⁵⁰ strategies, which will allow the model to adapt to a larger population.

Online content

Any methods, additional references, Nature Portfolio reporting summaries, source data, extended data, supplementary information, acknowledgements, peer review information; details of author contributions and competing interests; and statements of data and code availability are available at https://doi.org/10.1038/s41586-022-05498-z.

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Article Methods

Materials

Gallium–indium eutectic liquid metal, toluene, ethyl alcohol, acetone and isopropyl alcohol were purchased from Sigma-Aldrich. SEBS (G1645) was obtained from Kraton. Silicone (Ecoflex 00-30) was bought from Smooth-On as the encapsulation material of the device. Silicone (Silbione) was obtained from Elkem Silicones as the ultrasound couplant. Aquasonic ultrasound transmission gel was bought from Parker Laboratories. 1-3 composite (PZT-5H) was purchased from Del Piezo Specialties. Silver epoxy (Von Roll 3022 E-Solder) was obtained from ElS. Anisotropic conductive film cable was purchased from Elform.

Design and fabrication of the wearable imager

We designed the transducer array in an orthogonal geometry, similar to a Mills cross array (Supplementary Fig. 34), to achieve biplane standard views simultaneously. For the transducers, we chose the 1-3 composite for transmitting and receiving ultrasound waves because it possesses superior electromechanical coupling¹⁸. In addition, the acoustic impedance of 1-3 composites is close to that of the skin, maximizing the acoustic energy propagating into human tissues¹⁹. The backing layer dampens the ringing effect, broadens the bandwidth and thus improves the spatial resolution^{18,51}.

We used an automatic alignment strategy to fabricate the orthogonal array. The existing method of bonding the backing layer to the 1-3 composite was to first dice many small pieces of backing layer and 1-3 composite, and then bond each pair together one by one. A template was needed to align the small pieces. This method was of very low efficiency. In this study, we bond a large piece of backing layer with a large piece of 1-3 composite and then dice them together into small pieces with designed configurations. The diced array is then automatically aligned on adhesive tape with high uniformity and perfect alignment.

Electrodes based on eutectic gallium–indium liquid metal are fabricated to achieve better stretchability and higher fabrication resolution than existing electrodes based on serpentine-shaped copper thin film. Eutectic gallium–indium alloys are typically patterned through approaches such as stencil lithography⁵², masked deposition⁵³, inkjet printing⁵⁴, microcontact printing⁵⁵ or microfluidic channelling⁵⁶. Although these approaches are reliable, they are either limited in patterning resolution or require sophisticated photolithography or printing hardware. The sophisticated hardware makes fabrication complicated and time-consuming, which presents a challenge in the development of compact, skin-conformal wearable electronics.

In this study, we exploited a new technology for patterning. We first screen-printed a thin layer of liquid metal on a substrate. A key consideration before screen printing was how to get the liquid metal to wet the substrate. To solve this problem, we dispersed big liquid metal particles into small microparticles using a tip sonicator (Supplementary Fig. 2). When microparticles contacted air, their outermost layer generated an oxide coating, which lowered the surface tension and prevented those microparticles from aggregating. In addition, we used 1.5 wt.% SEBS as a polymer matrix to disperse the liquid metal particles because SEBS could wet well on the liquid metal surface. We also used SEBS as the substrate. Therefore, the SEBS in the matrix and the substrate could merge and cure together after screen printing, allowing the liquid metal layer to adhere to the substrate efficiently and uniformly. Then we used laser ablation to selectively remove the liquid metal from the substrate to form patterned electrodes.

The large number of piezoelectric transducer elements in the array requires many such electrodes to address each element individually. We designed a four-layered top electrode and a common ground electrode. There are SEBS layers between different layers of liquid metal electrodes as insulation. To expose all electrode layers to connect to transducer elements, we used laser ablation to drill vertical interconnect accesses²¹. Furthermore, we created a stretchable shielding layer

using liquid metal and grounded it through a vertical interconnect access, which effectively protected the device from external electromagnetic noises (Supplementary Fig. 8).

Before we attached the electrodes to the transducer array, we spin-coated toluene–ethanol solution (volume ratio 8:2) on the top of the multilayered electrode to soften the liquid-metal-based elastomer, also known as 'solvent-welding'. The softened SEBS provided a sufficient contact surface, which could help form a relatively strong van der Waals force between the electrodes and the metal on the transducer surface. After bonding the electrodes to the transducer array, we left the device at room temperature to let the solvent evaporate. The final bonding strength of more than 200 kPa is stronger than many commercial adhesives²².

To encapsulate the device, we irrigated the device in a petri dish with uncured silicone elastomer (Ecoflex 00-30, Smooth-On) to fill the gap between the top and bottom electrodes and the kerf among the transducer elements. We then cured the silicone elastomer in an oven for 10 min at 80 °C. As the filling material, it suppresses spurious shear waves from adjacent elements, effectively isolating crosstalk between the elements^{18,19}. With that being said, we think the main reason for the suppressed spurious shear waves is because of the epoxy in the 1-3 composite, which limits the lateral vibration of the piezoelectric materials. The Ecoflex as the filling material may have contributed but not played a chief role because the kerf is not too wide, only 100 to 200 µm. We lifted off the glass slide on the top electrode and directly covered the top electrode with a shielding layer. Then we lifted off the glass slide on the bottom electrode to release the entire device. Finally, screen-printing an approximately 50-µm layer of silicone adhesive on the device surface completed the entire fabrication.

Characterization of the liquid metal electrode

Existing wearable ultrasound arrays can achieve excellent stretchability by serpentine-shaped metal thin films as electrodes^{19,26}. The serpentine geometry, however, severely limits the filling ratio of functional components, precluding the development of systems that require a high integration density or a small pitch. In this study, we chose to use liquid metal as the electrode owing to its large intrinsic stretchability, which makes the high-density electrode possible. The patterned liquid metal electrode had a minimum width of about 30 μ m with a groove of about 24 μ m (Supplementary Fig. 3), an order of magnitude finer than other stretchable electrodes^{18,26,57}. The liquid metal electrode is ideal for connecting arrays with a small pitch⁵⁸.

This liquid metal electrode exhibited high conductivity, exceptional stretchability and negligible resistance change under tensile strain (Fig. 1b and Supplementary Fig. 4). The initial resistance at 0% strain was 1.74 Ω (corresponding to a conductivity of around 11,800 S m⁻¹), comparable with reported studies^{59,60}. The resistance gradually increased with strain until the electrode reached the approximately 750% failure strain (Fig. 1b and Supplementary Fig. 4). The relative resistance is a parameter widely used to characterize the change in the resistance of a conductor (that is, the liquid metal electrode in this case) under different strains relative to the initial resistance^{58–60}. The relative resistance is unitless. When the strain was 0%, the initial resistance R_0 was 1.74 Ω . When the electrode was under 750% strain, the electrode was broken and the resistance R at the breaking point was measured to be 44.87 Ω . Therefore, the relative resistance (R/R_0) at the breaking point was 25.79.

To investigate the electrode fatigue, we subjected them to 100% cyclic tensile strain (Fig. 1c). The initial 500 cycles observed a gradual increase in the electrode resistance because the liquid metal, when stretched, could expose more surfaces. These new surfaces were oxidized after contacting with air, leading to the resistance increase (Supplementary Fig. 4). After the initial 500 cycles, the liquid metal electrode exhibited stable resistance because, after a period of cycling, there were not many new surfaces exposed.

This study is the first to use liquid metal-based electrodes to connect ultrasound transducer elements. The bonding strength between them directly decides the robustness and endurance of the device. This is especially critical for the wearable patch, which will be subjected to repeated deformations during use. Therefore, we characterized the bonding strength of the electrode to the transducer element using a lap shear test. The liquid metal electrode was first bonded with the transducer element. The other sides of the electrode and the element were both fixed with stiff supporting layers. The supporting layer serves to be clamped by the tensile grips of the testing machine. Samples will be damaged if they are clamped by the grips directly. Then a uniaxial stretching was applied to the sample at a strain rate of 0.5 s⁻¹. The test was stopped when the electrode was delaminated from the transducer element. A SEBS film was bonded with a transducer element and we performed the lap shear test using the same method. The peak values of the curve were used to represent the lap shear strength (Fig. 1d). The bonding strength between the pure SEBS film and the transducer element was roughly 250 kPa, and that between the electrode and the transducer element was about 236 kPa, which were both stronger than many commercial adhesives (Supplementary Table 2). The results indicate the robust bonding between the electrode and the element, preventing the electrodes from delamination under various deformations. This robust bonding does not have any limitations on the ultrasound pressures that can be transduced.

Characterization of the transducer elements

The electromechanical coupling coefficient of the transducer elements was calculated to be 0.67, on par with that of commercial probes (0.58–0.69)⁶¹. This superior performance was largely owing to the technique for bonding transducer elements and electrodes at room temperature in this study, which protected the piezoelectric material from heat-induced damage and depolarization. The phase angle was >60°, substantially larger than most earlier studies^{18,62}, indicating that most of the dipoles in the element aligned well after bonding⁶³. The large phase angle also demonstrated the exceptional electromechanical coupling performance of the device. Dielectric loss is critical for evaluating the bonding process because it represents the amount of energy consumed by the transducer element at the bonding interface²⁰. The average dielectric loss of the array was 0.026, on par with that of the reported rigid ultrasound probes $(0.02-0.04)^{64-66}$, indicating negligible energy consumed by this bonding approach (Supplementary Fig. 1b). The response echo was characterized in time and frequency domains (Supplementary Fig. 1c), from which the approximately 35 dB signal-to-noise ratio and roughly 55% bandwidth were derived. The crosstalk values between a pair of adjacent elements and a pair of second nearest neighbours have been characterized (Supplementary Fig. 1d). The average crosstalk was below the standard -30 dB in the field, indicating low mutual interference between elements.

Characterization of the wearable imager

We characterized the wearable imager using a commercial multipurpose phantom with many reflectors of different forms, layouts and acoustic impedances at various locations (CIRS ATS 539, CIRS Inc.) (Supplementary Fig. 11). The collected data are presented in Extended Data Table 1. For most of the tests, the device was first attached to the phantom surface and rotated to ensure the best imaging plane. Raw image data were saved to guarantee minimum information loss caused by the double-to-int8 conversion. Then the raw image data were processed using the 'scanConversion' function provided in the k-Wave toolbox to restore the sector-shaped imaging window (restored data). We applied five times upsampling in both vertical and lateral directions. The upsampled data were finally converted to the dB unit using:

$$I_{\text{new}} = 20 \times \log_{10}(I_{\text{old}}) \tag{1}$$

The penetration depth was tested with a group of lines of higher acoustic impedance than the surrounding background distributed at different depths in the phantom. The penetration depth is defined as the depth of the deepest line that is differentiable from the background (6 dB higher in pixel value). Because the deepest line available in this study was at a depth of 16 cm and was still recognizable from the background, the penetration depth was determined as >16 cm.

The accuracy is defined as the precision of the measured distance. The accuracy was tested with the vertical and lateral groups of line phantoms. The physical distance between the two nearest pixels in the vertical and lateral directions was calculated as:

$$\Delta y = \frac{\text{imaging depth}}{N_{\text{pixel, vertical}} - 1}$$
(2)

$$\Delta x = \frac{\text{imaging width}}{N_{\text{pixel,lateral}} - 1}$$
(3)

We acquired the measured distance between two lines (shown as two bright spots in the image) by counting the number of pixels between the two spots and multiplying them by Δy or Δx , depending on the measurement direction. The measured distances at different depths were compared with the ground truth described in the data sheet. Then the accuracy can be calculated by:

Accuracy =
$$1 - \left| \frac{\text{computed distance}}{\text{ground truth}} - 1 \right|$$
 (4)

The lateral accuracy was presented as the mean accuracy of the four neighbouring pairs of lateral lines at a depth of 50 mm in the phantom.

The spatial resolutions were tested using the lateral and vertical groups of wires. For the resolutions at different depths, the full width at half maximum of the point spread function in the vertical or lateral directions for each wire was calculated. The vertical and lateral resolutions could then be derived by multiplying the number of pixels within the full width at half maximum by Δy or Δx , depending on the measurement direction. The elevational resolutions were tested by rotating the imager to form a 45° angle between the imager aperture and the lines. Then the bright spot in the B-mode images would reveal scatters out of the imaging plane. The same process as calculating the lateral resolutions was applied to obtain the elevational resolutions. The spatial resolutions at different imaging areas were also characterized with the lateral group of wires. Nine wires were located at ±4 cm, ± 3 cm, ± 2 cm, ± 1 cm and 0 cm from the centre. The lateral and axial resolutions of the B-mode images from those wires were calculated with the same method.

Note that the lateral resolution worsens with the depth, mainly because of the receive beamforming (Supplementary Fig. 15). There are two beamformed signals, A and B. The lateral resolution of the A point (x_1) is obviously better than that of the B point (x_2) . The fact that lateral resolution becomes worse with depth is inevitable in all ultrasound imaging, as long as receive beamforming is used.

As for different transmit beamforming methods, the wide-beam compounding is the best because it can achieve a synthetic focusing effect in the entire insonation area. The better the focusing effect, the higher the lateral resolution, which is why the lateral resolution of the wide-beam compounding is better than the other two transmit methods at the same depth. Furthermore, the multiple-angle scan used in the wide-beam compounding can enhance the resolution at high-angle areas. The multiple-angle scan combines transmissions at different angles to achieve a global high signal-to-noise ratio, resulting in improved resolutions.

The elevational resolution can only be characterized when the imaging target is directly beneath the transducer. For those targets that are far away from the centre, they are difficult to be imaged, which makes

their elevational resolutions challenging to calculate. When characterizing the elevational resolution, the device should rotate 45°. In this case, most of the reflected ultrasound waves from those wires cannot return to the device owing to the large incidence angles. Therefore, those wires cannot be captured in the B-mode images. One potential solution is to decrease the rotating angle of the device, which may help capture more wires distributed laterally in the B-mode image. However, a small rotating angle will cause the elevational image to merge with the lateral image, which increases the error of calculating the elevational resolution. Considering those reasons, we only characterized the elevational resolution of the imaging targets directly beneath the transducer array.

The contrast resolution, the minimum contrast that can be differentiated by the imaging system, was tested with greyscale objects. The collected B-mode images are shown in Fig. 2. Because the targets with +3 and -3 dB, the lowest contrast available in this study, could still be recognized in the images, the contrast resolution of the wearable imager is determined as <3 dB.

The dynamic range in an ultrasound system refers to the contrast range that can be displayed on the monitor. The contrast between an object and the background is indicated by the average grey value of all pixels in the object in the display. The grey value is linearly proportional to the contrast. The larger the contrast, the larger the grey value. Because the display window was using the data type 'uint8' to differentiate the greyscale, the dynamic range was defined as the contrast range with a grey value ranging from 0 to 255.

The object with -15 dB contrast has the lowest average grey value, whereas the object with +15 dB contrast has the highest (Supplementary Fig. 16). In our case, there are six objects with different contrasts to the background in the phantom. The highest grey value obtained from the object of +15 dB contrast was 159.8, whereas the lowest grey value from the object of -15 dB contrast was 38.7. We used a linear fit to extrapolate the contrasts when the corresponding average grey values were equal to 255 and 0, which corresponded to contrasts of 39.2 dB and -24.0 dB, respectively. Then the dynamic range was determined as:

Dynamic range =
$$39.2 - (-24.0) = 63.2 \text{ dB}$$
 (5)

The dead zone is defined as the depth of the first line phantom that is not overwhelmed by the initial pulses. The dead zone was tested by imaging a specific set of wire phantoms with different depths right beneath the device (Supplementary Fig. 11, position 4) directly and measuring the line phantoms that were visible in the B-mode image.

The bandwidth of the imager is defined as the ratio between the full width at half maximum in the frequency spectrum and the centre frequency. It was measured by a pulse-echo test. A piece of glass was placed 4 cm away from the device and the reflection waveform was collected with a single transducer. The collected reflection waveform was converted to the frequency spectrum by a fast Fourier transform. The full width at half maximum was read from the frequency spectrum. We obtained the bandwidth using:

$$Bandwidth = \frac{full width at half maximum}{centre frequency}$$
(6)

Contrast sensitivity represents the capability of the device to differentiate objects with different brightness contrasts²⁰. The contrast sensitivity was tested with the greyscale objects. The contrast sensitivity is defined as the contrast-to-noise ratio (CNR) of the objects having certain contrasts to the background in the B-mode image:

$$CNR = \frac{|\mu_{in} - \mu_{out}|}{\sqrt{\sigma_{in}^2 + \sigma_{out}^2}}$$
(7)

in which μ_{in} and σ_{in} are the mean and the standard deviation of pixel intensity within the object, and μ_{out} and σ_{out} are the mean and the standard deviation of pixel intensity of the background.

The insertion loss is defined as the energy loss during the transmission and receiving. It was tested in water with a quartz crystal, a function generator with an output impedance of 50 Ω and an oscilloscope (Rigol DS1104). First, the transducer received an excitation in the form of a tone burst of a 3-MHz sine wave from the function generator. Then the same transducer received the echo from the quartz crystal. Given the 1.9-dB energy loss of the transmission into the quartz crystal and the 2.2 × 10⁻⁴ dB (mm MHz)⁻¹ attenuation of water, the insertion loss could be calculated as:

Insertion loss =
$$20 \times \log_{10} \left(\frac{V_{\rm r}}{V_{\rm t}} \right) + 1.9 + 2.2 \times 10^{-4} \times 2d \times f_{\rm r}^2$$
 (8)

Simulation of the acoustic field

The simulation computes the root mean square of the acoustic pressure at each point in the defined simulation field. The root mean square is defined in the equation below and gives an average acoustic pressure over a certain time duration, which is pre-defined in a packaged function of the software. In the equation, x_i is the simulated acoustic pressure at the *i*th time step.

$$x_{\rm RMS} = \sqrt{\frac{1}{n} (x_1^2 + x_2^2 + \dots + x_n^2)}$$
(9)

Figure 2c is the simulated root mean square of the transmitted acoustic pressure field by the orthogonal transducers. The simulation was done using the MATLAB UltraSound Toolbox⁶⁷. Each one-dimensional phased array in the orthogonal transducers gives a sector-shaped acoustic pressure field. The simulation merges two such sector-shaped acoustic pressure fields. The imaging procedure was done with the same parameters as the simulations.

In the simulation, we defined the transducer parameters first: the centre frequency of the transducers as 3 MHz, the width of the transducers as 0.3 mm, the length of the transducers as 2.3 mm, the pitch of the array as 0.4 mm, the number of elements as 32 and the bandwidth of the transducers as 55%. Then we defined wide-beam compounding (Supplementary Fig. 13) as the transmission method: 97 transmission angles, from -37.5° to $+37.5^{\circ}$, with a step size of 0.78° . Then the acoustic pressure field was the overall effect of the 97 transmissions. Finally, we defined the computation area: -8 mm to +8 mm in the lateral direction, -6 mm to +6 mm in the elevational direction and 0 mm to 140 mm in the axial direction.

Data availability

 $\label{eq:alpha} All\,data\,are\,available\,in\,the\,manuscript\,or\,Supplementary\,Information.$

Code availability

The code that produced the findings of this study is available at https://github.com/UCSD-XuGroup/Wearable-Cardiac-Ultrasound-Imager.

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Author contributions H. Hu, H. Huang, M. Li, X.G. and S. Xu designed the research. H. Hu, H. Huang, M. Li and L.Y. performed the experiments. X.G., R.Q. and M. Li designed and trained the neural network. H. Hu, H. Huang, M. Li and Y.M. performed the data processing and simulations. H. Hu, H. Huang and S.Xu analysed the data. H. Hu, H. Huang, M. Li, R.S.W., R.Q., S. Xiang, J.W. and S. Xu wrote the paper. All authors provided constructive and valuable feedback on the manuscript.

Competing interests The authors declare no competing interests.

Additional information

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Correspondence and requests for materials should be addressed to Sheng Xu.

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Extended Data Fig. 1 | Schematics and optical images of the orthogonal imager. a, The orthogonal imager consists of four arms, in which six small elements in one column are combined as one long element, and a central part that is shared by the four arms. The blue and red boxes label a long element integrated by six small pieces in each direction. The number of elements in one direction is 32. The pitch between the elements is 0.4 mm. Optical images in

top view (**b**) and isometric view (**c**) showing the morphology of the orthogonal array. We used an automatic alignment strategy to fabricate the orthogonal array by bonding a large piece of backing layer with a large piece of 1-3 composite and then dicing them together into small elements with designed configurations. Inset in **c** shows the details of the elements. The 1-3 composite and backing layer have been labelled.



Extended Data Fig. 2 | Characterization of the effects of phase correction on imaging quality. B-mode images of a line phantom obtained from different situations (a). Left, from a planar surface. Middle, from a curvilinear surface without phase correction. Right, from a curvilinear surface with phase correction. b, The axial and lateral resolutions at different depths under these three situations. No obvious difference in axial resolution was found because it is mainly dependent on the transducer frequency and bandwidth. The lateral resolution of the wearable imager was improved after phase correction. Images collected in (**c**), the parasternallong-axis (PLAX) view of the heart and (**d**), the apical four-chamber view when measured by a planar probe (left panel), a curved probe without phase correction (middle panel) and a curved probe with phase correction (right panel). The left ventricular boundaries are labelled by white dashed lines in the images. **e**, Comparison of measured cardiac indices showing the impact of phase correction. Each measurement is based on the mean of five consecutive cardiac cycles (n = 5). The standard deviations are indicated by the error bars.



Extended Data Fig. 3 | **Optical images showing positions and orientations for ultrasound heart imaging. a**, Parasternal long-axis view. **b**, Parasternal short-axis view. **c**, Apical four-chamber view. **d**, Apical two-chamber view. The orthogonal wearable cardiac imager combines parasternal long-axis and short-axis views (e) and apical four-chamber and apical two-chamber views without rotation (f). The wearable imager can capture two parasternal views from a single position or two apical views from another single position. The sternum and ribs are labelled to indicate intercostal spaces.



Extended Data Fig. 4 | **B-mode images collected from a subject with different postures.** The four views collected when the subject is sitting (**a**), standing (**b**), bending over (**c**), lying flat (**d**) and lying on their side (**e**). The PLAX and PSAX views can keep their quality at different postures, whereas the quality

of A4C and A2C views can only be achieved when lying on the side. A2C, apical two-chamber view; A4C, apical four-chamber view; PLAX, parasternal long-axis view; PSAX, parasternal short-axis view.



Extended Data Fig. 5 | Continuous cardiac imaging during rest, exercise and recovery. Representative B-mode and M-mode images during rest (a), exercise (b) and recovery (c). The red line highlights the M-mode section corresponding to the current B-mode frame. More details can be seen in Supplementary Video 3.



Extended Data Fig. 6 | Segmentation results of the left ventricle with different deep learning models. By qualitatively evaluating the result, we found no 'jitteriness' in Supplementary Video 4. The segmented left ventricle contracts and relaxes as naturally as the B-mode video. The

segmentation boundaries are smooth with the highest fidelity. Compared with the original B-mode image, the FCN-32 model has the best agreement among all models used in this study.



Extended Data Fig. 7 | Waveforms of the left ventricular volume obtained with different deep learning models. Those waveforms are from segmenting the same B-mode video. Qualitatively, the waveform generated by the FCN-32 model gains the best stability and the least noise, and the waveform morphology

is more constant from cycle to cycle. Quantitatively, the comparison results of those models is in Supplementary Fig. 26, which shows that the FCN-32 model has the highest mean intersection over union, showing the best performance in this study.



Extended Data Fig. 8 | Different phases in a cardiac cycle obtained from B-mode imaging. The rows are B-mode images of A4C, A2C, PLAX and PSAX views in the same phase. The columns are B-mode images of the same view during ventricular filling, atrial contraction, isovolumetric contraction, end of ejection and isovolumetric relaxation. The dashed lines highlight the main features of the current phase. Bluish lines mean shrinking in the volume of the

labelled chamber. Reddish lines mean expansion in the volume of the labelled chamber. Yellowish lines mean retention in the volume of the labelled chamber. A2C, apical two-chamber view; A4C, apical four-chamber view; LA, left atrium; LV, left ventricle; LVOT, left ventricular outflow tract; RA, right atrium; RV, right ventricle; PLAX, parasternal long-axis view; PSAX: parasternal short-axis view.

Extended Data Table 1 | Full comparison of the imaging metrics between the wearable imager and a commercial ultrasound imager (Model P4-2v)

| | Wearable imager | Commercial imager (32 active elements) | Commercial imager (64 active elements) |
|-----------------------------------|--|--|--|
| Depth of penetration (cm) | > 16 | > 16 | > 16 |
| Axial accuracy (%) | 98.7 (40 mm) 96.6 (70 mm) 96.0 (110 mm) | 99.0 (40 mm) 95.9 (70 mm) 98.9 (110 mm) | 99.3 (40 mm) 95.6 (70 mm) 99.0 (110 mm) |
| Lateral accuracy (%) | 95.9 | 96.5 | 97.0 |
| Axial resolution (mm) | 0.59 (40 mm) 0.65 (70 mm) 0.62 (110 mm) | 0.58 (40 mm) 0.63 (70 mm) 0.61 (110 mm) | 0.54 (40 mm) 0.55 (70 mm) 0.60 (110 mm) |
| Lateral resolution (mm) | 1.55 (40 mm) 2.27 (70 mm) 3.49 (110 mm) | 1.26 (40 mm) 1.73 (70 mm) 2.52 (110 mm) | 1.26 (40 mm) 1.57 (70 mm) 2.28 (110 mm) |
| Elevational resolution (mm) | 3.65 (40 mm) 4.61 (70 mm) 6.41 (110 mm) | 4.36 (40 mm) 2.64 (70 mm) 3.45 (110 mm) | 3.87 (40 mm) 2.30 (70 mm) 3.14 (110 mm) |
| Contrast resolution (dB) | < 3 | < 3 | < 3 |
| Dynamic range (dB) | 63.2 (-24.0~39.2) | 56.9 (-30.6~26.3) | 50.3 (-28.7~21.6) |
| Dead zone (mm) | 6 | < 1 | < 1 |
| Bandwidth (%) | 55 | 74 | 74 |
| Contrast-to- noise ratio | 1.51 (-15 dB) 0.76 (-6 dB) 0.63 (-3 dB) 1.12 (+3 dB) 1.53 (+6 dB) 2.08 (+15 dB) | 2.01 (-15 dB) 0.69 (-6 dB) 0.26 (-3 dB) 0.73 (+3 dB) 1.23 (+6 dB) 1.49 (+15 dB) | 2.24 (-15 dB) 0.90 (-6 dB) 0.33 (-3 dB) 0.59 (+3 dB) 1.10 (+6 dB) 1.82 (+15 dB) |
| Insertion loss (dB) | 24.98 | 16.68 | 16.68 |

The overall performance of the wearable imager is comparable with that of the commercial one. Because the wearable orthogonal array has 32 elements in each direction, we also provided measurements of the commercial imager with only 32 elements activated.

Extended Data Table 2 | Summary of wearable ultrasonic devices for continuous monitoring of deep tissues

| Device form factor | Imaging | Sensing mode | In vivo | Real-time | Continuous sampling in-motion | Automatic image analysis | Number of quantified functions | Citation |
|--------------------|---------|-----------------|---------|-----------|----------------------------------|-----------------------------|-----------------------------------|-----------|
| Rigid module | Yes | 2D | Yes | Yes | No | No | 2 | 25 |
| Rigid module | No | 1D | Yes | Yes | No | No | 1 | 48 |
| Rigid module | No | 1D | Yes | Yes | Yes | No | 1 | 68 |
| Flexible patch | Yes | 2D | No | Yes | No | Yes | 0 | 32 |
| Flexible patch | Yes | 2D | No | No | No | No | 0 | 28 |
| Flexible patch | No | 1D | Yes | Yes | Yes | No | 1 | 69 |
| Flexible patch | No | 1D | No | No | No | No | 0 | 70 |
| Flexible patch | Yes | 2D | Yes | Yes | No | Yes | 0 | 47 |
| Flexible patch | Yes | 2D | No | Yes | No | No | 0 | 71 |
| Flexible patch | Yes | 2D | Yes | Yes | No | No | 0 | 72 |
| Flexible patch | Yes | 2D | No | Yes | No | No | 0 | 73 |
| Flexible patch | Yes | 3D | No | No | No | No | 0 | 74 |
| Flexible patch | No | 1D | Yes | Yes | No | No | 1 | 75 |
| Stretchable patch | Yes | 2D | Yes | Yes | No | No | 1 | 26 |
| Stretchable patch | No | 1D | Yes | Yes | No | No | 1 | 76 |
| Stretchable patch | No | 1D | Yes | Yes | Yes | No | 1 | 19 |
| Stretchable patch | Yes | 2D | No | Yes | No | No | 0 | 77 |
| Stretchable patch | Yes | 3D | No | No | No | No | 0 | 18 |
| Stretchable patch | Yes | Bi-planes of 2D | Yes | Yes | Yes | Yes | 7 | This work |

'Number of quantified functions' means the amount of physiological signals collected from images. For example, the number of quantified functions in this work is seven because we can extract myocardium displacement, left ventricular internal diameter, fractional shortening, stroke volume, ejection fraction, cardiac output and heart rate from those B-mode images.

Supplementary information

A wearable cardiac ultrasound imager

In the format provided by the authors and unedited

| 1 | Supplementary Information for |
|----------|--|
| 2 | |
| 3 | |
| 4 | A wearable cardiac ultrasound imager |
| 5 | |
| 6 7 | Hongjie Hu ¹ *, Hao Huang ¹ *, Mohan Li ² *, Xiaoxiang Gao ¹ *, Lu Yin ¹ , Ruixiang Qi ³ , Ray S. Wu ¹ , |
| 8 | Xiangjun Chen ⁴ , Yuxiang Ma ^{1,5} , Keren Shi ^{4,6} , Chenghai Li ⁷ , Timothy M. Maus ⁸ , Brady Huang ⁹ , Chengchangfeng Lu ² , Muyang Lin ¹ , Sai Zhou ⁴ , Zhiyuan Lou ¹ , Yue Gu ^{4,10} , Yimu Chen ¹ , Yusheng |
| 9 | Lei ^{1,11} , Xinyu Wang ¹ , Ruotao Wang ¹ , Wentong Yue ¹ , Xinyi Yang ⁴ , Yizhou Bian ¹ , Jing Mu ⁴ , |
| 10 | Geonho Park ¹ , Shu Xiang ¹² , Shengqiang Cai ^{4,7} , Paul W. Corey ¹³ , Joseph Wang ^{1,4} , Sheng |
| 11 | $Xu^{1,2,4,9,14\#}$ |
| 12 | |
| 13 | ¹ Department of Nanoengineering, University of California San Diego, La Jolla, CA 92093, USA. |
| 14 | ² Department of Electrical and Computer Engineering, University of California San Diego, La Jolla, |
| 15 | CA 92093, USA. |
| 16 | ³ Department of Computer Science and Engineering, University of California San Diego, La Jolla, |
| 17 | CA 92093, USA. |
| 18 | ⁴ Materials Science and Engineering Program, University of California San Diego, La Jolla, CA |
| 19 | 92093, USA. |
| 20 | ⁵ Department of Mechanical Engineering, Massachusetts Institute of Technology, Cambridge, MA |
| 21 | 02139, USA. |
| 22 | ⁶ Materials science and engineering program, University of California Riverside, Riverside, CA |
| 23 | 92521, USA. ⁷ Department of Machanical and Associate Engineering, University of California San Diago, La |
| 24 25 | ⁷ Department of Mechanical and Aerospace Engineering, University of California San Diego, La Jolla, CA 92093, USA. |
| 26 | ⁸ Department of Anesthesiology, UC San Diego Health-Sulpizio Cardiovascular Center, La Jolla, |
| 20 27 | CA 92037, USA. |
| 28 | ⁹ Department of Radiology, School of Medicine, University of California San Diego, La Jolla, CA |
| 29 | 92103, USA. |
| 30 | ¹⁰ Department of Neurosurgery, Yale University, New Haven, CT 06520, USA. |
| 31 | ¹¹ Department of Chemical Engineering, Stanford University, Stanford, CA 94305, USA. |
| 32 | ¹² Softsonics, Inc., San Diego, CA 92122, USA. |
| 33 | ¹³ Department of Anesthesiology, Sharp Memorial Hospital, San Diego 92123, CA, USA. |
| 34 | ¹⁴ Department of Bioengineering, University of California San Diego, La Jolla 92093, CA, USA. |
| 35 | *These authors contributed equally to this work. |
| 36 | [#] Email: shengxu@ucsd.edu |
| 37 | |

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- 147 Supplementary Discussion 1: Significance of the wearable imager
- 148
- 149 1.1 End-systolic volume (ESV), end-diastolic volume (EDV), heart rate, stroke volume, cardiac150 output, and ejection fraction

Normal cardiac function is essential for maintaining systemic tissue perfusion throughout the body¹. Cardiovascular diseases, especially in the elderly, impose a huge burden in terms of mortality, morbidity, disability, and healthcare costs². More than a million patients are admitted annually to U.S. hospitals with acute heart failure alone, together with a high median percentage of intensive care unit admission of 10%⁶⁸ and a high in-hospital mortality rate of around 4% to 7%⁶⁹. Also, cardiovascular failure is one of the leading causes of death in intensive care units⁷⁰ and nearly one-quarter of all deaths in intensive care units are attributed to it.

158

Accurate assessment of subtle changes in cardiac functions is essential for health management and disease prevention for healthy people, as well as diagnosis of pathogenesis and interventions for patients. The signals we can use to evaluate the cardiac functions include the ESV, EDV, heart rate, as well as their derivative signals such as stroke volume, cardiac output, and ejection fraction. ESV and EDV can be obtained by processing the apical four-chamber view B-mode images using a deep learning model. The heart rate can be observed based on the period of contraction in Mmode images. Based on these values, those derivative signals can be calculated by:

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- 167
- 168
- 169
- 170
- 171

 $Ejection\ fraction = \frac{Stroke\ volume}{EDV}\ (12)$

Stroke volume = EDV - ESV (10)

Cardiac output = Stroke volume * Heart rate (11)

172

173 Stroke volume indicates the absolute blood volume the left ventricle can pump out in a single 174 stroke. The cardiac output indicates the absolute blood volume the left ventricle can pump out 175 every minute. Ejection fraction indicates the relative fraction of the blood in the left ventricle that 176 the heart can pump out in a single stroke. Altogether, these indices provide insight into the 177 capability of the heart to deliver blood to tissues throughout the body.

178

179 1.2 Cardiac functions and common pathologies

180 Cells in the human body all require a steady supply of oxygen and nutrient for their 181 metabolism. The cellular metabolic rates are not static, but rather are subject to constant 182 fluctuations. Thus, the heart must not only be able to produce a cardiac output meeting the 183 metabolic demands of the body at a given time but also do so efficiently, such that enough 184 headroom is maintained to accommodate any heightened metabolic rates that can occur due to

- 185 circumstances such as strenuous exercise. The existence of a nonzero end-systolic volume itself
- also serves a similar purpose as a buffer³⁷. Thus, the occurrence of heart failure is marked either
- 187 by an inability of the heart to provide a cardiac output meeting the metabolic demands of the body,
- 188 and/or a compromised efficiency in function and consequent lack of headroom³⁸.
- 189

190 There are two forms of heart failure: systolic and diastolic. As the names suggest, systolic 191 heart failure results from the heart's lack of ability to pump blood during systole, while diastolic 192 heart failure results from the heart's lack of ability to fill with blood during diastole.

193

During systole, the goal of the left ventricle is to eject as much of its blood volume as possible. The ability of the left ventricle to do so can be viewed as a function of three factors: (a) the contractility of the myocardium, (b) the afterload in the left ventricle, and (c) the structural integrity of the left ventricle³⁸.

- (a) The contractility acts as the driving force for blood flow out of the left ventricle and through the aortic valve. If the contractility is compromised, the left ventricle's capacity to eject blood will be reduced. This can occur due to conditions such as dilated cardiomyopathy or ischemic heart disease, which cause the myocardium to contract weakly. Alternatively, it can be a result of pathologies such as arrhythmia, in which the myocardium activates asynchronously rather than produces a single, strong impulse during systole.
- (b) Afterload is the amount of resistance encountered by the left ventricle as it attempts to
 contract. The more resistance the left ventricle encounters, the more difficult it will be to
 eject blood. This can result from conditions such as aortic stenosis, where the aortic valve
 does not open fully and restricts the blood flow, or hypertension, in which there is elevated
 pressure in the left ventricle resisting contraction.
- 211

205

- (c) The structural integrity of the left ventricle is a prerequisite for its functions. For example,
 a mitral valve defect that results in an improper seal between the left ventricle and left
 atrium can cause blood to backflow into the left atrium during systole, instead of flowing
 through the aortic valve towards the rest of the body.
- 216

During diastole, the goal of the left ventricle is to fill with as much blood as possible. This ensures that there is enough blood available to be pumped out afterwards, during systole. Similar to systole, the diastolic function can also be broken down into three contributing factors³⁸: (a) the distensibility of the left ventricle, (b) external compression, and (c) structural factors.

- 221
- (a) In diastole, the distensibility of the left ventricle serves as the driving force, allowing it to
 expand and be filled with blood. A stiffer or thicker myocardium loses its ability to expand,

resulting in decreased distensibility. For example, the protein deposits caused by amyloidosis can lead to this effect. Hypertrophy can also be caused by chronic hypertension as a result of the heart working against high afterloads⁷¹. Gradual loss of distensibility is also a natural result of aging⁷².

- (b) External compression acts as the resistive force against the distensibility of the left ventricle. External compression restricts the volume of blood that can be filled, leading to diastolic heart failure. External compression is introduced from regions external to the left ventricle, such as the pericardium in constrictive pericarditis and cardiac tamponade. It can even come from the right ventricle, as is the case in cor pulmonale, in which right ventricular failure has excessive volume. This dilation of the right ventricle creates an external compression acting on the left ventricle.
- (c) Structural factors can also affect diastole, in the form of obstructions to filling.
 Obstructions prevent the left ventricle from filling optimally during diastole, leading to a
 reduction of the end-diastolic volume. For example, mitral stenosis can prevent the mitral
 valve from opening optimally, reducing blood flow into the left ventricle from the left
 atrium. A left atrial myxoma located in the way of the mitral valve could also cause a
 similar effect.
- 243

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228

244 Additionally, diastolic heart failure can also result from systolic heart failure. When the 245 ejection fraction decreases due to systolic heart failure, causing an initial drop in stroke volume 246 and cardiac output, the heart may compensate by increasing the end-diastolic volume with higher 247 filling pressure through neurohormonal pathways that increase the vascular tone and intravascular volume^{37,38}. This invokes the Frank-Starling relationship⁷³ by creating a higher pre-load, allowing 248 249 the stroke volume to return back to normal levels while the ejection fraction remains low. Because 250 diastolic filling must occur with elevated pressure if the cardiac output is to be maintained, diastolic 251 heart failure can happen in this situation. As with nearly all human physiology, this is one of many 252 examples showing how the final indices that we observe are often the result of a long cascade of 253 interactions.

254

255 As such, the numerical indices (i.e., ESV, EDV, heart rate, stroke volume, cardiac output, and 256 ejection fraction) by themselves can only point us towards whether systolic/diastolic heart failure 257 has occurred, or possibly both have occurred. For example, in systolic heart failure, one cannot 258 determine whether it stems from a lack of contractility, increased afterload, or loss of structural 259 integrity, based on the numbers alone. Therefore, while these indices may not provide a specific 260 diagnosis, they serve as highly efficient quantitative indicators for the early detection of a broad 261 range of cardiovascular diseases. The ability to continuously monitor these indices opens the 262 opportunity for more comprehensive examinations to be performed in a timely manner.

263

264 1.3 B-mode and M-mode images

265 In this study, we demonstrate direct ultrasound B-mode and M-mode imaging of cardiac 266 structures. This capability of the wearable imager is valuable in providing a more detailed 267 diagnosis after an initial indication of cardiovascular diseases from the numerical indices. A vast 268 number of factors contributing to heart failure can be identified visually just from imaging the 269 heart continuously. For example, ischemia may result in the death of myocardial cells, producing 270 effects such as fibrillation and hypokinesia/akinesia, which can be easily observed in continuously 271 recorded images. Features related to the structural integrity of the heart and obstructions to filling 272 may also be seen, such as valvular stenosis, valvular regurgitation, septal defects, and left atrial 273 myxoma. Hypertrophic cardiomyopathy can result in an obvious thickening of the myocardium 274 that can be seen on the ultrasound recording, while dilated cardiomyopathy results in a visible 275 thinning of the myocardium and dilation of the left ventricle. Hypertrophic cardiomyopathy can 276 also be differentiated from pathologies like amyloidosis due to the extreme brightness of the 277 amyloid proteins relative to the myocardium on the ultrasound image. These are a few notable 278 examples of the vast range of diseases that can be diagnosed conclusively through wearable 279 imaging.

280

281 In this study, four standard B-mode ultrasound views of the heart were implemented: 282 parasternal long axis, parasternal short axis, apical four-chamber, and apical two-chamber. 283 Parasternal long and short axis views are orthogonal, so are apical four- and two-chamber views. 284 Each of these views provides a different perspective of the heart, with its own viewing angle and 285 set of structures, allowing for the most comprehensive heart investigations. In addition, imaging 286 the heart from these standard views allows for a more accurate representation of cardiac structural 287 dimensions. Imaging from other angles can also reveal internal structures, but the displayed 288 dimensions may have deviations from what clinicians normally measure. From the four views, the 289 symptoms related to major abnormalities in cardiac functions such as changes in myocardial 290 thickness, can be easily and accurately observed. Furthermore, diseases and activities may increase 291 the heart rate, leading to faster valvular velocities, which could be quantitatively estimated through 292 a frame-by-frame observation method: the distance of the valve movement can be detected from 293 frames in B-mode images. A faster valvular velocity leads to fewer frames to reach the same 294 distance.

295

296 1.4 Unique innovation of the wearable imager

Emerging wearable electronics in recent years have gradually transformed ultrasound technologies from traditional large-scale equipment to miniaturized wearable devices. Wearable ultrasound devices have been demonstrated for other applications, including monitoring of blood pressure, blood flow, and tissue motions (Extended Data Table 1), continuous in-motion analytical B-mode imaging of deep tissues, a major aspiration for diagnostic ultrasound, has not been 302 achieved^{18,19,26,28,32,47,48,74-83.} Recently continuous B-mode imaging on various tissues of the human 303 body was reported based on a rigid ultrasound device²⁵. The authors used a soft adhesive material 304 to fix the rigid ultrasound device on the human body and acquired only snapshots of the heart when 305 the subject was in-motion. Moreover, there was a lack of analysis of these acquired images. The 306 data from wearable devices are only valuable when we can extract actionable information from

- 306 data from we307 them.
- 308

309 In this work, the ultrasound device is soft and can collect high-quality cardiac images during 310 exercise continuously. We reconstructed the B-mode images of phantoms and various tissues (e.g., 311 liver, abdominal aorta, inferior vena cava, and biopsy process, as seen in Supplementary Figs. 30 312 and 31) with a particular focus on the heart. Because the heart has a complex and fast-moving 313 anatomical structure, continuous cardiac imaging can provide new opportunities of understanding 314 pathologies of many cardiovascular diseases. We showed bi-planar imaging from multiple 315 standard views commonly used in transthoracic echocardiography examinations. We quantified 316 the displacement of myocardium for the diagnosis of myocardial ischemia. We also used a deep 317 learning model to extract key cardiac metrics such as stroke volume, ejection fraction, and cardiac 318 output from the continuous images automatically. These contributions have never been made by 319 any wearable devices. The novelty of this work is two-fold.

- 320 321
- (1) Innovation in device engineering, including array design, microfabrication, and transmitting strategy
- 322 323

First, we used 32 elements in each row of transducers with a pitch of 0.4 mm (shorter than one ultrasound wavelength), which considerably enhanced the signal-to-noise ratio and weakened grating-lobes, leading to better spatial resolutions, stronger penetration capabilities, and less artifacts compared to existing wearable phased array images.

328

329 Second, to achieve such a large size and small pitch of the array, new microfabrication 330 techniques were required. We replaced the conventional serpentine-based copper electrodes with 331 straight liquid metal electrodes that were inherently stretchable, and micropatterned the traces with 332 screen printing and laser ablation, which significantly increased the pattern resolution, with a 333 minimum width and kerf of 0.03 mm and 0.025 mm, respectively. Such a high pattern resolution 334 enables stretchable electronics with high-density. Additionally, we replaced the manual alignment 335 of hundreds of elements with a new automatic alignment method, which was more time-efficient 336 and had a higher success rate. Moreover, we applied a room-temperature bonding method and 337 made a dense backing layer to enhance the acoustic performance of each ultrasonic transducer in 338 the array.

Third, for the packaging strategy, in previous work, we used a type of silicone elastomer (Ecoflex-0030) as the substrate and superstrate, which provided mechanical support and waterproof encapsulation to the device^{18,19}. In this work, we used a triblock copolymer (SEBS) coated by liquid metal as the substrate and superstrate. This did not only provide mechanical support and waterproof encapsulation, but also serves as the ground electrode and the shielding layer to screen any interference from ambient electromagnetic waves that might otherwise induce imaging artifacts.

347

348 Specifically, to apply the SEBS as the encapsulation material, we have tried a variety of 349 polymers and many ways of coating liquid-metal. Liquid metal has a large surface tension and 350 does not easily wet the polymer substrate. We solved this problem by mixing SEBS, dissolved in 351 toluene solution, with liquid metal to form a composite electrode, which could readily wet and 352 adhere to the SEBS substrate. Silicone elastomers did not work because they cannot be dissolved 353 by common solvents so they failed to composite with the liquid metal. This new packaging strategy 354 enables more functions and use cases and has never been realized in any reported wearable 355 ultrasonic devices.

356

Fourth, an advanced wide-beam compounding transmission method was introduced to wearable ultrasonic technologies for the first time. Compared with other traditional transmission approaches, such as single plane-wave and mono-focus, the wide-beam compounding method provides B-mode images with enhanced spatial resolutions, signal-to-noise ratios, and contrast-tonoise ratios²⁷.

362

Based on the above innovations in device engineering, we realized high-quality B-mode ultrasound imaging using a wearable ultrasonic patch. High-quality B-mode imaging-guided applications and techniques, such as Doppler-based imaging⁸⁴, elastography⁸⁵, 3D/4D imaging⁸⁶, ultrafast/super-resolution imaging^{87,88}, and ultrasound stimulation⁸⁹, can all be possible on the wearable platform.

368

369 (2) Continuous cardiac imaging during exercise and associated automatic data processing by
 370 deep learning

371

Conventionally, cardiac imaging is carried out using bulky expensive equipment while the subject stays still. Existing wearable health devices allow monitoring the human body on the go but can only capture signals on the skin surface and generate signals in the form of data points and curves. Wearable imaging of central organs (e.g., the heart) remains a grand challenge in the field. Particularly, the heart is relatively difficult to image due to its complex and fast-moving anatomical structure and large depth from the skin.

In this study, we demonstrated continuous cardiac imaging with qualities comparable to thosefrom a commercial ultrasound probe. The innovation of this application is two-fold.

381

First, traditional stress echocardiography can only image the heart after exercise. Wearable cardiac imaging allows capturing the heart anatomy and performance before, during, and promptly after exercise continuously in real time. This technology enables evaluating cardiac abnormalities that are only present under stress, opening up new diagnostic possibilities with information that previously could not be acquired.

387

388 Second, conventionally, measuring cardiac indices such as stroke volume, cardiac output, and 389 ejection fraction usually requires cardiologists to contour the boundary of the left ventricle 390 manually, which is very time-consuming and operator-dependent. They typically collect only one representative cardiac cycle to evaluate those parameters³, giving only discrete data. Although 391 some research studies^{3,90} and commercial systems⁹¹ developed algorithms to derive these indices 392 393 over multiple cardiac cycles, it still requires cardiologists to operate the imaging probe manually. 394 Long-term continuous cardiac imaging by integrating the wearable ultrasound patch with deep-395 learning based image-processing can derive waveforms of these indices automatically, with an 396 unprecedented high temporal resolution.

397

In summary, the innovations in device engineering and automated image processing allow the wearable ultrasonic patch to perform long-term continuous cardiac imaging from multiple views during exercise. The output waveforms of cardiac indices such as cardiac output, stroke volume, and ejection fraction have never been achieved by existing wearables.

402

403 1.5 Bandwidth improvement of the wearable imager

The 55% transducer bandwidth is not as high as that of the commercial P4-2v imager (~74% bandwidth). In contrast to the rigid ultrasound probe, which has a very thick backing layer to effectively attenuate the reverberation, the wearable ultrasonic patch cannot be assembled with such a thick backing layer because it would severely impair the mechanical compliance of the device, preventing it from intimately conforming and coupling to the human skin. Therefore, the current design of a thin yet dense backing layer is a compromise between reducing reverberation and maintaining outstanding mechanical compliance of the device.

411

To quantify the efficacy of the thin yet dense backing layer in this study, we performed pulseecho testing of an element without the backing layer. The frequency spectrum and its -6 dB bandwidth are derived (Supplementary Fig. 1). The results show that the backing layer in this study improved the bandwidth by almost 25%.
In future work, we will develop new composites for backing layers^{92,93}. Specifically, we will incorporate glass bubbles and tungsten powders into the polyimide resin. By tuning the proportion of the three components, we can synthesize a composite with a small thickness, an appropriate acoustic impedance, and a high attenuation coefficient that is five times higher than the current material⁹³. This new composite will allow the wearable ultrasound probe to have both high bandwidth and skin-like mechanical properties.

Supplementary Discussion 2: Why wearable and why ultrasound?

423

424

- 425
- 426 2.1 Why wearable?

427 Not all diseases have regular and sustained symptoms or pathologies. Continuous monitoring over a long period is essential for providing reliable and accurate diagnosis^{3,4}. The insufficient 428 429 sampling of signals would possibly miss transient but critical signals and thus provide less 430 confidence to the diagnostic results. Thus, long-term monitoring is highly desired for those that 431 are sporadic and barely predictable. However, conventional methods to continuously monitor the 432 heart are invasive or semi-invasive, which are resource intensive and limit their use in bedside surveillance for critically ill or surgery patients^{94,95}. Invasive methods may require anesthesia that 433 could influence the measurement results^{96,97}. Their invasive nature is also associated with risk of 434 435 morbidity and mortality⁹⁴. Emerging wearable technologies address these challenges by enabling 436 long-term monitoring noninvasively.

437

Besides stress echocardiography demonstrated in this work, there are several other use casesthat can benefit from wearable long-term monitoring.

440

441 Sinus arrhythmia produced by the irregular release of impulses from the sinoatrial node is one typical example⁹⁸. Being prevalent among senior people⁹⁹, arrhythmia is the root cause of a variety 442 443 of other symptoms and diseases, such as blood clotting in the heart, hypotension that leads to 444 dizziness, and sudden death. Sinus arrhythmia includes sinus tachycardia, bradycardia, and arrest¹⁰⁰. Sinus arrhythmias may not have symptoms during the limited examination time because 445 it could be sporadic and barely predictable in daily life¹⁰¹. Hence, prolonged tracking time can help 446 447 record the timing and syndromes for a better diagnosis. Also, while it is simple to detect symptoms 448 with a quick test, it is imprudent to instantly diagnose signs as illness and intervene immediately. This is because sinus rhythm is very sensitive to factors such as mood and respiration, and after a 449 while the variations automatically revert to normal⁹⁸. Existing approaches for monitoring 450 451 arrhythmia are plagued by various problems. Auscultation of the heartbeat with a stethoscope is a 452 general but nonspecific measurement, because abnormity like premature or abnormal beats do not 453 always produce an audible pumping and may be missed and misdiagnosed¹⁰². An 454 electrocardiogram could be more effective and accurate. It provides abundant information because 455 different types of arrhythmias have different reflections on electrocardiograms. Doctors can tell

456 the type from an electrocardiogram directly and thus more quickly make a move. But motion 457 artifacts introduced by body movement will distort electrocardiogram signals badly, which challenges signal accuracy in daily monitoring¹⁰³. Additionally, the lack of structural information 458 459 prevents electrocardiograms from providing a comprehensive diagnosis. Photoplethysmography 460 fails to provide sufficient information for further diagnosing the type of arrhythmia¹⁰⁴, because it 461 can only monitor the heart rate. There are some studies on building mathematical models to 462 correlate photoplethysmography waveforms to the arrhythmia type, but the result is not accurate enough¹⁰⁵. Other invasive methods may be less commonly used¹⁰⁶. 463

464

465 Another example is paroxysmal atrial fibrillation, which intermittently occurs as a disorder of cardiac rhythm¹⁰⁷. This disease also suffers from the same problem that limited examination time 466 467 may not be enough for a thorough diagnosis, and this could introduce horrible results. Paroxysmal atrial fibrillation may develop into chronic atrial fibrillation, which occurs more frequently, a direct 468 469 result of misdiagnosis. Moreover, a common error in clinical management of atrial fibrillation is 470 to treat chronic sustained atrial fibrillation and paroxysmal atrial fibrillation similarly, despite some differences in management objectives¹⁰⁷. Atrial fibrillation is often associated with a high 471 472 risk of morbidity and mortality from heart failure, stroke, and thromboembolic complications^{107,108}. 473 Thus, the failure of detecting the early stage of paroxysmal atrial fibrillation is disastrous.

474

475 Similarly, coronary heart disease may also be undetectable in a transient test. Coronary heart 476 disease is caused by plaque built-up near the coronary arteries, which limits the blood supply¹⁰⁹. 477 Coronary heart disease often develops over decades and is frequently ignored because no 478 symptoms are detected. However, the unnoticed coronary heart disease could lead to an acute heart 479 attack due to the blockage of the artery, which is extremely lethal and results in a high mortality¹⁰⁹. 480 This heart attack caused by undetected coronary heart disease with no obvious symptoms may also 481 be called a "silent heart attack". Silent precursors to an imminent myocardial infarction can be 482 detected by observing segmental wall motion abnormalities in ultrasound images. 483

Finally, acute heart disease could barely be noticed when it is not occurring. Myocardial infarction is the lack of blood supply to the myocardium, which weakens cardiac activities¹¹⁰. Myocardial infarction has a high morbidity because many factors and triggers contribute to it, such as alcohol, physical exertion, and obesity. Myocardial infarction also has a high mortality. It cannot be effectively detected in current measuring procedures, because the short period of testing time cannot capture any signs of it unless it is occurring with symptoms.

490

We can summarize that previous efforts of wearable devices for non-invasive heart monitoring are mainly categorized into three fields: electrical probe, electromagnetic probe, and mechanical probe⁵¹. Specifically, the electrocardiogram sensor for local-field potential recording for analyzing cardiac rhythm^{12,111}; the electrical impedance tomography sensor based on internal electrical 495 conductivity mapping mainly for distinguishing systole and diastole phases¹⁶; the electromagnetic 496 sensor that measures stroke volume by relating the resonant frequency response of the sensor to 497 permittivity changes caused by blood volume¹¹²; the passive vibration sensor based on mechanical 498 waves usually designed for monitoring the heart rate¹¹³. However, those signals either are indirect 499 or suffer from low spatial resolutions, from which many clinically important cardiac 490 characteristics, such as the volume of heart chambers, the myocardium structure, and the 501 ventricular ejection function, cannot be visually and accurately evaluated¹¹⁴.

502

503 2.2 General requirements of imaging the heart

504 Echocardiography, as one of the most flexible diagnostic tools revealing the structural information of the heart, is widely adopted in hospitals¹¹⁵. However, current echocardiography 505 506 focuses on short-term recordings of the heart to make a diagnosis, but transient symptoms or 507 dysfunctions may not appear during the limited time of such recordings. Other methods based on 508 imaging can barely provide reliable and thorough monitoring of cardiac functions, including 509 magnetic resonance imaging¹¹⁶, computed tomography¹¹⁷, single-photon emission computed tomography¹¹⁸, positron emission tomography¹¹⁹, and optical coherence tomography¹²⁰. Here we 510 511 briefly outline the pros and cons of each method's working principles and the rationale behind 512 selecting the ultrasound platform in this study (Supplementary Table 1). We desire a device that 513 is wearable, can target the heart, and is capable of 2D or 3D imaging with a high spatial resolution 514 and sufficient contrast. These requirements can be summed up in a single term: wearable heart 515 imaging, which entails the following needs.

516

517 For a modality to be suitable in wearable devices, it must first be able to be packaged into a 518 small, lightweight, and minimal form factor that is both comfortable and non-invasive. The device 519 must ideally provide zero hindrance to the wearer, such that there is a negligible impact on their 520 comfort or normal activities when wearing it. A sufficient penetration depth is required to target 521 the heart non-invasively. Furthermore, the chosen modality must be robust, with no degradation 522 in the image quality over time or large variances in image quality under different conditions 523 encountered in daily life. Wearable devices are often desired to perform long-term monitoring, and 524 to be frequently removed and reapplied to the wearer. Therefore, the device must also withstand 525 the wear and tear of long-term use without a loss in image quality over time. Most importantly, 526 the device must be safe for indefinite use-for example, the modality should not be based on 527 ionizing radiation.

528

For a modality to target the heart, the major challenge to overcome is the high temporal resolution required. The heart may beat at rates of anywhere from 50 beats per minute at rest¹²¹ to nearly 200 beats per minute during exercise¹²², and as such, requires a modality that can achieve temporal resolutions of at least 33 ms^{5,123} to image continuously in real-time. Modalities that lack sufficient temporal resolutions can result in the lack of critical diagnostic information, as well asdegradation of image quality due to motion artifacts.

535

536 Lastly, to make a device useful for performing a wide range of general examinations, the 537 modality must be able to image in 2D or 3D with high spatial resolutions and good contrast. Most 538 of the work on wearable devices has primarily targeted measures such as pulse acquisition and 539 blood oxygen levels. While these are helpful signals, they cannot compare to the immense 540 diagnostic value that direct 2D or 3D imaging of the heart provides. Imaging provides a great 541 wealth of information, but its quality is crucial to being able to make these diagnoses. Poor image 542 quality may often obscure key indicators of diseases, lead to false positives, or make it difficult to 543 distinguish between different conditions.

544

545 Therefore, to summarize, the ideal modality should be robust, safe for long-term use, easily 546 scaled down to wearable and portable form factors, imaging in 2D or 3D with high spatial/temporal 547 resolutions, having a sufficient penetration depth to target the heart non-invasively, having a 548 sufficient contrast, and having a high signal-to-noise ratio. However, it should be noted that some 549 of these requirements have trade-offs. For example, by nature, imaging at high spatial resolutions 550 will tend to sacrifice the temporal resolution, due to the larger amount of data being acquired and 551 processed. Therefore, we can only select the most suitable modality that is able to balance all 552 factors while meeting sufficient requirements.

553

554 2.3 Magnetic resonance imaging

555 Magnetic resonance imaging works by using a powerful magnetic field to align the protons in 556 the body's tissues with the field. Radiofrequency waves are then pulsed to disturb this alignment, 557 followed by a subsequent release of secondary radio waves when the protons realign with the field. These signals are collected to form images¹²⁴. Magnetic resonance imaging provides high tissue 558 contrast and image quality in real-time with spatial and temporal resolutions in the range of 1.6 559 mm and 13-50 ms^{5,125}, respectively. In addition, it is non-ionizing because it uses a magnetic field 560 and radiofrequency waves. However, magnetic resonance imaging has several obvious drawbacks 561 562 that make it unsuitable for needs in this study. First, magnetic resonance imaging machines are 563 extremely bulky and expensive and are not scalable to wearable form factors. Furthermore, the 564 powerful magnetic field is an intrinsic requirement of the modality and can easily present a 565 significant hazard in everyday life in addition to being incompatible with devices like pacemakers 566 that are likely to be used by the target demographics in this study.

567

568 2.4 X-ray computed tomography

A computed tomography scanner consists of an X-ray source and a detector placed opposite of each other, which rotate around the subject to capture several images from multiple angles. The images are then used to reconstruct a 3D image of the subject¹²⁶. Although computed tomography 572 is relatively low cost and provides high spatial resolutions of around 0.3 mm⁶, the scanner contains

573 many moving parts. This type of moving design is not suitable for wearable devices, and the

574 physical constraint puts a strict limitation on the temporal resolution ($\sim 66 \text{ ms}^{6,127}$) that makes it

unsuitable for real-time cardiac imaging. Furthermore, the ionizing X-ray radiation makes it

575

576 fundamentally unsafe for long-term wearing.

- 577
- 578 2.5 Emission tomography

579 Single-photon emission computed tomography and positron emission tomography make use 580 of radiotracers injected into the body that radioactively decay over time as they travel through the 581 body following their designated molecular targets. These signals can then be collected using a gamma camera to quantify those molecular targets and metabolic events¹²⁸⁻¹³⁰. This allows 582 583 gathering unique types of information such as myocardial perfusion and cell metabolic activities¹²⁹. 584 In addition, positron emission tomography's temporal resolution is too slow (>2000 ms¹³¹) for 585 cardiac imaging, while single-photon emission computed tomography's poor image resolution (~10 mm¹³²) makes it largely impractical for more general diagnostic imaging. There is also a very 586 low signal-to-noise ratio in these modalities¹³⁰. Subjects are exposed to low levels of ionizing 587 588 gamma radiation, and the tracers must be replenished over time as they decay, making these 589 modalities unsuitable for wearable devices.

- 590
- 591 2.6 Optical coherence tomography

592 Optical coherence tomography uses light scattering within the 700-900 nm wavelength range, 593 also known as the "therapeutic window", to image the human tissue⁸. These wavelengths have low absorbance and high scattering in tissues on top of being non-ionizing, making them useful for 594 595 imaging tissues. In constructing the image, the time delay cannot be used to determine the location 596 of the scattered signals because the speed of light is too fast. Thus, interferometry is used instead, 597 in which the primary beam of light is split in half to target the tissue and a mirror simultaneously. 598 As the distance of the mirror is varied, constructive interference occurs with signals coming from 599 different locations within the tissue, allowing the position of the signals to be distinguished. Due 600 to the short optical wavelength, optical coherence tomography can achieve extremely high spatial resolutions of up to 1 μ m⁸, and temporal resolutions of 2 ms⁸. In addition, dyes can be injected 601 602 into the subject to add additional capabilities to optical coherence tomography. For example, the 603 use of voltage-reactive dyes can make it possible to form activation and conduction velocity vector 604 maps of the myocardium using optical coherence tomography⁸. Despite these advantages, optical 605 coherence tomography is unsuitable due to the bulky optics required, and limited tissue penetration 606 depth of $2-3 \text{ mm}^8$.

607

608 2.7 Why ultrasound?

Lastly, we look at the rationale for selecting the ultrasound modality in this study.Piezoelectric transducers generate ultrasound waves throughout the tissue. The same transducers

611 can then collect the backscattered signals and construct an image based on their strength and time delay at each pixel^{27,133,134}. This "all-in-one" capability of ultrasound allows the device to become 612 extremely compact and easily scaled down to wearable form factors²⁶. Ultrasound waves have low 613 614 absorption and backscattering and thus can penetrate over decimeters in the integumentary and musculoskeletal systems of the human body. Ultrasound has been able to achieve temporal 615 resolutions of less than 1 ms¹³⁵. The spatial resolution of ultrasound increases with the frequency. 616 617 However, this comes at the cost of penetration depth as higher frequencies are more strongly attenuated by tissues^{134,136}. In general, most conventional diagnostic ultrasound devices use sound 618 waves in the range of 2-18 MHz¹³⁴ and can achieve spatial resolutions in the range of 0.4-2 mm¹³⁷. 619 Additionally, even for long-term use, ultrasound has largely been considered fully safe as long as 620 the power output is kept at the minimum levels needed¹³⁸, below the safety threshold defined by 621 FDA ($I_{SPTA} \le 720 \text{ mW/cm}^2$, $I_{SPPA} \le 190 \text{ W/cm}^2$)¹³⁹. The mechanical energy itself is not toxic to 622 623 the body.

624

625 In the case of arrhythmia, a wearable device with live B-mode and M-mode images is perfect for several reasons. First, heart beats are directly recorded and visualized in echocardiography by 626 627 the wearable device, resulting in a high accuracy in calculating heart rates from B-mode videos or 628 M-mode images. Second, arrhythmia is very unpredictable and may only happen when the heart 629 is under high loads. The wearable device supports uninterrupted long-time monitoring in daily life 630 and can capture any irregularity in heart beats. Third, the abnormal cardiac structure causing 631 arrhythmia can be easily detected in B-mode images, which visualize the root cause of the disease 632 and inform better therapeutic decisions. For example, the gross dilations of the right atrium and 633 right ventricle are considered as an important indicator of arrhythmia¹⁴⁰.

634

The main drawback of ultrasound is that there is inherently a lot of noise in the signal, due to random scatterers in the tissue¹⁴¹. However, significant algorithmic progress has been made in this area to improve the image quality of modern ultrasound scanners. Especially given its versatile functions in clinical applications, ultrasound still remains extremely viable, as well as the best option as a wearable modality for imaging the heart.

640

As of the time of writing, there have been no other studies that have produced, specifically, wearable ultrasound heart imaging devices. A wearable device was made to target the heart¹⁴², but did not have imaging capability. Therefore, the wearable heart imaging capability introduced in this study addressed a critical unmet need.

645

646 2.8 Comparability of the wearable ultrasound to clinical ultrasound

Traditional ultrasound probes require either constant holding and/or repositioning of the probe
each time before taking a measurement. The wearable imager reduces the user dependency down
to a single placement of the probe at the start, after which no operator is necessary because the

patch remains adhered in place. Furthermore, in regions where sonographers are a scarce resource,

- such an advantage could allow a single sonographer to attend to multiple patients at a time, as they
- would not have to physically hold the probe for each patient.
- 653

In spite of these advancements, the wearable imager is not fully immune to user-dependency either. Because ultrasound cannot penetrate bone, an ultrasound probe must always be correctly positioned such that the region of interest is within the imaging window with a clear line of sight unobstructed by bones. This user-dependence can be further minimized by 3D imaging or 2D plane-steering, which is an effective way to eliminate the user-dependency.

659

660 The images from the wearable imager, while comparable to the commercial probes, are not at 661 the same level as those taken by a cardiologist using a clinical machine. The models of the 662 commercial probe and the driving system we used were P4-2v and Vantage 256 from Verasonics 663 company, which is widely known for manufacturing cutting-edge research ultrasound systems. 664 Nonetheless, the Verasonics system performance is still not as good as clinical ultrasound systems 665 manufactured by General Electric or Philips, which excel in designing ultrasound transmitting and 666 receiving modes and customizing image post-processing algorithms for each imaging target. We 667 acquired cardiac images taken by an experienced cardiologist using a clinical machine in the 668 hospital (Supplementary Fig. 32).

669

However, those clinical ultrasound systems have fixed adaptors and algorithms that cannot be adapted to driving the wearable imager in this study. Therefore, we used Verasonics, a programmable research platform, to drive the patch. In Figure 3a, we compared B-mode cardiac images tested by the wearable imager and the commercial P4-2v probe. Those two devices were controlled by the Verasonics Vantage 256 machine with the same transmitting mode and image post-processing algorithms. In this case, we could make a fair performance comparison between the wearable imager and the commercial P4-2v probe.

677

678 The size and cost of the back-end hardware can be readily reduced by replacing it with a 679 printed circuit board used for the commercial portable ultrasound probe. We will work on 680 prototyping a printed circuit board with pulsing, receiving, and data transmitting units and 681 integrating it with the wearable ultrasound imager. The printed circuit board will have a 682 miniaturized footprint. Additionally, the entire system can be powered by a lithium-polymer 683 battery with a voltage of only 3.7 V. The voltage can be amplified up to 200 V by a voltage 684 regulator. We will eventually prototype the printed circuit board in a compact and multilayered 685 structure and make it flexible or even stretchable.

686

Furthermore, the mechanical index, spatial resolution, and contrast-to-noise ratio can be further improved by using single crystal piezoelectric materials, adding matching layer and lens, 689 increasing the number of elements, implementing harmonic imaging, and shrinking the pitch

- 690 further to eliminate the artifacts. The near-field is excessively bright and can be improved by
- suppressing the ring-down with backing layer materials of high attenuation coefficients. The sector
- 692 angle of the sonographic window is limited and can be improved by shrinking the array pitch so 693 that the ultrasound beam can steer to a large angle without generating grating lobe-based artifacts.
- that the ultrasound beam can steer to a large angle without generating grating lobe-based artifacts.New algorithms will be explored to maintain a high frame rate while at an expanded sector angle.
- 695 We could also fabricate a large wearable imager so that it covers multiple positions simultaneously.
- 696 This large array will eliminate the need to repositioning when imaging different sites. To achieve

that, we will have to develop a new wiring strategy to individually address each transducer elementin the large array.

699

700 Supplementary Discussion 3: Configuration of the array

701

702 3.1 The configuration of the array in this study

703 The device is composed of two orthogonal 1D arrays, which can be separated into two parts: 704 four arms and a central area (Extended Data Fig. 1). Each arm consists of 13 elements, with each 705 element containing a row of 6×1 sub-elements. The 6×1 sub-elements are shorted and controlled 706 as one channel. We divided one element into 6×1 sub-elements to considerably improve the 707 stretchability of the device, which could conform more intimately to the curved human skin and 708 provide a better interfacial coupling condition. The central area consists of 36 individually 709 addressable sub-elements, whose activation can be configured through customized algorithms as 710 6×1 elements in either arm of the array. Because the array pitch is less than one wavelength of the 711 ultrasound wave, the device can be considered as two phased arrays with 32 elements in each.

712

713 The 88 channels were physically plugged into the control system, but not all of them were 714 activated all the time. The activating sequence was controlled by the customized algorithm to 715 reconstruct two sectorial images sequentially. Specifically, the 32 elements in one direction of the 716 orthogonal array transmit and receive ultrasound waves first. After a sectorial B-mode image has 717 been reconstructed, the system deactivated this direction and switched to controlling the other 718 direction of the orthogonal array to form the second image. Different from how traditional 2D 719 matrix probes work, this method is limited to only one pair of cross planes (one along each phased 720 array; Extended Data Fig. 1). The 2D matrix probe reconstructs a whole 3D image first, and then 721 extracts any pairs of cross planes from the whole 3D image. In other words, this method captures 722 extra unneeded data, which is more time-consuming, leading to a lower frame rate than the method 723 in this study. Because these standard echocardiographic views are orthogonal to each other and the 724 other cross planes in between are of less interest, our method optimizes the frame rate without 725 losing key data.

However, the disadvantage is that our method cannot do 3D imaging nor capture all cross
planes from a single probe position. To overcome these disadvantages, our future work will focus
on developing a wearable 2D matrix probe that can achieve 3D imaging and thus extract any pairs
of cross planes.

- 731
- 732 3.2 The Mills cross array

The configuration of Mills cross arrays¹⁴³⁻¹⁴⁶ is almost the same as that of our orthogonal array (Supplementary Fig. 34): two linear or phased arrays in orthogonal orientations with the central elements shared by two arrays. The three major differences between them are (1) the Mills cross array was rigid and the orthogonal array is stretchable; (2) the Mills cross array can reconstruct both 2D and 3D images¹⁴⁷, while our device can only conduct imaging in two 2D planes; (3) the manner of ultrasound transmission and receiving are different, as detailed in the following.

739

740 In the Mills cross array, one array serves for transmitting beamforming in the elevational 741 direction and the other for receiving beamforming in the azimuthal direction (Supplementary Fig. 742 34). In this way, a 2D image can be reconstructed with focusing in the elevational direction. 743 However, this method has a major limitation: the effective aperture in the azimuthal plane is very 744 small (determined by the shared center of the cross array). Such a small aperture diminishes the 745 reflected signals that go to the receive array, resulting in a low signal-to-noise ratio, and degrades 746 the lateral resolution in the azimuthal direction, causing low-quality of reconstructed images¹⁴⁵. 747 On the contrary, performing the transmit and receive beamforming using one common phased 748 array will not have this problem due to its long effective aperture in the azimuth plane, yielding 749 2D B-mode images with high qualities in this work.

750

751 3.3 The 1.5D/1.75D array

752 We did not create 1.5D or 1.75D arrays either because of three strong reasons. First, a 1.5D 753 or 1.75D array will need more electrodes. Specifically, the 1.5D and 1.75D arrays would include 104 and 260 additional electrodes compared to the current probe, respectively²⁰. This design 754 755 configuration would require fabricating more than 10 layers of electrodes, which would make the 756 top electrode excessively thick and compromise the device's mechanical compliance. Second, it 757 would significantly lower the imaging frame rate as it would involve an additional beamforming 758 process in the elevational direction. The frame rate is crucial for cardiac imaging because of the 759 highly dynamic cardiac movement. A low frame rate reduces the number of data points in the left 760 ventricular volume waveform, causing aliasing of critical features in the waveform of cardiac 761 cycles, such as diastasis and isovolumetric contraction (Fig. 5b right). The outcomes are inaccurate stroke volume, ejection fraction, and cardiac output. The frame rates for 1.5D or 1.75D arrays are 762 typically <10 Hz¹⁴⁸⁻¹⁵⁰, which could not support high-fidelity waveform extraction from the video. 763 764 Third, it is highly likely that there will be a strong grating lobe in the elevational direction due to 765 the transmitting beamforming implemented by a small number of elements, which can induce

artifacts into the images²⁰. Therefore, this work focuses on achieving cardiac 2D B-mode images
 using 1D phased array.

769 There is no focusing effect in the elevational direction for the orthogonal array in this work, 770 which may diverge ultrasonic energies transmitted to deep tissues. To mitigate this problem, we 771 used an advanced compounding transmission approach and high transducer performance with a 772 large electromechanical coupling coefficient in this work to maximize the transmission and 773 reflection energies in the azimuth and axial directions. This strategy compensates for the energy 774 loss due to the lack of the elevational focusing, resulting in bright and clear cardiac images even 775 at a depth of 14 cm. To further improve the elevational resolution, we will replace the linear arrays with 1.75D or 2D $\operatorname{array}^{20}$ in future studies. 776

778 Supplementary Discussion 4: Possible sources of electromagnetic interference

The noise in the ultrasound signals had such an ultra-wide bandwidth that the most likely source was radio communication signals. The electromagnetic wave frequencies beyond the ultrasound signals (1-5 MHz) in this study could be removed by applying digital filers to the received signals. For electromagnetic wave frequencies within this range, e.g., the amplitudemodulated radio and short-wave radio, they could not be removed by digital filters but could be mitigated by adding a shielding layer. For the latter, we could attribute the noise to the following two possible sources:

- (1) Electromagnetic noise received by the electrodes or transmission lines directly from the ambience¹⁵¹. Without the shielding layer, bare transmission lines and the electrodes can act as an antenna to receive those ambient electromagnetic waves.
- (2) Electromagnetic noise coupled from the human $body^{152}$. We observed an increasing noise 792 793 amplitude when the device got close to the human body, indicating an electrical field 794 approaching the human body. When electromagnetic waves pass by the human body, they 795 will generate magnetically-induced electrical potential in the human body according to the 796 Faraday's law of electromagnetic induction. In this case, the human body can be 797 considered as a circuit that generates an additive electrical field to the ambient 798 electromagnetic waves. Even though the conductivity of various parts of the human body 799 is low compared with metal conductors, some parts of the human body can still form a 800 loop and generate electrical potential. The potentials can be coupled to device signals as noise¹⁵³. 801

803 Supplementary Discussion 5: Measurement variations of imaging metrics

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Taking the measurements of axial resolutions as examples, there were cases where the axial resolution of wearable probe was better than that of commercial probe due to the variation in measurements (position of the probe, rotation of the probe, etc.). Because the surface of the commercial probe is not flat, the degrees of freedom of the commercial probe will be five, as indicated by the red arrows in the Supplementary Fig. 18. The degrees of freedom of the wearable probe will be three, as indicated by the red arrows in the Supplementary Fig. 18. The variations can be introduced in many forms:

- 812
- When the commercial probe is tilted in the side view (Supplementary Fig. 18a Side view),
 or the commercial and wearable probes are rotated in the top view (Supplementary Fig. 18
 Top views), the cross section of the filament is enlarged (from a circle to an ellipse). This
 will cause a longer radiofrequency signal, similar to the effect of reduced bandwidth. A
 longer radiofrequency signal will increase the size of the bright region generated by each
 filament target, causing a worse axial resolution.
- 819 2) When the commercial and wearable probes are moved laterally in the front view 820 (Supplementary Fig. 18 Front views), or the commercial probe is tilted in the front view 821 (Supplementary Fig. 18a Front view) the filament targets will be off-axis from the center. 822 Then the ultrasound beam intersecting the targets will be more diverged compared to the 823 center beam. The diverged beam has a greater tendency to collect more signals from 824 neighboring targets. This will also lead to a seemingly longer radiofrequency signal. As a 825 result, this kind of positioning will also increase the bright region generated by each 826 filament target and hence worsen axial resolution.
- 827 3) When the commercial and wearable probe are moved laterally in the side view 828 (Supplementary Fig. 18 Side views), the probes get closer to the phantom boundary. The 829 closer the probes are to the boundary, the higher the signals from the boundary are. Then 830 the background of the whole image gets brighter. And it is possible that the background 831 plus the noise in the radiofrequency signal will give a background gray value greater than 832 half of that of the deep filament targets (the deeper the target, the dimmer the pixels in the 833 image). This will lead to a higher pixel count, which leads to longer physical distance 834 (Equation 2, Methods), resulting in a worse axial resolution according to the discussion on 835 characterization of the wearable imager in the Methods section.
- 836

To reduce the impact of measurement variations, we repeated the same test five times. The axial resolutions from each test are shown in Supplementary Fig. 18. Even though the wearable imager sometimes shows a better axial resolution, the mean axial resolution of the commercial imager is superior due to its higher bandwidth. The error bars for all measurements in Figs. 2d, 2e and Supplementary Fig. 17 show the standard deviation of the measurements.

842

843 Supplementary Discussion 6: Imaging procedures

- 844
- 845 6.1 Imaging from different views

846 Four standard positions/orientations were used to obtain the best B-mode images of the heart. 847 The first position was on the left side of the sternum, between the second and fourth intercostal 848 spaces. A linear array device was pointing to the right shoulder. From this position, the imager 849 could inspect the parasternal long axis view of the left ventricle. By rotating the device 90 degrees 850 counterclockwise at the same position, with the device pointing to the left shoulder, a parasternal 851 short axial view of the left ventricle could be obtained. The second position was between the fourth 852 and fifth intercostal spaces. With the linear array device pointing to the left shoulder, the four 853 chambers of the heart could be observed from the apex in this view, also known as the apical four-854 chamber view. By rotating the device 90 degrees counterclockwise at the same position and aiming 855 towards the right shoulder, the device revealed the left ventricle, left atrium, and mitral valve, i.e., 856 apical two-chamber view.

857

After the transducer received the response echoes that carry the location and anatomic information of the heart, the echoes were demodulated, decimated, and compressed logarithmically to eventually generate the B-mode image. A graphical user interface for real-time phased array imaging was made up of display windows and control panels for customized settings (Supplementary Fig. 33).

863

In the apical four-chamber view, all four chambers could be seen simultaneously, so that ventricular interdependence and septal wall abnormalities between the chambers (e.g., in cor pulmonale) could be assessed. We could also measure left and right atrial lengths and areas, as well as right ventricular diameter, length, and area.

- 868
- 869

Apical two-chamber view could be used to measure the left atrial length and area.

870

871 In the parasternal long axis view, detectable structures include the left atrium, left ventricle, 872 mitral valve, aortic valve, interventricular septum, right ventricle, and left ventricular outflow 873 tract¹⁵⁴. The unique orientation of this view allows visualizing the full length of the mitral and 874 aortic valve leaflets during their closure and excursion, which makes it especially useful for 875 evaluating valvular functions. Measurements taken in the parasternal long axis view include the 876 interventricular septum end-diastole thickness, the left ventricular internal diameter end-diastole 877 (LVIDd), left ventricular internal diameter end-systole (LVIDs), and left ventricular posterior wall 878 dimensions. Among the measurements, the LVIDd and LVIDs are especially valuable because 879 they are correlated with the left ventricular volume. By increasing the imaging depth, the 880 pericardial and pleural spaces can also be seen from this view.

882 The parasternal short axis view was particularly useful for evaluating the left ventricular wall 883 motion. To do so, we adopted rings divided into a total of 17 segments to assist the mapping of 884 local pathologies. The 17-segment model is a standard tool used in transthoracic echocardiography 885 procedures. In the 17-segment model, the entire left ventricular wall is projected onto a 2D plane, 886 forming a circular projection area made up of multiple concentrically nested rings with varied 887 diameters. Each ring represents the myocardial wall from a different level of the left ventricle, 888 corresponding to the decrease in the diameter of the left ventricular wall from base to apex (viewed 889 from the parasternal short axis). The rings are further divided into a total of 17 segments, each of 890 which receives its blood supply from a different coronary artery. Segmenting the left ventricular 891 wall in this way provides a useful localization mapping that allows abnormal myocardial strain in 892 any given segment, due to cardiac malfunctions such as ischemic heart diseases or myocardial 893 infarction, to be traced back to its corresponding coronary artery of origin.

894

895 In the short-axis plane, the circular cross-section of the left ventricle was captured (the basal, 896 mid-cavity, and apical views/slices), and the contractility and distensibility were accessible as the 897 motion of the walls was along the plane. The basal slice captures the ring with the largest diameter 898 (segments 1-6). The mid-cavity slice captures a smaller diameter ring nested inside the basal view 899 (segments 7-12). Likewise, the apical slice is nested inside the mid-cavity slice (segments 13-16). 900 Segment 17 is the apex. Therefore, the relative changes in the cross-sectional diameter, along with 901 the uniformity of wall motions, could be easily assessed. Using these assessments, we can identify 902 the specific segment of the left ventricular wall as pathological.

903

904 To quantitatively evaluate the segment displacement, we set a target area for each segment to 905 facilitate further processing. We blurred the original images in the area to reduce the impact of 906 speckles on the feature refinement, and then computed the edge information using a Canny 907 operator based on the blurred image. By indexing the edge in the binary map, the myocardium 908 displacement could be recorded. With these measurements, we could monitor the potential risk 909 factors for myocardial infarction or akinesia, precisely localize ischemic heart diseases, and easily 910 assign hypokinetic or akinetic regions of the left ventricular wall to their governing coronary arteries by tracking the relevant myocardium displacement¹⁵⁵. 911

- 912
- 913 6.2 Phase correction on nonplanar surfaces

Applying phase correction allows the wearable imager to cover a more diverse population, including obese people. Also, the phase correction is critical for providing accurate cardiac monitoring. Because we used a wide-beam compounding transmission in this study, the delay calculation for the aperture could vary a lot from traditional plane-wave compounding transmission^{27,156}. As an intrinsic feature of wide-beam transmission, the focal point of the aperture was set at the opposite side to the imaging area, with the distance between the focal point and the center of the aperture kept constant during the transmission (Supplementary Fig. 13). 921

922 Assuming that a linear array of the wearable imager was attached to a planar surface, the 923 transmission and time delay could be calculated following the general approach. Considering the center of the linear array located at (0,0) by default, we can define the distance between the focal 924 point and the center of the aperture as $df = \sqrt{x_{focal}^2 + z_{focal}^2}$, in which $z_{focal} < 0$. Also, the 925 location of the i^{th} transducer could be defined as $(x_i, 0)$, where $x_i = (i - 16.5) \times pitch$ 926 927 because the device had 32 transducers in each imaging plane. Additionally, we defined the aperture size as A and the angle departure from the vertical direction as θ , where $\theta > 0$ when $x_{focal} >$ 928 0. For a given position of a pixel (x_p, z_p) , the time of the wavefront to go to the pixel since the 929 930 earliest trigger on the transducers was:

932
$$t_{\theta,p}^{e} \approx \frac{\sqrt{(x_p - df \times \sin \theta)^2 + (z_p - df \times \cos \theta)^2 - dt}}{c}$$
(13)

933

934 where
$$dt = df \times \cos \theta$$
 when $|df \times \sin \theta| < \frac{A}{2}$ and $dt =$
935 $\sqrt{(df \times \sin \theta - sign(\theta) \times \frac{A}{2})^2 + (df \times \cos \theta)^2}$ when $|df \times \sin \theta| \ge \frac{A}{2}$. And the time for the
936 reflected wave to get back to the i^{th} transducer was:

937

938
$$t_{i,p}^{r} = \frac{\sqrt{(z_p - z_i)^2 + (x_p - x_i)^2}}{c} (14)$$
939

940 The total time delay since the trigger should be

941

942
$$t_{i,\theta,p}^{dt} = t_{\theta,p}^e + t_{i,p}^r$$
(15)

943

where *c* was the speed of sound that we assumed to be constant in the medium. Considering the delay differences among transducers, the delay of trigger of i^{th} transducer was:

946

947
$$t_{i,\theta}^{t} \approx \frac{\sqrt{(df \times \sin \theta - x_i)^2 + (df \times \cos \theta - z_i)^2} - dt}{c}$$
(16)

948

949 As a result, the delay of the i^{th} transducer for beamforming at a given pixel was:

951
$$t_{i,\theta,p}^{total} = t_{i,\theta,p}^{dt} - t_{i,\theta}^{t}$$
(17)

since each channel started to receive data after transmission. When the device was placed on a nonplanar surface, the time delay formula above was no longer valid (Supplementary Fig. 15). Assume the curvature radius of the nonplanar surface was r. The previous location the i^{th} transducer would then move to (x'_i, z'_i) , and $x'_{i} = sign(x_{i}) \times r \times sin \varphi$ (18) $z'_i = r \cdot (1 - \cos \varphi) \ (19)$ where $\varphi = \frac{x_i}{r}$. With the phase correction, the time of the wavefront to go to the pixel since the earliest trigger on the transducers would be changed to $t_{\theta,p}^{\prime e} \approx \frac{\sqrt{(x_p - df \times \sin \theta)^2 + (z_p - df \times \cos \theta)^2 - dt'}}{c}$ (20) $dt' = \sqrt{df^2 + r^2 + 2 \times df \times r \times \cos \theta} - r$ when $|\varphi| < \frac{A}{2r}$ and dt =where $\sqrt{(df \times sin\theta - sign(\varphi) \times |x_1'|)^2 + (df \times cos\theta - ssign(\varphi) \times |z_1'|)^2}$ when $|\varphi| \ge \frac{A}{2\pi}$. Meanwhile, the time for reflected waves to get back to the i^{th} transducer was changed to $t_{i,p}^{\prime r} = \frac{\sqrt{(z_p - z_i^{\prime})^2 + (x_p - x_i^{\prime})^2}}{c}$ (21) and the delay of triggering the i^{th} transducer was changed to $t_{i,\theta}^{\prime t} \approx \frac{\sqrt{(df \times \sin \theta - x_i)^2 + (df \times \cos \theta - z_i)^2} - dt'}{c}$ (22) Finally, the delay of the i^{th} transducer for beamforming at a given pixel after phase correction became:

983
$$t_{i,\theta,p}^{\prime total} = t_{\theta,p}^{\prime e} + t_{i,p}^{\prime r} - t_{i,\theta}^{\prime t}$$
(23)

984

985 In the receive beamforming, the value of each pixel could be computed as:

986

$$I_p = 20 \log_{10} \sum_{\theta=\theta_{min}}^{\theta=\theta_{max}} \sum_{i=1}^{\# of \ channel} RF_i(t_{i,\theta,p}^{\prime total} \times f_s)$$
(24)

988

991

989 where RF_i is the radiofrequency signal collected for the i^{th} transducer and f_s is the sampling 990 frequency. Any value larger than 255 would be cut off to adapt to the 8-bit display.

992 In Extended Data Fig. 2, for the images labeled with "Planar", we fixed the probe on a planar 993 glass slide and used ultrasound couplant to compensate the gap between the probe and the skin. 994 The images labeled with "Curved" were obtained with the probe naturally laminated on the curved 995 skin surface. Exemplary images of the parasternal long axis view were collected on an obese 996 subject, whose curvature radius at the imaging site was ~82 mm. The size and shape of the left 997 ventricle were obviously distorted without the phase correction. The measured values of the stroke 998 volume, cardiac output, and ejection fraction tend to be smaller pre-phase correction than the 999 results post-phase correction, due to the distorted size and shape of the left ventricle.

1000

1001 6.3 Validation for long-term use

Motion artifacts plague ultrasound imaging¹⁵⁷⁻¹⁶⁰. To verify the performance of the wearable imager under daily circumstances, we attached the device to subject's chest and tested the imaging quality under different postures. In the experiments, there was no obvious deterioration of imaging quality, due to the intimate contact of the device to human skin (Extended Data Fig. 4), attesting the stable performance of the device.

1007

We compared the imaging results along the four standard views with ultrasound gel and silicone as the couplant (Supplementary Fig. 20). The results showed that the contrast between different structures and details in the heart anatomy were well conserved when the couplant was changed from ultrasound gel to silicone. We recycled the device and repeated the tests on ten subjects and obtained reliable and reproducible results (Supplementary Fig. 24).

1013

In addition, the biocompatibility of the device was also a potential concern for long-term use, which may adversely irritate the skin. We attached the device to the subject's chest with a commercial adhesive (e.g., Tegaderm) for 24 hours (Supplementary Fig. 22), during which the skin experienced all kinds of scenarios, including exercising and showering. After removing the device, no allergy was observed. The major issue was the reddish area around the device attachment location caused by peeling off the Tegaderm adhesive.

1020

1021 6.4 Simultaneous measurements of M-mode images and electrocardiogram

At the beginning of the test, the wearable imager was attached to the parasternal site to image along the parasternal short axis view. We triggered the recording of B-mode images and electrocardiogram (by a commercial device, Cyton Biosensing Board 8 channels) simultaneously. With the concurrent ending of the recording, the time (i.e., frame number in this case) of B-mode images and electrocardiogram could be aligned. Furthermore, we plotted the M-mode image by selecting pixels in one line in the B-mode images and correlating this array of pixels to the frame number.

1029

1031

1030 Supplementary Discussion 7: Stress echocardiography

1032 7.1 Significance

1033 Under a steady state of a healthy subject, the left ventricular volume changes consistently, so 1034 the stroke volumes, cardiac outputs, and ejection fractions (three of the most significant indicators of cardiac functions) do not vary significantly¹⁶¹. However, the heart state may change frequently 1035 1036 and could be extremely intricate at times. Abnormal fluctuations exist in estimating the cardiac 1037 function of patients with heart diseases, and the symptoms could be very unpredictable, which also 1038 incur dramatic changes in the heart in a relatively short time. Symptoms of cardiovascular diseases 1039 like myocardial ischemia only emerge under stress, where the induced wall motion abnormalities 1040 are more noticeable¹⁶². Therefore, stress echocardiography is carried out. Valvular pathologies are easily observed by individually tailored stress echocardiography¹⁶³. In addition, stress 1041 echocardiography is valuable for studying pulmonary circulation¹⁶⁴. Stress echocardiography can 1042 also help determine limits for safe exercise before starting a cardiac rehabilitation program or 1043 1044 recovering from a cardiac event, and sometimes help evaluate the cardiac status before heart 1045 surgery.

1046

1047 7.2 Limitations of existing procedures

1048 The process of stress echocardiography sometimes suffers from inter-observer variabilities. 1049 Current evaluation approach requires manually tracing the contour of the left ventricle in the 1050 echocardiographic images and calculating its end-systolic and end-diastolic volumes for further 1051 analysis. Assessments are carried out typically based on only one cycle of heartbeat, even if the 1052 standard guidelines recommend tracking and averaging five cycles before making a 1053 conclusion^{165,166}. It potentially brings huge variance and uncertain accuracy to diagnostic results, 1054 especially for those less-trained medical assistants in low-resource regions.

1055

1056 Stress echocardiography also requires extremely sophisticated procedures to find an 1057 appropriate imaging location/orientation in a short time⁴⁰. The acquired views must be of the same 1058 regions of the myocardium before and after exercise⁴⁰. Same regions in the ultrasonographic 1059 window are required to make sure the images are from the same or a very similar 1060 location/orientation of the heart. Otherwise, the measurements will be incorrect, because the1061 chamber volume varies in images from different locations/orientations.

1062

In addition, the end-point when the patient should stop exercising is dependent on whether patients achieve the target heart rate, experience moderate angina/severe chest pain, or reach tolerable exercise limits. These subjective criteria may result in suboptimal cardiac testing outcomes.

1067

Furthermore, patients may be vulnerable to certain diseases that show up only during exercise, which is not monitored by existing procedures. For example, myocardial ischemia does not show in the resting state but appears only in stress, resulting in hypokinesia, akinesia, and dyskinesia. These symptoms are usually mild but can develop to be acute and lethal if neglected. However, because the patient is not monitored during exercise, the initiation of myocardial ischemia is unknown, not only missing valuable data but also leaving the patient at risk.

1074

1075 7.3 Echocardiography by the wearable imager

1076 Those limitations of existing echocardiography can potentially be addressed with the wearable 1077 imager demonstrated in this study. The aperture of the wearable imager in this work is relatively 1078 small. The entire array length is 12.7 mm, which is shorter than the intercostal space (~14.5 to 19.7 1079 mm¹⁶⁷). Therefore, imaging is not challenged by the ribs because we only image from one position 1080 at a time, rather than using a big patch to cover many locations simultaneously. We imaged in the 1081 parasternal axis views and apical views at different locations (Extended Data Fig. 3). In the 1082 parasternal long axis view, the wearable imager was attached to the chest of the subject 1083 continuously during the entire process. The heart rate was estimated to be average 70 beats per 1084 minute for the rest stage, which changes occasionally though, but still within a normal range. Then, 1085 the subject exercised on a cycling bike while the device continuously monitored the heart status. 1086 As the exercise began, the heart rate gradually rose. The subject exercised as hard as possible to 1087 reach the possible highest heart rate. The whole exercise duration took around 12 mins. After 1088 exercise, the heart rate slowed down back to normal.

1089

1090 The LVIDs and LVIDd were measured during different stages. Before exercise, the average 1091 LVIDd and LVIDs were 45 mm and 27 mm, respectively. During exercise, growing need of blood 1092 supply accelerated the heart beats, but the LVIDd and LVIDs were slightly dropping because a 1093 shorter pumping period allowed fewer muscle extensions. At the peak of exercise, LVIDd roughly 1094 dropped to 38 mm and LVIDs went down to 22 mm. After exercise, the LVIDd and LVIDs 1095 recovered to approximately 41 mm and 24 mm, respectively. Fractional shortening, the reduction 1096 of end-diastolic diameter during systole, is a measure of the cardiac muscular contractility. It was 1097 calculated as the difference between the LVIDd and LVIDs divided by the LVIDd.

1099 The parasternal long axis view contains information mostly about the left ventricle and atrium, 1100 while the apical four-chamber view provides a more comprehensive window of all four chambers 1101 and is more precise for estimating the left ventricular volume. Because the apical four-chamber 1102 view requires the patient to be tilted in the lying side position so ultrasound can enter from the 1103 apex, we could not collect these data during exercise. In the apical four-chamber view, we could 1104 see that both end-systolic volume and end-diastolic volume were increasing after exercise. This process of heart restoration is sometimes defined as heart volume reversal¹⁶⁸. Reasonably, exercise 1105 leads to increases in both contractility and afterload of chambers, resulting in a physiological 1106 1107 decrease in chamber volumes immediately.

1108

1109 Changes in ventricular size under stress may provide useful information regarding cardiac 1110 functions. For example, the end-systolic volume reversal may provide complementary information 1111 for risk stratification of cardiac diseases¹⁶⁸. A decreasing end-diastolic volume may be a critical 1112 indicator of hypovolemia resulting from poor oral intake, emesis, or myocardial loss during cancer 1113 treatment¹⁶⁹. An unsteady end-systolic volume recovery also portends the possibility of diseases 1114 like septal defects¹⁷⁰ or valvular regurgitation¹⁷¹.

- 1115
- 1116 Supplementary Discussion 8: Acoustic coupling of wearable imager
- 1117

In this study, the 50-micron layer of silicone underneath the device is in a liquidous state (Silbione part A, A-4717, Factor II Inc.). It was not used to encapsulate the array but instead as an acoustic couplant and adhesive layer. The silicone layer is a structural material that improves the acoustic coupling and adhesive properties of the device without limiting its flexibility.

1122

Rigid probes cannot provide meaningful B-mode images without couplants due to incompatible mechanical properties between the rigid probe and the soft human skin (Supplementary Fig. 21d left panel). The stretchable ultrasound patch is packaged by skin-like soft polymers, i.e., SEBS in this case. The mechanical compliance of the patch allows intimate conforming to the human skin without external holding, which reduces interfacial air gaps and enhances acoustic coupling between the transducer and human skin. Therefore, the patch can image the human body even without coupling materials.

1130

With that being said, microsized air bubbles may still exist at the wearable patch/skin interface. To achieve the best acoustic coupling, we added uncured Silbione (part A, A-4717, Factor II Inc.), a thin layer of liquid silicone whose acoustic impedance (1.03 MRayl)¹⁷² is very close to the human skin, as the couplant underneath the device. The couplant can eliminate microsized air bubbles and enhance the coupling between the device and the skin. In addition, traditional hydrophilic ultrasonic gels are volatile and cannot last for long periods of time. The silicone couplant does not evaporate at room temperature. Therefore, it can provide a good acoustic coupling over the longterm.

1139

To illustrate the performance of the silicone couplant, we compared the image quality of the wearable patch on the human body with and without the silicone layer (Supplementary Fig. 21d). The cardiac anatomic structures are clearly displayed. The image with silicone couplant is brighter than that without, indicating a stronger signal-to-noise ratio due to better interfacial acoustic coupling.

1145

1146 To quantify the coupling performance of the liquidous silicone, we have used a single 1147 transducer to sense a specific target. Specifically, we excited the transducer with a thin layer of 1148 liquidous silicone as couplant, and then acquired the reflected signals. For comparison, we also 1149 did the same test using commercial ultrasound gel as couplant. We did the experiments on a 1150 commercial phantom (Aquaflex, Parker Laboratories) with a reflector 18 mm underneath the 1151 surface. The results show a comparable signal-to-noise ratio between commercial ultrasonic gel 1152 and liquidous silicone (Supplementary Fig. 21), indicating the outstanding acoustic coupling 1153 performance of the silicone.

1154

In addition, we also imaged the heart from four different views with commercial ultrasound gel as the couplant. The resulting images were compared with Supplementary Fig. 20 generated with liquidous silicone as the couplant. Images from all views show that the contrast between different structures and details in the cardiac anatomy are highly similar, and no obvious structural differences are found. These results show the practicality and excellent performance of liquidous silicone-based couplant.

1161

1162 The acoustic attenuation of the silicone is negligible. The attenuation depends on the 1163 ultrasound frequency and the material thickness. Given the attenuation coefficient of ~0.11 dB/mm 1164 for liquidous silicone at 3 MHz¹⁷³, a 50 μ m thick couplant only causes 0.005 dB attenuation, which 1165 has practically no negative effect on the acoustic properties of the wearable ultrasonic array.

1166

Furthermore, the liquidous silicone has a high viscosity (60 Pa·s)¹⁷⁴, which makes the device less susceptible to sliding on the skin, ensuring a consistent imaging window on the same target during long-term continuous monitoring.

1170

1171 Supplementary Discussion 9: Continuous cardiac performance monitoring

1172 9.1 Monitoring ejection fraction, cardiac output, and stroke volume simultaneously

1173 Commonly monitored vital signs contain body temperature, respiration rate, peripheral blood 1174 pressure, and heart rate. The first three parameters cannot directly reflect the heart status, while 1175 the heart rate only tells how quickly the heart is pumping but does not reveal the actual performance of the heart. Other parameters, like ejection fraction, cardiac output, and strokevolume should be explored for a more comprehensive and conclusive diagnosis.

1178

1179 Ejection fraction represents the fraction of the blood ejected from the left ventricle per cycle. 1180 It is an indicator of the left ventricle's overall systolic performance (Equation 23). Cardiovascular diseases lurk in an abnormal fluctuation of ejection fraction. Normal left ventricular ejection 1181 fraction should be >50%, and a reduced one may manifest itself as heart failure¹⁷⁵: a moderately 1182 1183 reduced ejection fraction is within 40-49%, and a reduced one is <40%. Chronically attenuated 1184 ejection fraction is undoubtedly a danger sign, and an unreasonably high ejection fraction also entails troubles like hypertrophic cardiomyopathy¹⁷⁶, a common cause of sudden cardiac arrest. 1185 As one of the most clinically significant indices of cardiac function, ejection fraction is key in 1186 1187 differentiating systolic versus diastolic heart failure, and is well correlated with mortality in stable outpatients with coronary artery disease and heart failure^{177,178}. 1188

1189

1190 Cardiac output is a volumetric blood flow rate and is an indicator of the tissue oxygenation (Equation 24). The cardiac output, the product of the heart rate and the stroke volume, the blood 1191 volume pumped from the left ventricle every minute¹⁷⁹, can aid in the diagnosis of heart failure 1192 conditions, monitor patient status during surgeries and intensive care, and evaluate the overall 1193 cardiac functions¹⁸⁰. The cardiac output is widely monitored in anesthesiology and emergency 1194 care¹⁸¹. Measurement of cardiac output is specifically essential in unstable patients whose 1195 1196 condition may undergo dramatic changes in a short time, as it indicates an overall systemic oxygen delivery and tissue perfusion¹⁸². Many pathologies besides cardiovascular diseases lead to changes 1197 in cardiac output. Abnormally decreased cardiac output could be a sign of heart failure caused by 1198 valvular heart diseases¹⁷⁹, but also could be intoxications like acute azotemia¹⁸³, indicating a severe 1199 dysfunction of the kidney. Aberrant high cardiac output may be a complication of sepsis¹⁸⁴, 1200 hyperthyroidism¹⁸⁵, or anemia¹⁸⁶. When evaluating the cardiac output, oftentimes a patient may 1201 appear asymptomatic during resting conditions due to a wide range of physiological regulatory 1202 processes such as vasodilation and minor increases of heart rate within the physiological range. 1203 1204 Therefore, maximal cardiac output measurements during exercise are also of particular interest for their ability to reveal underlying problems in otherwise normally asymptomatic subjects. 1205

1206

Stroke volume is the difference between the EDV and the ESV, representing the absolute blood volume ejected from the left ventricle in a single cycle. Stroke volume, when used together with cardiac output and/or ejection fraction, can provide a much more comprehensive overview of the cardiac status. For example, as mentioned above, cardiac output can often appear asymptomatic. In those cases, the presence of a lowered stroke volume may reveal underlying heart failure. In other cases, a patient might appear asymptomatic if only considering the ejection fraction. Then, the presence of a decreased stroke volume would be able to indicate diastolic heart failure.

1215 There are several other examples demonstrating why monitoring any one of these parameters 1216 in isolation may lead to inaccurate diagnosis results. For example, ejection fraction may overstate 1217 cardiac functions in left ventricular hypertrophy, which can lead to heart failure. In this case, ejection fraction remains normal, but stroke volume and cardiac output are dropping¹⁸⁷⁻¹⁸⁹, which 1218 is also well known as heart failure with preserved ejection fraction. Diagnosis only based on 1219 1220 ejection fraction would be wrong, while a comprehensive analysis on ejection fraction combined 1221 with cardiac output and stroke volume can generate correct results. Conversely, diastolic and systolic dysfunctions of the left chambers sharply reduce ejection fraction¹⁹⁰, but the cardiac output 1222 1223 can maintain in the normal range with compensation of increased heart rate. Besides cardiac 1224 diseases, some surgical procedures may also have impacts on cardiac functions that require 1225 monitoring these three indices simultaneously.

1226

1227 This scenario is also paralleled in the measurement of blood pressure, which is commonly 1228 done in current practice. Blood pressure reflects a composition of multiple contributing factors, 1229 for example, cardiac pre-load and vascular resistance. Much like the scenarios presented by stroke volume, ejection fraction, and cardiac output, a normal blood pressure reading could obscure 1230 1231 underlying abnormalities in the cardiac pre-load, and vascular resistance if measured in isolation. 1232 Measurement of the left ventricular volume can serve as an indicator for these factors contributing 1233 to blood pressure. The cardiac pre-load is how much the myocardium is stretched prior to contraction, and is reflected in the EDV¹⁹¹. Vascular resistance can be estimated from cardiac 1234 output using lumped parameter models of the circulatory system^{192,193}. The ability to monitor the 1235 volume of the left ventricle can thus provide insight into the contributing factors to blood pressure 1236 1237 and reveal diseases that might otherwise be obscured.

1238

1239 Therefore, ejection fraction, cardiac output, and stroke volume are important parameters for 1240 evaluating cardiac performance¹⁹⁴, and together provide a comprehensive analysis of the blood 1241 delivery capabilities of the heart. Continuously monitoring these indices of the heart for the long 1242 term is of strong prognostic value and has great potential to decrease the mortality and morbidity 1243 of many cardiovascular diseases and conditions.

1244

1245 9.2 Monitoring the left ventricular volume by 3D imaging

1246 From Equations 22 to 24, the ejection fraction, cardiac output, and stroke volume are directly 1247 related to the volume of the left ventricle (EDV and ESV). Thus, the most direct method to measure 1248 these indices would be to monitor the left ventricular volume. Ideally, the most accurate and direct 1249 approach would be to capture 3D images of the left ventricular chamber throughout the cardiac 1250 cycle and use those 3D images to calculate the volume. Common methods of capturing these 3D 1251 images include traditional ultrasound, computed tomography, radionuclide imaging, and magnetic 1252 resonance imaging. However, these common cardiac imaging techniques have a host of limitations, 1253 including device bulkiness, low temporal resolutions, and long-term toxicity to the body.

Furthermore, in many cases, it is desirable to continuously monitor cardiac output in the operating theater to prevent complications during surgery such as shock¹⁹⁵, which is not viable for the aforementioned techniques given these drawbacks.

1257

1258 In addition, the calculation based on manual image labelling suffers from interobserver 1259 variability. Specifically, when labelling the left ventricular dimensions, the endocardial border 1260 requires to be traced continuously from one side of the mitral annulus to the other side. However, 1261 because the endocardial is hypoechoic, no sharp boundary can be easily seen in the image, yielding 1262 large discrepancies of left ventricular dimensions between clinicians' observations⁴². The 1263 discrepancies can be further magnified in calculating stroke volume, cardiac output, and ejection 1264 fraction, which results in an inaccurate diagnosis and introduces uncertainties in subsequent 1265 treatment regimens. In addition, less severe abnormality suffers from greater interobserver 1266 variability, where early symptoms will easily slip away if not well measured by two or more 1267 experienced echocardiographers.

1268

1269 9.3 Monitoring the left ventricular volume by 2D imaging

1270 The next most direct method of obtaining the left ventricular volume would be to approximate 1271 the 3D volume of the left ventricle through 2D imaging. This approach faces similar problems due 1272 to the use of imaging but is computationally faster and can achieve higher temporal resolutions 1273 than 3D imaging.

1274

1275 While 2D imaging with ultrasound is a widely used approach for its convenience, traditional 1276 ultrasound has a bulky housing and requires the probe to be manually held in place. Point-of-care ultrasound probes transcend the limitations of traditional medical imaging¹¹ and promote precision 1277 1278 medicine for household use, but they still require an external force to maintain a stable coupling 1279 with the skin. Otherwise, the image window will change and generate unfair comparisons, which results in misdiagnoses. In addition, their use highly depends on the clinicians' experience, which 1280 is prone to generating inter-observer variabilities and operational errors¹⁹⁶. Robotic arms have been 1281 1282 applied in such a case, but the higher cost and the even more bulky design are the new problems¹⁹⁷, 1283 making it inaccessible in most cases and impractical for continuous and long-term measurements. 1284

1285 9.4 Monitoring the left ventricular volume by model estimations

1286 Therefore, the traditional approaches to continuously monitoring stroke volume, cardiac 1287 output, and ejection fraction do not tend to employ imaging, but instead use models to estimate 1288 these parameters from other indirect measurements. Here, we provide a review of some of the 1289 major relevant non-imaging methods.

1290 1291

(a) Fick's method

According to the Fick principle, the ratio between the rate at which oxygen is absorbed into the blood and the rate at which blood is delivered through the body, is directly represented by the difference in blood oxygen content between mixed venous blood and arterial blood. This principle is summarized by¹⁸⁰:

 $CO = \frac{VO_2}{C_a - C_v}$ (25)

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- 1297
- 1298

where *CO* is cardiac output, VO_2 is the amount of pure gaseous oxygen consumed per unit time, and C_a and C_v are oxygen content of arterial and mixed venous blood, respectively. Typically, *VO*₂ is measured using a spirometer, while C_a and C_v require catheters to be inserted into the patient. C_v is measured from the pulmonary artery or vena cava, while C_a is often measured from the brachial or femoral artery¹⁸⁰.

1304

1305 While Fick's method can be highly accurate, in the range of 5 to $10\%^{180}$, and is often used as 1306 a benchmark for other methods, it is highly invasive and requires catheterization. Furthermore, to 1307 take accurate readings, the patient's cardiac output and oxygen consumption must be stable for 1308 several minutes at a time¹⁸⁰. It is also inconvenient for infants, or during surgery, because a 1309 sufficient blood volume is required for this technique¹⁸⁰.

1310

1311 (b) Indicator dilution techniques

Indicator dilution is a variation of the Fick principle and works in a similar way. Rather than
using oxygen as an indicator, other indicators can be injected into the bloodstream at a single point,
and their concentration is analyzed downstream. The most common indicators used in dilution are
indocyanine green dye (dye dilution), or cold saline (thermodilution).

1316

In the case of dye dilution, the dye concentration is measured using a densitometer, based on the optical density of the blood. This is then plotted over time, and the area under the curve can be related to the cardiac output and the amount of dye initially injected using the Stewart-Hamilton equation¹⁹⁸:

1321

1322
$$CO = \frac{V_{indicator}}{\int_0^\infty C(t)dt}$$
(26)

1323

1324 where the numerator is the initial amount of indicator injected, and C(t) is the concentration of 1325 the indicator measured downstream over time.

In reality, because the dye is not removed from the bloodstream by the kidneys sufficiently
 fast, the concentration curve peaks multiple times as the dye recirculates¹⁸⁰. Therefore, the curve
 must be extrapolated so that it returns to zero concentration.

- For thermodilution, the Stewart-Hamilton equation is modified to account for the heat transfer
 between blood and saline^{180,198}:
- 1333

1334

1330

$$CO = \frac{K * V_{indicator} * (T_{blood} - T_{indicator})}{\int_0^\infty T_{measured}(t)dt}$$
(27)

- 1335
- 1336 $K = \frac{c_{indicator}\rho_{indicator}}{c_{blood}\rho_{blood}} (28)$
- 1337

1341

1338 where c is the specific heat, ρ is the density, $V_{indicator}$ is the volume of indicator initially 1339 injected, T_{blood} and $T_{indicator}$ are the initial temperatures of the blood and indicator, respectively, 1340 and $T_{measured}(t)$ is the temperature measured downstream over time.

1342 The injected saline lowers the temperature of the blood as it travels through the bloodstream, 1343 followed by warming as it is mixed and diluted in the blood. This is measured using a thermistor. 1344

Indicator dilution is unsuitable for continuous monitoring as the temporal resolution is too low.
One must wait for roughly 1 minute for the indicator to circulate and fully dilute¹⁸⁰. In addition,
the densitometer requires a lengthy calibration process¹⁸⁰. Unlike the Fick method, there is no selfsustaining constant flux of indicator, so periodic reinjection is required. Over time, this can
increase the risk of embolism, infection, and fluid imbalances¹⁸⁰.

1350

1351 These methods can also be error-prone. For example, thermodilution techniques will have limited accuracy given unstable body temperature, but this may be uncontrollable in certain cases, 1352 like liver transplantation surgery¹⁹⁹. In addition, arterial pulse waves could also be transferred to 1353 1354 cardiac measurements, but this is also unreliable for monitoring on trends because it cannot 1355 compensate for circulatory changes. Also, the Stewart-Hamilton model itself assumes ideal fluid 1356 flow. Many of the assumptions of ideal fluid flow are violated by the human circulatory system, such as single inflow and single outflow tracts, complete mixing of the fluids, steady flow, and no 1357 recirculation of indicator¹⁹⁸. 1358

1359

Additionally, in dye dilution, the downslope of the concentration curve must be extrapolated extensively due to recirculation of the dye, limiting the accuracy. In thermodilution, one must carefully maintain the temperature of the injectate, or else it may cause the temperatures to deviate. Thermodilution is also unreliable in low cardiac output situations, where it tends to overestimate the cardiac output. Under low-flow conditions, the indicator has more time to equilibrate with the surrounding tissues, leading to diminished measured temperature changes and a smaller area under the curve²⁰⁰, because heat exchange is assumed to only occur between blood and saline.

1367 1368

(c) Conductance catheterization

1369 In conductance catheter measurements, a catheter with multiple electrodes segmented along 1370 its length is inserted into the left ventricular chamber. The blood within the chamber is segmented 1371 into different volumes stacked together, with their boundaries defined by the left ventricular wall and the equipotential surfaces through the electrodes²⁰¹. Using the dielectric and conductive 1372 properties of the blood, each separate volume of blood can then be treated as a resistor and 1373 capacitor in parallel²⁰¹. The height is the distance between electrodes. The cross-sectional area is 1374 the cross-sectional area of the blood inside the left ventricle²⁰¹. This cross-section varies over time 1375 throughout the cardiac cycle. The conductance, 1/R, can be modeled by²⁰¹: 1376

1377

1378
$$\frac{1}{R(t)} = \frac{\sigma V(t)}{L^2}$$
(29)

1379

1380 where σ is the conductance of the blood, *V* is the volume of the segment, and L is the length 1381 between electrodes. From there, the volumetric contribution of each segment to the stroke volume 1382 can be found²⁰¹:

 $\Delta V = \frac{L^2}{\sigma} \left(\frac{1}{R_{he}} - \frac{1}{R_{ee}} \right)$ (30)

- 1383
- 1384
- 1385

1387

1386 where R_{be} and R_{ee} are the resistances at the beginning and end of ejection, respectively.

1388The summation of each segment then gives the total stroke volume, and from there, ejection1389fraction and cardiac output may also be determined.

1390

This method has the disadvantage of being highly invasive, requiring catheterization of the left ventricle. The conductivity of blood must also be calibrated. This can lead to errors, as the conductivity of blood changes throughout the cardiac cycle (see the Bioimpedance section). The model also assumes the equipotential surfaces to be parallel and the blood in the ventricle to be the only conducting objects, which may deviate from the real situation²⁰¹.

- 1396
- 1397 (d) Radionuclide angiography

1398 Traditionally, gamma cameras used in radionuclide angiography were too slow to perform 1399 continuous monitoring of left ventricular volume, ejection fraction, and cardiac output. However, 1400 now there are small and lightweight scintillation probes, which can perform radionuclide angiography right at the bedside. A radionuclide indicator is injected into the circulation, and the
 radionuclide count density is measured throughout the cardiac cycle^{202,203}. The maximum count

- 1403 density throughout the cycle represents end-diastole, while the minimum represents end-systole²⁰².
- 1404 In this way, the ejection fraction may be determined with the following equation^{202,203}:
- 1405

1406

$$EF = \frac{end\ diastolic\ counts - end\ systolic\ counts}{total\ diastolic\ counts - background\ counts}\ (31)$$

1407

1408 Count averaging is used (roughly 5 heartbeats) to increase the reliability of the measurement and 1409 raise the count density²⁰². The drawback of this approach is namely the ionizing radiation, which 1410 is safe in shorter timeframes, but unsuitable for extended monitoring. Because count averaging is 1411 used, irregular activities such as fibrillation or ectopic activities can affect the accuracy²⁰².

1412

1413 (e) Velocity measurements

1414 Cardiac output can also be found by measuring the blood velocity at a vessel, while knowing 1415 its diameter. Ways of measuring the blood velocity include Doppler ultrasound and 1416 electromagnetic flow probes. In Doppler ultrasound, the changes in the signal frequency are related 1417 to the blood flow velocity²⁶:

- 1418
- $\frac{Fd}{Fs} = \frac{u}{x} (32)$
- 1420

1423

1421 where Fd is the frequency shift, Fs is the source frequency, u is the velocity, and x is the sound 1422 speed. Thus, by measuring the frequency shift, the flow velocity can be measured.

In electromagnetic flow probing, a probe is implanted and slipped around a blood vessel.
According to Faraday's law of magnetic induction, a conductor (i.e., the blood in this case) moving
through a magnetic field generates a voltage that is proportional to the velocity¹⁸⁰:

- 1427
- 1428 1429

1430 where e is the induced voltage, B is the magnetic flux density, L is the spacing between 1431 electrodes, and u is the flow velocity.

e = BLu (33)

1432

Velocity measurements must first be calibrated by measuring the blood vessel diameter before it can be converted to cardiac output¹⁹². In the case of Doppler, the angle of the probe to the blood flow must additionally be identified¹⁸⁰. While studies have found Doppler to be inaccurate in measuring cardiac output, especially in children, its strength is in monitoring the trends and its capability of detecting rapid changes¹⁹². This makes it useful for monitoring surgical or fluid administration procedures in clinical settings. In addition, Doppler is one of the few non-invasivemethods that measure cardiac output.

1440

1441 Electromagnetic flow probes face significant drawbacks in that they require the blood vessel 1442 to be fully exposed through surgery such that the probe can be positioned around the blood 1443 vessel¹⁸⁰. The accuracy of this method also depends strongly on how well the surgical exposure is 1444 done¹⁸⁰. Overall, it is an unfavorable method in most situations.

1445

1446 (f) Pulse contour analysis

1447 In pulse contour analysis, the arterial pulse wave is recorded to produce a measurement of 1448 cardiac output. The area under the pulse wave curve is related to the stroke volume, and therefore the area under the curve times the heart rate can be related to the cardiac output¹⁹². To derive the 1449 relationship for reliable monitoring, a three-element Windkessel model of the aorta's mechanical 1450 characteristics and the peripheral resistance of the body was used to represent the circulation^{192,193}. 1451 1452 To calibrate the model, the relationship between the aortic pressure and cross-sectional area must be derived, so that the vessel compliance can be estimated¹⁹³. Studies have also further developed 1453 the model to be able to identify the stressed and unstressed states of the left ventricle and 1454 1455 additionally calculate the ejection fraction²⁰⁴.

1456

Due to the significant amount of calibrations required for this approach, the accuracy is limited. In addition, as with most models, ideal conditions must be satisfied for the model to work well. Indeed, current commercial devices for measuring cardiac output via pulse contour analysis demonstrate a lack of ability to properly account for peripheral circulatory changes¹⁹². The pulse contour analysis method can be unreliable during surgeries, such as during liver transplantation, where cirrhosis patients have high cardiac output and low fluctuating peripheral resistance¹⁹².

- 1463
- 1464 (g) Bioimpedance

Bioimpedance techniques relate the changes of electrical impedance across the thorax to the cardiac cycle to monitor the cardiac performance. Bioimpedance is also one of the few noninvasive techniques for monitoring cardiac output. Classical methods of processing the impedance signals were dubious in their accuracy and whether they measured the blood flow of the heart at all¹⁹². Patients with conditions such as pulmonary edema, where excessive lung water was present, repeatedly had poor results in validation studies of bioimpedance¹⁹².

1471

Recently, more reliable methods of processing the bioimpedance signals have been developed.
Electrical velocimetry is based on the idea that red blood cells are randomly oriented when there
is no blood flow in the aorta, and subsequently become aligned when the aortic valve opens. This
alignment produces a change in the conductivity of the blood.

1477

The overall bioimpedance measured can be modeled with the following equation:

- 1478 1479
- 1480

1483

1481 where Z_0 is the quasi-static background impedance, ΔZ_R is the artifact from respiration, and ΔZ_C 1482 is the impedance change due to the cardiac cycle.

 $Z(t) = Z_0 + \Delta Z_R + \Delta Z_C \quad (34)$

1484 From the impedance curve of ΔZ_c , one can determine the acceleration of the blood flow based 1485 on the time taken to align the red blood cells. An average velocity can then be estimated and 1486 converted to stroke volume and cardiac output based on body mass.

1487

1488 In adults, electrical velocimetry is just within the 30% limits of agreement²⁰⁵, but is not reliable 1489 in children, likely due to the body mass-based assumptions¹⁹².

1490

1491 In summary, these methods tend to have several sources of error due to their reliance on 1492 models and assumptions. The lack of imaging-derived left ventricular volume measurements 1493 causes them to be reliable only under ideal conditions and to include extraneous signal sources in 1494 their measurements. The use of imaging is not viable in long-term continuous monitoring either, 1495 with the currently available devices. Non-imaging techniques can monitor for slightly longer 1496 periods of time, but are generally too invasive and inconvenient, with many requiring 1497 catheterizations and injections, cannot be used in everyday life or during exercise, and are also not 1498 viable solutions for long-term monitoring. In addition, those invasive approaches require expertise in the placement and manipulation of the catheters to get the correct results. Only well-trained 1499 clinicians can handle this measurement modality⁹⁷. As a result, many of these techniques are 1500 1501 typically only used within the intensive care unit or during an operation, where their benefits 1502 sufficiently outweigh the drawbacks. Thus, there is a significant technological gap that can be 1503 filled by a wearable, versatile, and non-invasive imager that can do direct imaging-derived measurements of the left ventricular volume. 1504

1505

1506 9.5 Anatomical considerations of imaging posture

1507 In the apical four-chamber view, we could only collect data during recovery but not during exercise (Fig. 5). This is not to do with a technological limitation, but rather an intrinsic anatomical 1508 1509 fact. Extended Data Fig. 4 has shown that the apical four-chamber view is best captured in the lying side position, which has limited the continuous monitoring during exercise. From our 1510 supplementary data (Supplementary Fig. 28) and literatures^{206,207}, it is generally accepted that the 1511 1512 apical four-chamber view is best captured in the lying side position and will be partly 1513 blurred/blocked in other positions. The lying side position is the clinically required position to obtain a qualified apical four-chamber view^{206,207}. 1514

The positional requirements for the apical four-chamber view are very strict for two reasons. First, only when the patient is in the lying side position does the ultrasound beam enter the heart's apex at a straight angle (Supplementary Fig. 28). Other positions (e.g., standing) will yield incorrect measurements of chamber dimensions and thus cardiac indices.

1520

1521 Second, we compared the apical four-chamber images from a commercial imager taken by an 1522 experienced cardiologist and from a wearable imager taken by a trained amateur, both with the 1523 subject in the standing position (Supplementary Fig. 28). The cardiac chamber morphologies 1524 cannot be fully reconstructed in either image. The right ventricle and right atrium have been totally 1525 shadowed by the lungs. While we can roughly see the morphology of the left ventricle and left 1526 atrium, the image quality is poor. The left ventricular endocardium cannot be visible, and the left 1527 atrium is malformed, because part of the ultrasound array has been blocked by the lungs and only a small fraction of the elements play an effective role in the imaging process²⁰⁷ (Supplementary 1528 1529 Fig. 28). It results in images with low spatial-resolutions and low contrast-to-noise ratios.

1530

1531 On the contrary, lying in the lying side position brings the relative orientations of the heart 1532 and the transducer array out of the lung shadow. Therefore, the inability to collect high-quality and 1533 morphologically representative apical four-chamber images in the standing position is more of an 1534 intrinsic anatomical fact rather than a limitation of the commercial imager or the wearable imager. 1535 Thus, we imaged the heart in the apical four-chamber view only with the lying side position.

1536

1537 However, the apical four-chamber view is not the only source of information in 1538 echocardiography. We have also presented continuous monitoring of the parasternal long axis view 1539 throughout the whole process (before exercise, during exercise, and during recovery; Fig. 4), as 1540 the anatomy does not limit this view as strictly as it does for the apical four-chamber view. The 1541 M-mode recording in Figure 4b is extracted from the B-mode recording. Extended Data Fig. 5 1542 shows that the patch is capable of taking B-mode images during exercise and illustrates how the 1543 M-mode is extracted. The chamber motion is noticeable, and the change of heart rate is also 1544 computed in Figure 4c. Abundant information, such as left ventricular internal diameter end 1545 diastole/systole, the fractional shortening ratio, and the anatomic integrity of the left ventricle are 1546 also determined. In addition, Figure 4d and Figure 4e are presenting the details from different 1547 stages, demonstrating the capability of wearable imager in the continuous monitoring during 1548 exercise.

1549

In addition, monitoring the cardiac status can be valuable during many surgeries and other interventions to prevent further complications²⁰⁸. Given that the lying side position is the standard position during the echocardiography measurement for ejection fraction, our device can be directly applied to numerous situations where monitoring of cardiac function is essential and lying side position is easy to reach. For example, during general anesthesia as well as the postoperative period, there is always the risk of cardiac events for patients who undergo noncardiac surgeries, such as myocardial infarction and heart failure²⁰⁸⁻²¹². Therefore, the monitoring of the cardiac function is valuable in this context especially for the patients with risk factors²⁰⁸. During the beginning of general anesthesia, the lying side position can be easily reached, thus providing a preliminary evaluation of the cardiac function preoperatively with our device.

1560

In this work, two-dimensional cardiac images can generate an approximate left ventricular volume by the Modified Simpson's rule, which is widely accepted in clinical diagnosis even though it comes with approximation and inaccuracy. Future work can follow up on developing solutions for imaging in the parasternal long axis view during exercise/stress testing. For example, we can fabricate a 2D matrix probe to reconstruct images three-dimensionally. The cardiac volume and other related parameters can be directly and more accurately computed from threedimensionally reconstructed images.

1569 Supplementary Discussion 10: Detailed left ventricle working processes

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1571 A new cardiac cycle starts from the end of left ventricular relaxation in the last cycle. When 1572 the pressure in the left ventricle drops below that in the left atrium, the pressure difference pushes 1573 the mitral valve to open, so blood flows from the left atrium to the left ventricle. The inflow of 1574 blood triggers the first volume increment of the left ventricle, which is called rapid inflow. The 1575 blood inflow builds up the pressure in the left ventricle and at the same time releases the pressure 1576 in the left atrium. As the pressure difference decreases and the inflow rate slows down, the volume of the left ventricle hits the first equilibrium, and the mitral valve tends to close. This period is the 1577 interval between passive blood flow and active blood flow caused by atrium contraction, which is 1578 1579 named diastasis. Afterward, the left atrium starts to contract to build the second pressure difference. 1580 The pressure difference again enforces the mitral valve to open and causes the second stage of 1581 blood inflow to the left ventricle. Simply, the process of this pressure generation is called atrial 1582 systole or atrial contraction. At the end of this process, the volume of the left ventricle reaches its 1583 peak as the second equilibrium.

1584

1585 Then, the left ventricle starts to contract to raise the pressure inside. Because the mitral valve and aortic valve keep closed during the contraction, the volume of the left ventricle holds. 1586 1587 Therefore, this contraction is called isovolumetric contraction. The mitral valve is a one-way valve, 1588 so there is no blood countercurrent back into the left atrium and only the aortic valve opens after 1589 the isovolumetric contraction. The blood in the left ventricle is then ejected into the aorta, which 1590 is called rapid ejection. The rapid ejection results in an obvious drop in the left ventricular volume 1591 and pressure so there is a sharp slope in the curve. Like the rapid inflow, as the pressure difference 1592 between the aorta and left ventricle vanishes, the aortic valve closes. Following the ejection, a 1593 small portion of blood in the left ventricle remains. Because the aortic valve is also a one-way

valve to keep the unidirectional blood flow, the left ventricular volume keeps stable during relaxation, which is also named isovolumetric relaxation. At the end of the isovolumetric relaxation, next cardiac cycle commences.

1597

1598 The precise monitoring of cardiac events can provide more insights into pathologies of cardiac 1599 dysfunctions. For example, the ejection rate based on the volume curve can reveal the myocardium 1600 contractility and stenosis of aortic valves.

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1602 Supplementary Discussion 11: Neural network for continuous ultrasound imaging

1604 11.1 Developing the deep learning model

We surveyed different deep learning models in the literature and made the selection based on their popularity and relevance to this study. Currently, widely-adopted image segmentation deep learning models can be classified into two categories: U-Net models, which are known for fast and precise biomedical image segmentation with multiple variants²¹³, and Fully Convolutional Networks (FCN) models, an end-to-end deep learning method, which first uses a series of fully connected convolutional layers to find useful features from the input image and then use an upsampling layer to restore the output image size to the same as the input image size²¹⁴.

1612

1613 Specifically, U-Net models are based on the well-established U-Net architecture for fast and precise biomedical image segmentation²¹³. U-Net architecture²¹³ has gained wide adoption. Many 1614 researchers have proposed variant models based on the U-Net architecture. In this study, in 1615 1616 addition to the original U-Net model, we also worked with its variants, including Attention-U-Net (U-Net model with an attention gate)²¹⁵, U-Net++ (U-Net with redesigned skip pathways, dense 1617 skip connections, and deep supervision)²¹⁶, NAS-UNet (Neural Architecture search with a U-Net 1618 architecture)²¹⁷, U-Net+ResNet50 (U-Net with ResNet encoders)²¹⁸, and U-Net mini (a simplified 1619 structure of the original U-Net²¹⁹). 1620

1621

The FCN model is widely used for semantic segmentation²¹⁴. FCN gained wide adoption for 1622 its outstanding performance and no restriction on the input image size. Specifically, an FCN model 1623 1624 first uses a series of fully connected convolutional layers to find useful features from the input 1625 image and then uses an upsampling (backwards strided convolution) layer to restore the output 1626 image size to the same as the input image size. If we return the output directly after solely using 1627 the upsampling layer, we call the network FCN-32 based on the number of upsampling operations. 1628 It is believed that spatial location information could be lost when going deep in the neural network. 1629 One solution for this is to fuse the predictions of multiple stages of the convolution layers. FCN-1630 8, for example, fuses the predictions of the final three layers, and therefore only needs to upsample for eight times. Unless otherwise specified, all of our models used vanilla encoder as auto-encoder. 1631

We also experimented with other types of auto-encoders, such as VGG and ResNet, in which caseswe specify the auto-encoder names in the model name.

1634

To ensure accuracy of the pre-processing of input images, we performed frame-level segmentation/labelling of the left ventricle under the guidance of a cardiologist, and followed the Modified Simpson's rule²²⁰ for volume calculation. For the commercial imager, we collected totally 500 frames on static subjects, 221 out of which are manually labelled for training. For the wearable imager, we collected overall 500 frames on the static subjects and 5858 frames on the moving subjects, 201 and 2029 out of which, respectively, are manually labelled for training.

1641

We used data augmentation techniques to expand the limited labelled dataset. We applied rotation, scaling, and gaussian noise to the data to augment the dataset. Specifically, we generated four additional images for each of the labelled images by rotating the original data clockwise/counterclockwise by 5 degrees, rescaling it by a factor of 1.2, and applying gaussian noise to it. The results indicated that the data augmentation slightly improved the performance of the deep learning model (Supplementary Fig. 27).

1648

We evaluated the models' performance quantitatively using Mean-Intersection-Over-Union (mIoU), which is one of the most widely used evaluation metrics for benchmarking object detection²²¹. Specifically, intersection-over-union (IoU) is used to compare the similarity between two shapes²²¹ and is calculated using:

 $IoU = \frac{Area \ of \ intersection}{Area \ of \ union} \ (35)$

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- 1654
- 1655

The mIoU is then calculated by taking the mean of IoU for all images in a dataset. In this study, we calculated mIoU by taking the mean of IoU of each pair of the deep learning model predicted image segmentations and the manually labelled image segmentations. Specifically, for each (predicted-image, labelled-image)" pair, its IoU equals to the number of pixels within the left ventricle in both images divided by the total number of unique pixels within the left ventricle by either of the two images. The evaluation results for different models are listed in Supplementary Fig. 26 and Supplementary Table 5.

1663

During exercise, small amounts of the recorded B-mode images were corrupted when the heart was blocked by the deep-breathing lung. To solve this problem, we applied an imputation algorithm to complement the missing parts in these corrupted images to generate a continuous waveform (Supplementary Fig. 29).

1668

1669 11.2 Detailed analysis of the neural network

1670 Wearable sensors can generate continuous images. A milestone in the roadmap of developing 1671 wearable sensors is to integrate deep-learning algorithms to process the continuous images automatically. There are some commercial systems⁹¹ and research studies^{3,90} on developing 1672 algorithms or deep-learning models for quick calculation of ejection fraction and cardiac output, 1673 1674 but the data output is still discontinuous because the commercial probe needs to be operated 1675 manually and can only produce intermittent data points. Integrating the wearable patch with deep-1676 learning models can continuously output a large amount of actionable information and clinical 1677 insights, which have not been achieved in the literature.

1678

For the deep learning component, our primary goals are (1) to identify image segmentation models to automatically extract the left ventricular volume from B-mode images; (2) to integrate the best-performing models in our research settings and analyze their performance. In the following sections, we provide more analysis and discussions about training paradigm comparison and model selection, data collection and processing (containing sample sizes), model implementation and parameter tuning strategy (containing learning rate), model performance, and future directions.

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- 1687

(1) Training paradigm comparison and model selection

1688 Researchers have made tremendous progresses in improving automatic image segmentation 1689 using various types of machine learning models in recent years. Most researchers focused on using 1690 supervised learning models^{222,223}, which are models that learn from a training dataset with "correct" 1691 ground-truth labels. Many researchers have reported promising results using supervised learning 1692 models for medical image segmentation^{3,223}. Although supervised learning models achieve high 1693 accuracy in image segmentation tasks, their weakness is in the expensive training cost because 1694 they acquire the training set through manual labeling in most cases.

1695

1696 In this study, we mainly focused on integrating supervised deep learning models into our research settings because segmentation results with a high accuracy are a prerequisite for 1697 1698 extracting accurate cardiac performance metrics from the images. The weakness of supervised 1699 learning can be considerably mitigated by few-shot learning and reinforcement learning (detailed 1700 discussions are provided in future directions below). Among various supervised learning image segmentation models, those gaining the most popularity²²⁴ are Convolutional Neural Networks, 1701 which are deep learning models that directly learn from raw pixel data by adaptively assigning 1702 weights to different areas/pixels of the input²²⁵. Specifically, in the field of medical image 1703 1704 segmentation, fully Convolutional Neural Networks have achieved superior results compared to traditional Convolutional Neural Networks models in many studies²¹³. Therefore, we chose to 1705 1706 work with different variations of FCN models and U-Net models that are fully Convolutional 1707 Neural Networks performing particularly well within medical research settings^{222,223,226}. We have

discussed each of the specific variations of these models in the Methods and SupplementalInformation.

1710

1711 (2) Data collection and processing

1712 The deep learning models in this work were implemented with readable, consistent, and well-1713 structured codes, high-quality datasets, and reproducible results. The sample size is listed in 1714 Supplementary Table 4. All data were used by all models. Notably, we worked with two types of 1715 datasets in this study. The first type was "static" datasets, which contained left ventricle ultrasound 1716 B-mode images we collected from static subjects. The second type was "after exercise" datasets, 1717 which contained left ventricle ultrasound B-mode images we collected from subjects after intense 1718 physical exercise. The "after exercise" datasets were more challenging to process because (1) the 1719 size of the left ventricle was continuously and rapidly changing, leading to a higher variability in 1720 the dataset; (2) it contained noisier images, in which the left ventricle was sometimes blocked by 1721 the lung because the subjects were breathing heavily. To the best of our knowledge, this study 1722 represents the first to continuously segment cardiac ultrasound images with intensively changing 1723 left ventricular volumes. After collecting the data, we cropped the image size to be 512×512 1724 pixels and manually segmented them to create the ground-truth as the training dataset. Furthermore, 1725 we used data augmentation techniques such as rotating, scaling, and adding gaussian noises to 1726 expand the size of the training dataset.

1727 1728

(3) Model implementation and parameter tuning strategy

1729 We implemented the models based on open-source codes listed in Supplementary Table 3. 1730 The final hyperparameters after careful tuning in this study are also listed in Supplementary Table 1731 3. Specifically, when training the models, we selected adaptive moment estimation (Adam) as the 1732 optimizer for all models. We found Adam to be the best optimizer among other optimization 1733 algorithms because it not only trains the best-performing models but also converges faster and 1734 requires less time for tuning. As an adaptive optimizer, Adam adaptively changes learning rates for different parameters, and, therefore, the models' performance was not much affected by the 1735 1736 initial learning rate. Thus, we applied each model implementation's default learning rate in most 1737 cases.

1738

1739 Because most of the neural network models in this study were very deep, batch normalization 1740 was applied to make the networks faster and more stable. Specifically, when training deep neural 1741 networks, an update in the training process could change parameters in all layers, causing them to 1742 have different distributions after each update. This makes the optimization process like chasing a 1743 "moving" target and thus causes the network to become unstable and slow to train. We applied 1744 batch normalization to solve this problem. Specifically, each Convolutional Neural Network layer 1745 was connected with a batch normalization layer. The batch normalization layers standardized the 1746 network layer's output by applying a transformation that makes each layer's output mean close to

1747 0 and standard deviation close to 1. This will effectively keep each layer's input distribution1748 unchanged, and thus stabilize the network and speed up the optimization process.

1749

1750 (4) Model performance

We have compared the models' performance both quantitively and qualitatively (Extended Data Figs. 6 and 7, and Supplementary Fig. 26). Specifically, we found that images segmented by FCN-32 model achieved not only the highest mIoU but also the highest image fidelity. We achieved automatic image segmentation using the FCN-32 model.

1755

1756 One of the reasons that FCN-32 achieved better results than U-Net models was that FCN-32 did not have skip-connections. As introduced in the Supplementary Fig. 25, FCN-32's structure 1757 1758 included a series of fully-connected convolutional layers and a single upsampling layer at the end. 1759 Compared with FCN models, U-Net models had more upsampling layers and concatenated outputs 1760 of previous layers using skip connections. Although these differences made U-Net models more 1761 robust to scale variations, there might be risks that the connections brought back some unnecessary 1762 features. Because all of our data were of similar scales, the robustness to scale variations was not 1763 a priority in our dataset. Therefore, the additional robustness brought by the skip-connections in 1764 U-Net models was unable to make significant improvements to the model. On the other side, the 1765 skip-connections and additional up-sampling layers possibly brought back redundant features that 1766 harmed the model's performance. We found FCN models usually output one singular region of 1767 left ventricle, while U-Net models sometimes output a few small detached external pieces 1768 (Extended Data Fig. 6). Additionally, the edges of U-Net's segmentations also appeared to be 1769 "fuzzier" than those of FCN's, proving that FCN was better at removing redundant features (Extended Data Fig. 6). Similarly, although the fusing step in FCN-8 could restore some location 1770 1771 information, it could also bring back redundant information that was already filtered by previous 1772 layers, and hence made FCN-8's performance slightly worse than FCN-32's.

1773

1774 (5) Future directions

1775 The FCN-32 model with the current dataset was sufficient for image segmentation in this 1776 study, which facilitated the frame-by-frame output of those cardiac indices. The FCN-32 had good 1777 generalization capability regarding different left ventricular volumes and different image noise 1778 levels. As for the application to new transducers and perspective data on new subjects in future 1779 studies, the generalizability of the current model could be further improved by expanding the training dataset or optimizing the network. For example, few-shot learning⁴⁹, an advanced version 1780 1781 of supervised learning, can achieve promising results with the model trained on only a few labeled images for each subject²²⁷. Using this model, it is easier to expand the cohort size to achieve a 1782 1783 higher data diversity by collecting images from a small number of subjects. Another potential strategy is reinforcement learning⁵⁰, which is another training paradigm alongside supervised 1784 1785 learning and unsupervised learning. It can optimize models' training and data augmentation
- 1786 approaches so that those models can be better tuned and less overfitted²²⁶, thus improving the
- 1787 models' generalizability. By integrating those strategies, the FCN-32 model will potentially be
- adapted to a larger population with nosier data.



Supplementary Fig. 1 | Characterization of the transducer array. a, Electrical impedance
spectrum with the amplitude and phase angle. It shows the 3 MHz resonance frequency and 3.87
MHz antiresonant frequency. The calculated electromechanical coupling coefficient is 0.67. b,
Map of the dielectric loss of all transducer elements in the orthogonal imager. c, The pulse-echo

- 1793 response and corresponding frequency spectrum of the transducers, showing a wide bandwidth of
- 1794 ~55% and a central frequency of 3 MHz. **d**, Crosstalk between a pair of adjacent elements and a
- pair of second nearest neighbors, which is lower than the standard -30 dB indicated by the dashed
- 1796 line. e, The pulse-echo response and corresponding frequency spectrum of the transducers,
- 1797 showing a wide bandwidth of ~30% and a central frequency of 3 MHz.



1798 Supplementary Fig. 2 | Fabrication processes of the wearable imager. a, Sonicate liquid metal

- in toluene with SEBS to homogenize the materials. b, Print the liquid metal composite on a SEBS
- 1800 substrate using a doctor blade. c, Pattern the liquid metal-based composite electrode using laser
- 1801 ablation. d, Print a subsequent layer of liquid metal insulated with a layer of SEBS on the previous
- 1802 electrode. More layers of electrodes can be fabricated by repeating this step. e, Drill vertical-
- 1803 interconnect-accesses (VIAs) using laser ablation to allow electrical connection between the top
- 1804 electrodes and transducers. **f**, Pattern the shielding layer, bottom electrode, and alignment mask
- 1805 using laser ablation. **g**, Dice the transducer array together with the backing layer. **h**, Spin coat
- 1806 toluene-ethanol solution onto the electrodes to allow adhesion between electrodes and transducers.
- 1807 **i**, Bond the top electrodes to the transducer array. **j**, Bond the bottom electrode to the transducer
- 1808 array. **k**, Irrigate the gap between the two glass slides with Ecoflex to encapsulate the device. **l**,
- 1809 Lift off the glass slides to release the device. **m**, Soften the shielding layer and bond it to the device.



1810

1811 Supplementary Fig. 3 | Images showing the fabrication resolution of the liquid metal

1812 composite electrodes. a, Optical image with reflected illumination and b, scanning electron

- 1813 microscope, showing the minimum width of the liquid metal composite electrodes. c, Optical
- 1814 image with transmitted illumination and **d**, scanning electron microscope, showing the narrowest
- 1815 grooves on the liquid metal composite electrodes patterned by laser ablation.



1816 Supplementary Fig. 4 | Mechanical testing of the liquid metal composite electrodes. a, The 1817 high stretchability of the electrodes allows ~700% maximum strain. Optical and scanning electron 1818 microscope images of the liquid metal composite electrodes b, before and c, after uniaxially 1819 stretched for 100 % strain. There are no visible cracks in the electrode after stretching, indicating 1820 its excellent intrinsic stretchability.



1821 Supplementary Fig. 5 | Results of lap shear strength tests. Both liquid metal electrode and pure

1822 SEBS are characterized. The curves peak values represent the bonding strength.



3rd layer



Shielding layer

- 1823 Supplementary Fig. 6 | Optical images of the multilayered liquid metal composite electrodes.
- 1824 **a**, Ground electrode. **b**, The first layer. **c**, The second layer. **d**, The third layer. **e**, The fourth layer.
- 1825 **f**, The shielding layer. All images share the same scale bar.



1826 Supplementary Fig. 7 | The thickness of the SEBS substrate and the printed liquid metal

1827 **composite.** The thicknesses of **a**, the SEBS film and **b**, the liquid metal composite layer were

1828 measured by a Dektak profilometer. The thin thicknesses of the substrate and the electrode

1829 contribute to the overall low form-factor of the wearable imager.



1830 Supplementary Fig. 8 | Characterization of noise levels after applying different strategies 1831 sequentially. a, B-mode images and b, noise amplitude after Step 1: Shielding electrode. Step 2: 1832 Electrical matching between the transducer and the pulser. Step 3: Ground wire modification by adding an inductor and capacitor in series to the ground wire. The modification rendered the 1833 ground wire to be more resistive at around the transducer center frequency and drain noise at 1834 1835 around the transducer center frequency to the ground better. Other noise can be effectively 1836 removed by filters. Step 4: Signal accumulation, which is a built-in function provided by 1837 Verasonics. The signal accumulation overlaps recent frames to counteract the running noise. We 1838 normalized the noise level by dividing all noise levels by the highest noise level for better 1839 comparison. c, Signals from the cardiac myocardium mixed with noises without shielding. d, 1840 Signals with much reduced noises after shielding.



1841 Supplementary Fig. 9 | Stress-strain curve of the entire device. The cardiac imager was
1842 stretched uniaxially, from which the Young's modulus of the entire device was calculated to be
1843 921 kPa. It showed the device had a similar modulus to the human skin (420 to 850 kPa)²⁴. The

1844 test was performed under nominal stress, which equals to force divided by initial cross-sectional

1845 area of the entire device.



After stretching



- 1847 and ground layers are opaque, we patterned dots on the SEBS substrate to represent the location
- 1848 of each transducer element. Optical images show the device **a**, before and **b**, after stretching. The
- 1849 spatial distribution of arrayed elements is comparable to that obtained through simulation in Figure
- 1850 1e, verifying the accuracy of the simulation.



1851 Supplementary Fig. 11 | The structure of the phantom for device characterizations. We used 1852 a commercial phantom (CIRS 539) to characterize multiple properties of the wearable imager. The 1853 signal-to-noise ratio, axial, lateral and elevational resolutions at different depths, and axial 1854 accuracy were tested when the device was put at the position 1. The lateral accuracy, as well as 1855 axial and lateral resolutions at different lateral distances, were tested when the device was put at 1856 the position 2. The dynamic range, contrast-to-noise ratio, and contrast resolution were tested when 1857 the device was put at the position 3. The dead zone was tested when the device was put at the 1858 position 4.



1859 Supplementary Fig. 12 | Characterization of resolutions and acoustic fields with different

transmission methods and angles. a, The lateral and axial resolutions of wide-beam 1861 compounding, mono-focus, and plane-wave transmission^{156,228}. The wide-beam compounding

- 1862 transmission has the best resolutions among all three. Acoustic fields of **b**, plane-wave, **c**, mono-
- 1863 focus, and **d**, wide-beam compounding transmission methods, with transmission angles of -21°,
- 1864 0°, and 21° . The plane-wave strategy produces the worst resolutions. This is because this mode
- 1865 only transmits a single plane-wave, resulting in a low echoic signal-to-noise ratio, and poor spatial
- 1866 resolutions. The mono-focus strategy yields a greater signal-to-noise ratio, and better spatial
- 1867 resolutions. However, the resolution will deteriorate outside the focal zone. The wide-beam
- 1868 compounding transmission has a stronger and more uniform acoustic wave intensity over a larger
- area than the other two.



1870 Supplementary Fig. 13 | The mechanism of wide-beam compounding B-mode imaging. a, 1871 Multiple frames are first acquired with multiple transmissions at different angles. The multipleangle scan compensates the low echoic energies from regions away from the center, expanding the 1872 1873 insonation area from being rectangular to sector-shaped. The enhanced echoic energy improves 1874 the resolution at high steering angles. The frames are collected at the same rate as the high pulse repetition frequency. The final image is obtained by the superposition of acquired frames, which 1875 1876 achieve synthetic focusing with improved resolution over the entire ultrasonographic window. 1877 Additionally, the superposition helps eliminate the random noise in the images. b, Schematics of the acoustic field simulation setup. All key parameters are labeled and set the same as the practical 1878 1879 imaging procedure.



1880 Supplementary Fig. 14 | Signal-to-noise ratio as a function of step size and number of steering 1881 angles of the wide-beam compounding imaging. The image's signal-to-noise ratio firstly rises 1882 and then falls with the angle step size but increases monotonically with the angle number. When 1883 the angle step size initially increases, the most constructive interference of multiple acoustic fields 1884 is in the region of interest, ensuring the highest signal-to-noise ratio for the image reconstruction. 1885 As the angle step size keeps increasing, the overlap between individual acoustic fields decreases, 1886 resulting in a reduced signal-to-noise ratio. The signal-to-noise ratio increases as the number of 1887 angles grows, because all individual acoustic fields are more or less coherently integrated to 1888 reconstruct images. However, an excessive number of angles sacrifices the imaging temporal 1889 resolution. We used 96 steering angles with a 1° step, which gives adequate penetration depth and 1890 spatial resolutions while maintaining an acceptable frame rate of 20-30 Hz.



1891

1892 Supplementary Fig. 15 | The flow chart of receive beamforming. a, The reflected echoes are 1893 received by the transducer elements, whose signals are amplified by the time gain control to 1894 enhance the weak signals from deep objects. The amplified signals are then converted to digital signals by an analog to digital converter (ADC), and then sent into a delay calculator for phase 1895 difference correction and signal alignment. Direct summing of the synchronized signals may result 1896 1897 in significant side-lobe artifacts. Therefore, adaptive apodization assigns varying weights to the various signals, which are eventually summed together as beamformed signals with an enhanced 1898 signal-to-noise ratio. **b**, Schematic calculation for phase correction. (x_{focal} , z_{focal}) is the focal point. 1899 (x_i, z_i) is the ith transducer. (x_p, z_p) is the pixel of interest. θ is the steering angle. df is the focal 1900 depth. r is the curvature radius. φ is the angle departure of the ith transducer from z-axis on the 1901 1902 curvature. c, Schematic receive beamforming of ultrasound signals. There are two beamformed 1903 signals A and B. The lateral diffusion in A is less than that in B, which indicates a better lateral 1904 resolution of A. In other words, the closer the imaging area to the transducer elements, the better 1905 the lateral resolution. 1906



| 1907 | 0 Pixel value 255 |
|------|--|
| 1908 | Supplementary Fig. 16 Gray scale B-mode images of phantoms and selected windows for |
| 1909 | calculating the dynamic range. Red: The +15 dB contrast gives an average pixel value of 159.8 |
| 1910 | Orange: The +6 dB contrast gives an average pixel value of 127.3. Yellow: The +3 dB contrast |
| 1911 | gives an average pixel value of 110.1. Green: The -15 dB contrast gives an average pixel value o |
| 1912 | 38.7. Cyan: The -6 dB contrast gives an average pixel value of 68.5. Blue: The -3 dB contrast give |
| 1913 | an average pixel value of 80.9. These pixel values are labelled on the pixel scale bar at the bottom |
| 1914 | The dynamic range is thus calculated to be 63.2 dB, which is well above the 60 dB threshold |
| 1915 | usually used in medical diagnosis to give adequate details of the echo patterns in the images. |



1916 Supplementary Fig. 17 | Detailed comparison of the imaging metrics between the wearable 1917 and the commercial imagers. a, Accuracy of the detected scatter positions as a function of the 1918 scatter depth. b, Elevational resolution as a function of depth. c, Lateral resolution as a function of 1919 depth. d, Axial resolution as a function of depth. Each test was repeated for five times. Each 1920 column or point is defined by the mean and the standard deviation (the error bar) of the results 1921 from five tests.



1922 Supplementary Fig. 18 | Schematic experimental setups of resolution tests. The setups are for
a, the commercial imager and b, the wearable imager. The degrees of freedom of each imager are

- 1924 labeled in red arrows. c, The axial resolutions of the wearable imager and the commercial imager
- 1925 at each depth were measured five times. The five degrees of freedom of the commercial imager
- 1926 and the three degrees of freedom of the wearable imager introduced measurement variations. The
- 1927 higher the degrees of freedom, the higher the possibility of worsening the measured axial
- 1928 resolutions. Even though the commercial imager bandwidth is larger than that of the wearable
- 1929 imager, some data points of the commercial imager are worse than those of the wearable imager
- 1930 because of the measurement variations.



1931 Supplementary Fig. 19 | Processes of evaluating the surface curvature for phase correction.
1932 a, Scan the imaging sites on the subject using a 3D scanner. b, Obtain 3D surface reconstruction
1933 from the scanning. c, Select the two sites of interest in this study and build intersection networks

- in the Catia software. d, Zoomed-in schematics of the two intersection networks from a
 reconstructed 3D surface. e, Collect the average curve radii from curves in the intersection
 networks. f, Fit every intersection curve with a smooth curve. Select the intersection curve whose
- 1937 fitting radius is the closest to the mean radius of all curves from the site. The fitting curve is then
- 1938 used to correct the phased distortion induced by the surface curvature of human body.



1939 Supplementary Fig. 20 | B-mode images collected with different couplants. a-d, The PSAX,

- PLAX, A2C and A4C views collected with evaporative ultrasound gel. e-h, The PSAX, PLAX,
 A2C and A4C views collected with non-evaporative silicone. No obvious structural differences
- are found in the comparison. PSAX: parasternal short axis view; PLAX: parasternal long axis view;
- are found in the comparison. PSAX: parasternal short axis view; PLAX: parasternal long axis view;
- 1943 A2C: apical two-chamber view; A4C: apical four-chamber view.



1944 Supplementary Fig. 21 | Quantitatively evaluation of different coupling conditions. 1945 Comparison of the coupling performance between commercial ultrasonic gel and liquidous 1946 silicone couplants. Received signals of the single transducer with **a**, commercial ultrasonic gel, 1947 and **b**, liquidous silicone couplant. **c**, The two couplants result in comparable signal-to-noise ratios 1948 of the received signals. **d**, Cardiac images taken under different coupling conditions. The left panel 1949 shows the image from a commercial rigid ultrasound imager without couplant. The middle panel 1950 shows the image from a wearable ultrasound imager without couplant. The right panel shows the 1951 image from a wearable ultrasound imager with silicone couplant.



1952 **Supplementary Fig. 22 | Optical images of attaching the wearable imager to the chest for** 1953 **long-term.** Optical images of the chest **a**, before attaching the probe, **b-i**, 0, 1, 2, 4, 8, 12, 16, and 1954 24 hours after the attachment, and **j**, after detaching the imager from the human body. Sweat 1955 droplets can be seen in the zoomed-in inset in **e**, after the subject finished working out. Water 1956 droplets can be seen in the zoomed-in inset in **g**, after the subject finished showering.



1957 Supplementary Fig. 23 | Continuous surface temperature and heart rate monitoring for 1

hour. a, Recording the surface temperature of the device by a thermal camera every minute for 1
hour. The highest temperature is ~33 °C, which is harmless to the human body. b, Monitoring the

heart rate using a oximeter every half minute with and without the device attachment. No obvious

1961 difference is observed, showing the safety of the wearable cardiac imager for long-term monitoring.



- 1962 Supplementary Fig. 24 | Images of the parasternal long axis view from 10 subjects using a
- **recycled device.** Expanding the testing cohort size validates the reproducibility and reliability of
- 1964 the wearable imager.



1965 Supplementary Fig. 25 | The structure of the FCN-32 neural network. The FCN-32's structure

- 1966 includes a series of fully-connected convolutional neural network (CNN) layers and an upsampling
- 1967 layer at the end. We used the AlexNet structure for the downsampling process. The input layer is
- 1968 first connected to five groups of connected CNN layers, then connected to two additional CNN-
- 1969 dropout bilayers, and finally connected to an upsampling layer to restore to its original size.
- 1970 Specifically, the five CNN groups have similar structures but different dimensions. The input of
- 1971 each group is first zero-padded and sent into a 2D convolutional layer. Then, we used batch
- 1972 normalization to standardize the CNN's output and activate it with a Rectified Linear Unit and
- 1973 downsample it with max-pooling.



1974 Supplementary Fig. 26 | The comparison of the intersection over union among different 1975 models used in this study. The comparison is made a, before and b, after exercise. For a pair of 1976 predicted images and manually-labelled ground truth image, its intersection over union equals to 1977 the number of pixels that are classified as within the left ventricle in both images divided by the 1978 total number of unique pixels that are classified as within the left ventricle by either of the two 1979 images. The figure shows each model's Intersection over Union on a testing dataset. The FCN-32

- 1980 performs the best with the highest intersection over union and its variation is among the lowest.
- 1981 1.5 interquartile range (IQR) is a common rule in statistics to differentiate the outliers. Data points
- 1982 outside of this range are regarded as outliers.





1983 Supplementary Fig. 27 | The types and results of data augmentation. a, Four types of data

augmentation and corresponding segmentation results. We applied rotation, scaling, and gaussian

- 1985 noise to the data to augment the size of the dataset. **b**, The data augmentation increases the average
- and reduces the variation of mean intersection over union.




- 1996 images. **g**, Because a part of the transducer array was blocked by the lung, only the unblocked part
- 1997 could send ultrasound to the LV and LA, which led to a low signal-to-noise ratio and a lower image
- 1998 quality in the standing position than that in the lying side position. Both the right ventricle and
- 1999 right atrium were mostly obscured due to lung shadow.



2000 Supplementary Fig. 29 | Validation of the image imputation algorithm. To validate the 2001 reliability of this image imputation algorithm, we manually erased three periods from **a**, an 2002 originally continuous volume wave to get **b**, the erased result before imputation. After imputation, 2003 c, the completed new wave has a good agreement with the original one with a 0.93 Pearson 2004 correlation coefficient. To define the hyperparameter N, we compared different imputation results 2005 with **d**, N=3, **e**, N=4, and **f**, N=5. The results show the differences between the generated waves 2006 from various N numbers are negligible. We decide to use N=3 in practice for its simplicity and 2007 algorithm efficiency.



Supplementary Fig. 30 | B-mode images of the abdominal area and liver from the wearable and the commercial imagers. a, B-mode images of the abdominal area with a depth of ~6 cm. Similar structures including the inferior vena cava and abdominal aorta can be recognized in both images. b, B-mode images of the liver with a depth of ~8 cm. The complete boundary and fine

2012 structures such as the hepatic vein can be observed in both images.



- Supplementary Fig. 31 | B-mode images of biopsy tests on a commercial phantom (CIRS 052).
 a, The cross section and longitudinal section of the area of interest before inserting the biopsy
 needle. b, The cross section and longitudinal section of the area of interest after inserting the biopsy
 needle. c, The cross section and longitudinal section of the area of interest after releasing the inner
 stylet. d, The cross section and longitudinal section of the area of interest after releasing the biopsy
 needle. The positions and the behaviors of the biopsy needle are clearly recorded in two orthogonal
 orientations simultaneously by the wearable imager. The uniquely enabling capability of the
- 2020 wearable imager is to forgo the need for an operator to constantly hold the device.



Supplementary Fig. 32 | B-mode images of cardiac anatomies tested by an experienced cardiac sonographer. B-mode images from a, parasternal long axis view, b, parasternal short axis view, c, apical two-chamber view, and d, apical four-chamber view using a clinical ultrasound machine.



Supplementary Fig. 33 | Photograph and schematics of the imaging system. a, Experimental
 setup of the imaging system. Key components have been labelled. b, Working flow chart of the
 system. Red dashed box: Analog signal conditioning path. The low-noise amplifier pre-amplifies
 the raw signal with high fidelity to facilitate the following conditioning. The time gain control is a

2029 programmable amplifier that can selectively amplify electrical signals induced by echoes from 2030 different depths. The intensity loss from deep regions is compensated in the time-gain control. The 2031 programmable gain amplifier allows the overall pixel level to be instantly adjustable when imaging. The anti-aliasing filter is a low-pass filter that cuts off the high-frequency component beyond the 2032 2033 Nyquist frequency to make unambiguous analog-to-digital sampling. The corner frequency of the 2034 anti-aliasing filter is determined before each run according to the sampling rate. Blue dashed box: 2035 Digital signal conditioning path. The 23-tap FIR filter is programmed based on the transducer's 2036 center frequency and filters out signals with frequencies over four times of the central frequency. 2037 The decimator decimates the signals by N times, where N depends on the central frequency and sampling rate to lower the required bandwidth for data transmission. The 41-tap FIR bandpass 2038 2039 filter only allows data near the center frequency to pass, which refines the signals. Another 2040 decimator downsamples the data stream according to the setting of the bandpass filter and 2041 sampling rate to maximize the transmission efficiency. Finally, apodization is applied to the data 2042 in each channel to fulfill the requirement for gradual aperture tapering. ACF: anisotropic conductive film; ADC: analog digital converter; FPGA: field-programmable gate array; PCIe: 2043 2044 peripheral component interconnect express; FIR: finite impulse response; I/O: input/output.



Supplementary Fig. 34 | Configuration of a Mills cross array. It includes a transmit array and a receive array. Transmit and receive beamforming applied by both arrays helps focus in the elevational direction. But the signal-to-noise ratio and the lateral resolution are limited in the reconstructed image due to the small effective aperture.

| Parameters | Spatial | Temporal | Radioactivi | Invasivenes | Mappin | Citatio |
|-----------------|------------|------------|-------------|-------------|--------|---------|
| Modality | resolution | resolution | ty | S | g | n |
| Magnetic | | | | | | |
| resonance | 1.6 mm | 33.3 ms | No | No | 3D | 5 |
| imaging | | | | | | |
| X-Ray | | | | | | |
| computed | 0.3 mm | 50 ms | Yes | No | 3D | 6 |
| tomography | | | | | | |
| Single photon | | | | | | |
| emission | 10 mm | 37.5 ms | Yes | No | 3D | 7 |
| computed | 10 11111 | 57.5 1115 | 105 | 110 | 50 | |
| tomography | | | | | | |
| Positron | | | | | | |
| emission | 2 mm | 2000 ms | Yes | No | 3D | 7 |
| tomography | | | | | | |
| Optical voltage | 1 mm | 0.25 ms | No | No | 2D | 8 |
| map | 1 111111 | 0.25 1115 | no | NO | 20 | |
| Optical | | | | | | |
| coherence | 0.001 mm | 85 ms | No | No | 3D | 9 |
| tomography | | | | | | |
| Ultrasonography | 1 mm | < 1 ms | No | No | 3D | 10,11 |

Supplementary Table 1 | Summary of existing imaging methods for the heart. Spatial resolution, temporal resolution, radioactivity, invasiveness, and mapping capability are evaluation parameters in this study. Comprehensive analysis of these parameters of cardiac imaging technologies helps us understand standards of medical imaging technologies, which serve as guidance for developing the wearable cardiac imager.

| Model | Bonding strength (kPa) | Reference |
|--|---------------------------|-----------|
| Histoacryl | 2 | 22 |
| Tisseel | 6 | 22 |
| Bioglue | 8 | 22 |
| 3M Electrically Conductive Adhesive Transfer Tape 9703 | 13 | 229 |
| Coseal | 18 | 22 |
| 3M Electrically Conductive Adhesive Transfer Tape 9712 | 20 | 230 |
| 3M Electrically Conductive Adhesive Transfer Tape 9713 | 22 | 231 |
| 3M Electrically Conductive Adhesive Transfer Tape 9719 | 27 | 232 |
| 3M EMI Copper Foil Shielding Tape 1181 | 27 | 233 |
| 3M 3313 Copper Foil Tape | 228 | 234 |
| Liquid metal electrode | 236 kPa | This work |
| Pure SEBS | 250 kPa | This work |

Supplementary Table 2 | Comparison between the bonding strength of the liquid metal
 electrode, pure SEBS, and commercial adhesives. The bonding strength between the electrode
 and the transducer element is stronger than many commercial adhesives, preventing the electrodes
 from delamination under various deformations.

| Model type | Training paradigm | Input Image Resolution | Learning rate | Optimizer | Code availability |
|---------------------|---------------------|---------------------------|------------------|-----------|----------------------|
| U-Net | Supervised learning | 512×512 | 10-3 | Adam | 235 |
| FCN-32 | Supervised learning | 512×512 | 10-3 | Adam | 236 |
| FCN-8 | Supervised learning | 512×512 | 10-3 | Adam | 237 |
| Attention-U- Net | Supervised learning | 512 × 512 | 10-5 | Adam | 238 |
| U-Net++ | Supervised learning | 512×512 | 10-4 | Adam | 239 |
| FCN-8 VGG | Supervised learning | 512×512 | 10-3 | Adam | 240 |
| VGG-U-Net | Supervised learning | 512×512 | 10-3 | Adam | 241 |
| Resnet50-U- Net | Supervised learning | 512 × 512 | 10-3 | Adam | 242 |

Supplementary Table 3 | Model parameters and code availability. Key parameters of all
 implemented models include model type, training paradigm, input image resolution, learning rate,

2060 optimizer, and code availability. The models' hyperparameters were carefully tuned and were

2061 implemented based on readable, consistent, and well-structured open source code.

| | Static, commercial probe | Static, wearable patch | After exercise, wearable patch (without data augmentation) | After exercise, wearable patch (with data augmentation) |
|------------------------------|--------------------------------|------------------------------|--|---|
| Size of labeled data | 221 | 201 | 2029 | 10145 |
| Size of unlabeled data | 279 | 299 | 3829 | 3829 |

Supplementary Table 4 | Sample sizes for all models. All models were trained with all data
 collected in static and after intensive exercise.

| Model Name | Mean Intersection over Union (static) | Mean Intersection over Union (after exercise) |
|-----------------|--|--|
| Attention-U-Net | 0.77 | 0.64 |
| U-Net++ | 0.78 | 0.65 |
| Nas-U-Net | 0.69 | 0.52 |
| Resnet50-U-Net | 0.91 | 0.84 |
| FCN-8 | 0.92 | 0.85 |
| FCN-8 VGG | 0.91 | 0.83 |
| U-Net | 0.85 | 0.74 |
| U-Net-mini | 0.16 | 0.09 |
| FCN-32 | 0.93 | 0.87 |

2064 Supplementary Table 5 | Mean Intersection over Union among different models. The table 2065 displays each model's mean Intersection over Unions on a testing dataset when the subject is static and after exercise. The evaluation of Intersection over Union is based on the comparison of a pair 2066 2067 of predicted image and ground truth image. Specifically, the Intersection over Union equals to the number of pixels in the overlaid area of both images divided by the number of pixels in the 2068 2069 combined area of both images. Mean Intersection over Union is the average Intersection over 2070 Union across all images in the testing dataset. The FCN-32 model has the highest mean Intersection 2071 over Union among all models.

2072

| 2073 | Supplementary Video 1. Cardiac long and short axis views imaged by an orthogonal array. |
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| 2074 | |
| 2075 | Supplementary Video 2. Cardiac apical four- and two- chamber views imaged by an |
| 2076 | orthogonal array. |
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| 2078 | Supplementary Video 3. Continuous cardiac imaging during rest, exercise, and recovery. |
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| 2080 | Supplementary Video 4. Left ventricle segmentation results by FCN-32. |
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| 2082 | Supplementary Video 5. Imaging guided biopsy on a phantom by an orthogonal array. |
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