Supplementary information

A self-powered intracardiac pacemaker in swine model

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Supplementary Note1. Energy output of SICP during each cardiac motion cycle

SICP coverts biomechanical energy from cardiac motion to electricity with timedependent. The average output power \overline{P} was related to the load resistance. The maximum energy output per cycle of SICP can be derived by the following equation ¹:

$$E = \overline{P}T = \int_0^T VIdt = \int_{t=0}^{t=T} VdQ = \oint VdQ$$
$$E_{max} = \frac{1}{2}Q_{sc,max}(V_{OC,max} - V_{OC,min})$$
$$E_{max} = \frac{1}{2}Q_{sc,max}\Delta V$$

Here, E_{max} represents the maximal output energy per cycle. The $Q_{SC,max}$ and ΔV of SICP in vivo were about 6.0 V and 8.5 nC, respectively. Therefore, the E_{max} of SICP for per cycle is about 0.026 µJ.

The pacing threshold energy can be derived by the following equation:

$$E_t = \int_0^T V_t \times I dt = V_t \times I \times T = \frac{V_t^2 \times T}{R}$$

Here, E_t represents the pacing threshold energy, V_t is the pacing threshold voltage. *R* represents the pacing resistance. *T* stands for stimulus pulse durations.

The mean pacing threshold voltage of SICP is 1.5 V with a pulse width of 0.5 ms, the mean pacing impedance of swine is about 953 Ω^2 . Therefore, the mean pacing threshold energy of SICP is 1.18 µJ in animal experiment. On the other hand, Ritter, P. et al. reported early performance clinical test of a miniaturized leadless cardiac pacemaker - Medtronic's Micra TPS³. The mean pacing capture threshold at the 3-month visit for the 60 patients measured with a pulse width of 0.24 ms was 0.51 V (95% CI, 0.45–0.56; P<0.0001), meeting the efficacy objective. Among these 60 patients, the mean electrical values for R-wave sensing amplitude, pacing impedance, and pacing capture threshold at a pulse width of 0.24 ms were as follows respectively: 11.7 ± 4.5 mV, 719 ± 226 ohm, 0.57 ± 0.31 V at implant, 15.6 ± 4.8 mV, 662 ± 133 ohm, 0.48 ± 0.21 V at 1-month, and 16.1 ± 5.2 mV, 651 ± 130 ohm, 0.51 ± 0.22 V at 3-months.

 $E_{maximum pacing threshold} = (0.88 V)^2 \times 0.24 \text{ ms} \div 493 \Omega = 0.377 \, \mu\text{J}$

 $E_{meanmum pacing threshold} = (0.51 V)^2 \times 0.24 \text{ ms} \div 719 \Omega = 0.087 \, \mu\text{J}$

 $E_{minimum pacing threshold} = (0.26 V)^2 \times 0.24 \text{ ms} \div 945 \Omega = 0.017 \, \mu\text{J}$

Therefore, based on the rough calculation we can draw the following conclusion:

 $E_{minimum pacing threshold} \le E_{max} = 0.026 \ \mu J = 1/3.3$ $E_{meanmum pacing threshold} = 1/14.5 \ E_{maximum pacing threshold}$

Supplementary Note2. Maximal Power output of SICP

$$P_{max} = \frac{E_{max}}{T} = E_{max}f$$
$$P_{max} = \frac{1}{2}Q_{SC,max}\Delta Vf$$

Here, P_{max} is the maximal power output of SICP. f denotes the operating frequency that drove by heart, which is about 1.5 Hz. Therefore, the P_{max} of SICP is about 0.039 μ W.

Leadless Pacemaker	<u>SICP</u>	Nanostim LCP	Aveir LSP202V	Micra TPS
Self-powered	Yes	No	No	No
Cost	Low	High	High	High
Length (mm)	<u>42</u>	42	38	25.9
Diameter (mm)	<u>6.8</u>	5.99	6.5	6.7
Volume (cc)	<u>1.52</u>	1	1.1	0.8
Mass (g)	<u>1.75</u>	2	2.4	2
Fixation Mechanism	<u>Helix/Hook</u>	Helix	Helix	Hook
MRI compatibility	Yes	Yes	Yes	Yes

Supplementary Table 1. Comparison of representative commercial leadless pacemakers.



Supplementary Fig.1|The surface of SICP before and after deposited by Parylene-C (Scale bar = $10 \ \mu m$).



Supplementary Fig.2|SEM images of the surface of POM and PTFE without treatment by ICP (Scale bar = $5 \ \mu m$)



Supplementary Fig.3 | Schematic illustration of EHU when working under different tilt angles.



Supplementary Fig.4 A photograph of SICP fixed on the endocardium of the right ventricle in isolated heart (Scale bar = 0.5 cm).



Supplementary Fig.5 | a, V_{oc} , I_{sc} and (b) power density of EHU with different resistance. Source data are provided as a Source Data file.



Supplementary Fig.6 Stability and durability tests of EHU. Source data are provided as a Source Data file.



Supplementary Fig.7| a, V_{oc} , (b) I_{sc} and (c) Q_{sc} of EHU at high frequency operation (~6.5 Hz). Source data are provided as a Source Data file.



Supplementary Fig.8 The PM powered by a capacitor with a capacity of 47 μ F. Source data are provided as a Source Data file.



Supplementary Fig.9|The cytoskeletal structures and cell nucleus of L929 cells stained by immunofluorescence at day 1, 2, and 3, respectively. (Scale bar=200 μm)



Supplementary Fig.10|The viability of L929 cells on the encapsulation film tested by the Cell Counting Kit-8 (n=3, Data are presented as mean \pm SD. ns, no significant differences). Source data are provided as a Source Data file.



Supplementary Fig.11. Localized tissues of the skin to deep layer muscle from the implantation location of the materials after 3 months implantation stained by Hematoxylin and Eosin (H&E).



Supplementary Fig.12|a, Hemolysis (n=3) and (b) coagulation test for the encapsulation materials (scale bar = 2 μ m). Source data are provided as a Source Data file.



Supplementary Fig.13 | A Photograph of the external jugular vein exposed with a small incision. (Scale bar = 1 cm)



Supplementary Fig.14 Photographs of (**a**) delivery system (1: inside sheath, 2: outside sheath, 3,4: delivery catheter), (**b**) delivery catheter with SICP advancement, and (**c**) Sutured incision after device implantation. (Scale bar = 1 cm)



Supplementary Fig.15|The schematic diagram of SICP implantation process.



Supplementary Fig.16 The voltage of a capacitor from 0 V to 3 V within 9000 s at the same electrical output of SICP *in vivo*.

	Postoperative wound	
Number of animals	Experimental purposes	Experimental results
		The AV/B experimental animal

1# (a swine)	 Induce a atrioventricular block (AVB) animal model by radiofrequency ablation Explore the pacing efficiency for PM of SICP <i>in vivo</i> 	The AVB experimental animal model was successfully constructed. The PM of SICP successfully regulated the AVB animal heart rate from 33 bpm to 90 bpm.
2#-5# (four swine)	 Evaluate the performance of the homemade introducer and dilator advancement Energy harvesting of SICP <i>in vivo</i> Evaluate pacing effect of SICP in acute phase 	The homemade introducer and dilator advancement successfully implanted the device into the right ventricle. The <i>in vivo</i> open circuit voltage and short circuit current of SICP were about 6.0 V and 0.2 μA. The SICP achieved successful regulation of animals heart rate (from 90 bpm to 108 bpm) <i>in vivo</i> .
6#-8# (three swine)	Evaluate the long-term stability (energy harvesting, pacing, fixed effect and biocompatibility) of SICP <i>in vivo</i>	The SICP converts cardiac motion energy to electricity and maintains endocardial pacing function during three weeks follow-up period.

Supplementary Fig.17| Photographs of the postoperative animal and statistical analysis of animal

experiments.

Reference

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