1	Supplementary Information
2	A Hybrid Transistor with Transcriptionally Controlled Computation and Plasticity
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#### 77 Methods

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#### 79 Chemicals and Reagents

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PEDOT:PSS aqueous suspension (Clevios<sup>™</sup> PH1000, Heraeus Epurio LLC), ethylene glycol 81 82 (anhydrous 99.8%, Sigma-Aldrich), sulfuric acid (H<sub>2</sub>SO<sub>4</sub>, 95.0-98.0%, Sigma-Aldrich), hydrogen peroxide (H<sub>2</sub>O<sub>2</sub>, 30 wt% in water, Sigma-Aldrich), sodium DL-lactate (NaC<sub>3</sub>H<sub>5</sub>O<sub>3</sub>, 60% in water, 83 84 TCI), sodium fumarate (Na<sub>2</sub>C<sub>4</sub>H<sub>2</sub>O<sub>4</sub>, 98%, VWR), HEPES buffer solution (C<sub>8</sub>H<sub>18</sub>N<sub>2</sub>O<sub>4</sub>S, 1 M in 85 water, pH = 7.3, VWR), potassium phosphate dibasic ( $K_2$ HPO<sub>4</sub>, Sigma-Aldrich), potassium 86 phosphate monobasic (KH<sub>2</sub>PO<sub>4</sub>, Sigma-Aldrich), sodium chloride (NaCl, VWR), ammonium 87 sulfate ((NH4)<sub>2</sub>SO<sub>4</sub>, Fisher Scientific), magnesium(II) sulfate heptahydrate (MgSO<sub>4</sub>·7H<sub>2</sub>O, VWR), 88 (ATCC), casamino Wolfe's Trace Mineral Mix acids (VWR), isopropyl ß-D-1thiogalactopyranoside (IPTG, Teknova), anhydrotetracycline hydrochloride (aTc, Sigma-Aldrich), 89 90 3-oxohexanoyl-homoserine lactone (OC6, Sigma-Aldrich), kanamycin sulfate (C<sub>18</sub>H<sub>38</sub>N<sub>4</sub>O<sub>15</sub>S, 91 Growcells), Riboflavin 5' -monophosphate sodium salt hydrate (>93 %, TCI), and LIVE/DEAD<sup>®</sup> 92 BacLight<sup>™</sup> Stain (L7012, Invitrogen), were used as received. Two-part silicone elastomer 93 (Sylgard<sup>™</sup> 184, Electron Microscopy Sciences) was used according to manufacturer instructions. 94 All media components were autoclaved or sterilized using 0.22 µm PES filters.

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#### 96 Bacteria Strains and Culture

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Bacterial strains and plasmids are listed in Table S1. Cell cultures were prepared from bacterial 98 99 stocks stored in 20% glycerol at -80 °C. The stocks were streaked onto agar plates containing 100 LB (for wild-type and knockout strains) or LB with 25 µg/mL kanamycin (for plasmid-harboring 101 strains), and subsequently grown overnight at 30 °C for Shewanella and 37 °C for E. coli. Single 102 colonies from the plates were picked and inoculated into Shewanella Basal Medium (SBM, Table 103 S2) amended with 0.05% trace mineral supplement, 0.05% casamino acids, and supplemented 104 20 mM sodium lactate (2.85 µL of 60% w/w sodium lactate per 1 mL culture) for Shewanella and 105 20 mM glucose (10 µL of 2 M glucose per 1 mL culture) for E. coli as the electron donor, unless 106 otherwise noted. Aerobic cultures were pregrown in 15 mL culture tubes at 30 °C and 250 rpm 107 shaking. Anaerobic cultures were pregrown using the same procedure outlined above, but with 108 argon purged growth medium in a nitrogen-filled glovebox (S1200, Vigor) and supplemented with 109 40 mM sodium fumarate (40 µL of 1 M sodium fumarate per 1 mL culture) as the electron acceptor. 110 Additional 25 µg/mL kanamycin (10 µL of 2.5 mg/mL kanamycin per 1 mL culture) was 111 supplemented to the pregrowth medium of plasmid-harboring strains. Aerobically pregrown 112 cultures were washed 3x using the sterile SBM growth medium and adjusted to an OD<sub>600</sub> of 1-3.5 113 (NanoDrop 2000C) before being transferred into the glovebox. For steady-state protein 114 expression, strains were pregrown anaerobically without inducer(s) for 6 h before being diluted 115 1:25 into inducer-containing media (from 1000x stocks) and grown for 18-24 hours.

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# 117 Device Fabrication

119 Three versions of the OECTs were used: small channel OECTs, 2-electrode versions of the small 120 channel OECT without the gate, and a large channel OECT. The small channel OECTs were 121 used for the majority of experiments with the following exceptions. In direct channel reduction 122 experiments (Figure 2h, Figure S3g), the 2-electrode versions of the small channel OECT were used and noted as 2-electrode devices or no gate. In UV-vis spectroscopy (Figure 2g, Figure S3c 123 124 - S3e), the large channel OECTs were used to fit the laser aperture of the instrument. The large 125 channel OECTs were labeled as 'large channel OECTs' and the small channel OECTs were noted 126 as original three-terminal OECTs or without any specific naming. OECTs were fabricated 127 according to prior work<sup>1</sup>. Quartz microscopic slides (FQ-S-003, AdValue Technology) were 128 cleaned with soapy water, acetone, and isopropyl alcohol, and dried with nitrogen before oxygen 129 reactive-ion etching (RIE, 150 W, 50 sccm, 120 s). Quartz slides were coated with a photoresist 130 layer (AZ5209E) and lithographically patterned to define the electrode layout. Subsequently, Au 131 electrodes (100 nm) with a Ti adhesion layer (10 nm) were thermally evaporated on the quartz 132 slides, and excess materials were removed with acetone lift-off. Another photolithographic pattern 133 was formed to define the PEDOT:PSS region over the channel and the tip of the gate to obtain a 134 width/length of 150 µm x 10 µm and 500 µm x 500 µm, respectively. The PEDOT:PSS (Clevios<sup>™</sup> 135 PH1000) was filtered (0.22 µm PES filters) and spun cast on the pattered quartz slides, followed 136 by hot plate drying at 90 °C for 15 min and acetone lift-off. To increase the conductivity, 137 PEDOT:PSS films were immersed in ethylene glycol at 90 °C for 3 min over a hot plate. The as-138 fabricated PEDOT:PSS film had an average thickness of 26.5 nm for the channel and 38.3 nm for the gate-tip layer. Polydimethylsiloxane (PDMS, Sylgard<sup>™</sup> 184) with 9 % wt curing agent was 139 140 drop cast and cured for 48 hours at room temperature to ensure surface smoothness. The OECT 141 chambers and fluid access ports in the PDMS sheets were manually cut with a hole punch.

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The OECTs with the larger channel size were only used for UV-vis measurement. The electrodes were fabricated in the same way as the smaller OECT. The PEDOT:PSS channel was fabricated by drop-casing 80 μL PEDOT:PSS over the entire slide, followed by 20 minutes of air-drying and heating at 90°C for 20 minutes. Afterward, the excessive PEDOT:PSS film was removed with cotton swabs soaked with 70% ethanol. Conductivity enhancement was achieved by EG treatment of 10 min at 90 °C. Silicon spacers (0.5 mm, GBL664581, Sigma-Aldrich) were hand cut to create the OECT chambers and only used with the large channel OECTs.

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# 151 Device Operation and Electrochemistry

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153 Shewanella Basal Medium (SBM) amended with 0.05% casamino acids and 1X Wolfe's Trace 154 Mineral Mix was used as the base electrolyte except for the carbon source comparison 155 experiments (Figure 2e, Figure S2d) where the SBM was only amended with 1X Wolfe's Trace 156 Mineral Mix. When noted, the SBM was further supplemented with 40 mM sodium fumarate to 157 support cell growth. Before each experiment, the OECT slides and PDMS sheets were autoclaved 158 separately and assembled in the biosafety cabinet. OECT experiments were conducted in a 159 nitrogen-filled glovebox to create the anaerobic condition except for UV-vis spectroscopy which 160 was conducted under ambient conditions. Media and solutions stocks were purged with argon for 161 15 min and stored in the glovebox. A multichannel potentiostat (MultiPalmSens4, PalmSens BV) 162 was used for the electrochemical measurements. During continuous OECT operation, unless

otherwise noted, the gate ( $V_{GS}$ ) and drain ( $V_{DS}$ ) voltages were biased at 0.2V and -0.05 V, respectively. Transfer curves were measured with a gate scan rate of 20 mV/s, except in the artificial synaptic hysteresis measurement where a rate of 10 mV/s was used. For all the experiments, OECTs were stabilized in the glovebox with abiotic electrolytes and constant bias voltages for 72 hours before inoculation or measurements.

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169 In the electrode potential measurements with Ag/AgCl pellet reference electrode (550010, A-M 170 Systems), the Ag/AgCl electrodes were directly inserted into the OECT chamber without salt 171 bridges. The same Ag/AgCl pellet electrodes were used as the gate for the large channel OECTs 172 in abiotic UV-vis measurements. Hybrid OECT experiments with electroactive bacteria were 173 conducted between 18 - 36 hours after inoculation (initial OD<sub>600</sub> at 0.01). Synaptic measurements 174 were conducted after I<sub>DS</sub> was stabilized for at least 1 minute at V<sub>GS</sub> = 0 V.

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#### 176 Inoculation Procedure

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178 For aerobic culture growth, SBM supplemented with 20 mM lactate was used for cell growth. 179 Following aerobic overnight growth, the cultures were triple-washed by centrifugation using the 180 same growth medium to remove byproducts. The optical density of the washed cultures was 181 measured before transferring them into the glovebox, side the glovebox, the cell cultures were 182 diluted to create an intermediate inoculum with a cell density 10-fold higher than the target OD for 183 final inoculation. This dilution used the same electrolyte solution that filled the OECT. Finally, the 184 OECTs were inoculated using a 1:9 volume ratio of the intermediate cell culture to the OECT 185 electrolyte. For instance, S. oneidensis supplemented with 1 µM exogenous FMN with inoculum 186 OD<sub>600</sub> of 0.05 was prepared from aerobically grown cell cultures. The triple-washed cell cultures 187 (average  $OD_{600}$  at 3.38) were brought into the glovebox and diluted to the intended  $OD_{600}$  of 0.5 188 with SBM supplemented with 1 µM exogenous FMN, creating the intermediate cultures. Then 5 189 µL of these intermediate cultures were inoculated into the OECTs containing 45 µL of SBM 190 supplemented with 1 µM FMN, achieving a final inoculation OD<sub>600</sub> at 0.05.

191

192 For steady-state expression conditions, cells were anaerobically cultured in SBM supplemented 193 with 20 mM lactate and 40 mM fumarate, along with specific inducer(s). To maintain consistent 194 induction strength and to prevent cell growth from affecting OECT readouts, electrolyte in the 195 OECTs were SBM supplemented with 20 mM lactate and specific inducer(s), but lacked fumarate. 196 Prior to induction, strains were grown statically and anaerobically without inducer(s) for 6 hours. 197 Subsequently, cell cultures were diluted 1:25 into media containing inducers (prepared from 198 1000x concentrated stocks) and incubated for an additional 18-24 hours. After this induction 199 period, the cell cultures were directly inoculated into the OECTs at a 100-fold overall dilution 200 without prior washing. Specifically, cells were first diluted 10-fold with the fumarate-free OECT 201 electrolyte forming the intermediate cultures. The final inoculation of the OECTs was achieved by 202 adding the intermediate cell culture to the OECT electrolyte at a 1:9 volume ratio.

203

In the carbon source experiment, SBM amended with 1X Wolfe's Trace Mineral Mix was used as the base electrolyte. Aerobically grown *S. oneidensis* MR-1 cells culture were triple-washed with SBM without a carbon source, then the cell cultures were kept at room temperature for 3 hours to induce starvation conditions<sup>2</sup>. Afterward, cell cultures were washed again and their  $OD_{600}$  was measured. Subsequently, cell cultures were brought into the glovebox and diluted to obtain the intermediate stocks with an intended  $OD_{600}$  of 0.1. The dilutions were performed with SBM supplemented with either 20 mM lactate, 20 mM pyruvate, 20 mM acetate, or no carbon source. Finally, the intermediate stocks were inoculated into OECTs containing the SBM supplemented with the respective carbon source or no carbon source at a ratio of 1:9, achieving a final inoculation  $OD_{600}$  of 0.01.

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215 In cell viability experiments, aerobically grown S. oneidensis MR-1 cell cultures were triple-216 washed with SBM supplemented with 20 mM lactate. For E. coli cultures, the lactate was replaced 217 with 20 mM glucose. The densities for the washed cell cultures were measured by OD<sub>600</sub> and the 218 S. oneidensis cultures were allocated to 3 parts: live, heat-killed, and lysed cells. Heat-killed cells 219 were obtained by incubating at 80 °C for 20 min. Lysed cells were obtained by sonication (Qsonica 220 55, Qsonica LLC) for 90 s at 4 °C. The output power was set to 15 W with "ON" and "OFF" intervals 221 of 10 s and 5 s, respectively. Then cell cultures were brought into the glovebox and diluted with 222 SBM supplemented with 20 mM lactate to obtain the intermediate cultures at an intended OD600 223 of 0.1. The intermediate cultures were inoculated into the OECTs at a ratio of 1:9, achieving a 224 final inoculation  $OD_{600}$  of 0.01. The S. oneidensis supernatants were obtained by filtering (0.22) 225 µm PES filters) the cell cultures (initial OD600 at 0.01) anaerobically grown in the glovebox with 226 SBM supplemented with 20 mM lactate. The entire volume of OECT electrolyte was replaced with 227 the supernatant during inoculation.

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# 229 Spectroscopy

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The UV-Vis spectroscopy of the PEDOT:PSS channel was measured from 190 nm to 1100 nm (Agilent 8453 UV-Visible Spectroscopy System). A custom sample holder was 3D printed to fit the OECT slides to the instrument. Measurements were blanked with devices lacking the PEDOT:PSS channel. When bacteria cells were present in the sample, the blank devices were likewise inoculated with the same inoculum.

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# 237 Fluorescence Microscopy

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239 Microscopy was performed using a Nikon Ti2 Eclipse inverted epifluorescence microscope. 240 Immediately after the 24-hour operation in the glovebox, OECTs were gently washed 2x by 241 refreshing the electrolyte with the sterile SBM supplemented with 0.05% trace mineral supplement, 242 0.05% casamino acids. Then, the PDMS sheets were replaced with a 0.5 mm silicon spacer to 243 ensure the sample thickness was compatible with the working distance of the microscope. 244 Subsequently, the OECTs were gently washed with SBM containing 0.05% trace mineral supplement, 0.05% casamino acids, and LIVE/DEAD<sup>®</sup> BacLight<sup>TM</sup> Stain mix (3 µL of SYTO 9 and 245 246 propidium stocks per 1 mL) at a final solution volume of 10 µL per OECT chamber. The OECTs 247 were then sealed with coverslips, covered with aluminum foil, and transferred out of the glovebox 248 for microscope imaging. Bacterial counts were performed using ImageJ software.

#### 250 Atomic Force Microscopy

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252 The AFM scans were conducted using a DriveAFM (Nanosurf AG). The cantilevers (Dyn190AI) 253 were driven with the photothermal laser (CleanDrive) under dynamic mode. OECTs were 254 randomly selected from two fabrication batches: 3 slides of the as-fabricated OECTs (8 OECTs 255 per slide) were used as pre-inoculation samples, and 3 cleaned OECT slides post the 48-hour 256 inoculation served as post-inoculation samples. Two OECTs per slide were randomly chosen for 257 AFM scans using Nanosurf CX software for data acquisition. Images for PEDOT:PSS film 258 thickness were acquired at 90 µm x 90 µm and processed with Nanosurf CX software to correct 259 background variations. Topology and phase images were acquired at 500 nm x 500 nm and 260 analyzed without further process. Grain sizes were fitted by the Watershed method with the 261 resulting histograms generated using Gwyddion software.

#### 263 OECT Data Processing

Measured OECT data were processed using GraphPad Prism9 and MATLAB (R2021b update 1). The measured  $I_{DS}$  data were normalized to the initial value before inoculation ( $I_{DS0}$ ) before fitting. The  $I_{DS}$  decay rate constants for all samples except the lysed *S. oneidensis* were obtained by fitting  $I_{DS}/I_{DS0}$  data with an exponential decay model:

$$i_{DS}(t) = e^{k \cdot t} \tag{1}$$

Where *t* is time in hours, *k* is the fitted  $I_{DS}$  decay rate constant. Fitting was performed with the built-in one phase decay function in GraphPad Prism9. The  $I_{DS}/I_{DS0}$  data for lysed *S. oneidensis* samples were fitted to a simple linear regression model:

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$$i_{DS}(t) = I_{DS0} + \beta t \tag{2}$$

278 Where *t* is time in hours, the slope  $\beta$  is used as the rate of change. Fitting was performed with 279 the built-in simple linear regression function in GraphPad Prism9.

The response function of FMN concentrations was modeled with a four-parameter logistic regression function in terms of  $I_{DS}$  decay rate constants (noted here as r):

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 $\frac{r}{R_{max}} = \frac{[S]^n}{\left(K_{\frac{1}{2}}\right)^n + [S]^n}$ (3)

284

285 Where *n* is the Hill coefficient, [*S*] is the FMN concentrations,  $R_{max}$  is the maximum rate constant, 286  $K_{\frac{1}{2}}$  is the half-maximum concentration constant. Fitting was performed with the built-in four-287 parameter dose-response function in GraphPad Prism9.

288 In synaptic experiments, MATLAB scripts were used to process the measured data. The

289 'Vgs\_pulse\_pair.m' script was used to identify and extract the channel current  $I_{DS}$  peak values

using islocalmax function (Figure 5d, 5e, 5g - 5i, Extended Data Figure 5a - 5e). The

291 'IDS\_VGS\_Pul\_long.m' script implemented median filtering to remove pulses and extract the

292 baseline for channel current  $I_{DS}$  responding to continuous gate pulse inputs. The filtering was

- 293 achieved by using the medfilt1 function (Figure 5f and Figure S5f). Extracted  $I_{DS}$  curves were 294 fitted to an exponential model  $b(1)^*exp(b(2)^*x(:,1))+b(3)$  with fitnlm function to obtain the rate 295 constants, where b(1), b(2), and b(3) are fitting coefficients, x is time in seconds.
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- 297 All scripts are available through the Texas Data Repository.
- 298

# 299 Statistical Methods

- Independent samples t-tests were conducted using the built-in t-test function with the unpaired
   option in GraphPad Prism9.
- 303

304 In the 2-input Boolean gate figures (Figure 4c and 4d, Figure 5h and 5i), statistical tests were 305 performed using R (version 4.2.3) with multicomp package (version 1.4-23). In brief, we employed 306 a linear model fitting approach to determine the interaction term between the two inputs noted as 307 Factor1 and Factor2, specifically Im(Results ~ Factor1 \* Factor2 - 1). Then, based on the 308 significance of the interaction terms, different linear contrasts were tested for general linear 309 hypotheses, specifically glht(model, linfc = contrast). The significance of the interaction term was 310 examined for p < 0.05. If the interaction term was significant, the following linear contrasts were 311 evaluated for each specific logic gate and model parameterization:

- 312 313
- NAND gate:  $H_0: \mu_{++} + \frac{-1}{3}(\mu_{--} + \mu_{+-} + \mu_{-+}) = 0$  (4)

NOR gate: 
$$H_0: \frac{-1}{2}(\mu_{++} + \mu_{+-} + \mu_{-+}) + \mu_{--} = 0$$
 (5)

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The contrast matrices for the NAND and NOR logic gates in the general linear hypothesis test are as follows: (2/3, -2/3, -2/3, -1/3) for NAND gate, and (-2/3, 2/3, 2/3, 1) for NOR gate. These contrast matrices were calculated by algebraic substitution given the following coefficient/parameter relationships defined by our parameterization of the linear model, specifically:  $\mu_{++} = \text{coef}[1], \mu_{+-} = \text{coef}[2], \mu_{-+} = \text{coef}[2], \mu_{--} = \text{coef}[2] + \text{coef}[3] + \text{coef}[4].$ The resulting p-values and the interaction term are outlined in Table S3.

323 If the interaction term was not significant, a linear model without interaction was fit to the data, 324 and the model was tested with new contrast matrix as:  $(1, -\frac{1}{3}, -\frac{1}{3}, -\frac{1}{3})$  for the NAND gate, 325 and  $(-\frac{1}{3}, 1, -\frac{1}{3}, -\frac{1}{3})$  for the NOR gate. The p-values are outlined in Table S3 without an 326 interaction term.

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# 337 Tables

#### 

# **Table S1**. Strains and plasmids used in this study.

Strain and Plasmid	Description	Source
Strains		
Escherichia coli MG1655	Wild-type strain	Lydia Contreras, University c Texas at Austin
Shewanella oneidensis		
MR-1	MR-1 (ATCC700550) wild type strain	American-Type Culture Collection
∆mtrC	JG596, deletion of genes <i>mtrC</i> , <i>omcA</i> , and <i>mtrF</i> .	Jeffrey Gralnick, University c Minnesota <sup>3</sup>
∆Mtr	JG1194, deletion of genes <i>mtrC</i> , omcA, mtrF, mtrA, mtrD, dmsE, SO4360, cctA, and recA.	Jeffrey Gralnick, University c Minnesota <sup>4</sup>
∆bfe	deletion of genes SO0702.	Jeffrey Gralnick, University o Minnesota⁵
∆lysis	S2933, deletion of genes SO2966 and SO2974.	Lydia Contreras, University o Texas at Austin <sup>6</sup>
Plasmids		
pCD8	Empty Buffer gate	Ref. <sup>7</sup>
pCD24r1	<i>mtrC</i> Buffer gate (sRBS1 <sub>mtrC</sub> )	Ref. <sup>7</sup>
pAT4	<i>Mtr</i> Buffer gate	This work
pNAND- <i>mtrC</i>	mtrC NAND Boolean logic gate	Ref. <sup>8</sup>
pNAND-sfgfp	sfgfp NAND Boolean logic gate	Ref. <sup>8</sup>
pNOR- <i>mtrC</i>	mtrC NOR Boolean logic gate	Ref. <sup>8</sup>
pNOR-sfgfp	sfgfp NOR Boolean logic gate	Ref. <sup>8</sup>

#### Table S2. Shewanella Basal Medium (SBM) formulation 342

Ingredient	Quantity per 1 L
K <sub>2</sub> HPO <sub>4</sub>	225 mg
KH <sub>2</sub> PO <sub>4</sub>	225 mg
NaCl	460 mg
(NH <sub>4</sub> ) <sub>2</sub> SO <sub>4</sub>	225 mg
MgSO <sub>4</sub> *7H <sub>2</sub> O	117 mg
HEPES	100 mL of 1 M HEPES
Casamino acids	5 mL of 10% casamino acids in ddH <sub>2</sub> O, if needed
Wolfe's Mineral Mix	5 mL of Wolfe's Mineral Mix, if needed
ddH <sub>2</sub> O	Adjust volume to 1 L and pH to 7.2

#### 343 344

# Table S3. Relevant statistical values

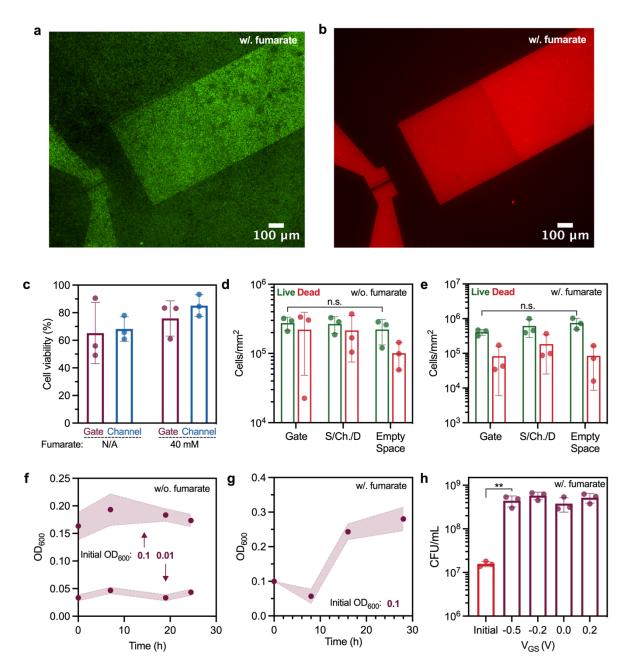
Data Sets	pNAND	pNOR
Source	2-way ANOVA interaction p= 0.00313	2-way ANOVA interaction p= 1.84 x 10 <sup>-5</sup>
Potential (V <sub>s</sub> )	Contrast test p = 0.00204	Contrast test p = $2.36 \times 10^{-7}$
Synaptic Conductance Change (AG)	1-way ANOVA contrast test p = 0.00306	1-way ANOVA contrast test p = 0.00152

# Change (△G)

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# 360 Supplementary Figures

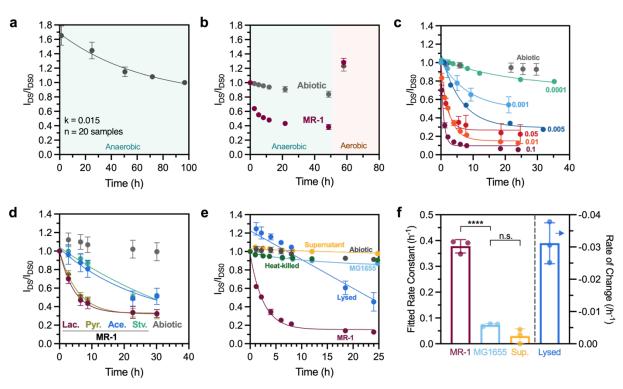




363 Figure S1. Cell viability and growth in OECTs under varying culture and operation conditions. 364 Representative fluorescent microscopy images showing OECTs operated with constant drain 365 voltage  $V_{DS}$  = -0.05V and gate voltage  $V_{GS}$  = 0.2V. Cells were supplemented with 20 mM lactate 366 and 40 mM fumarate. LIVE/DEAD<sup>®</sup> BacLight<sup>™</sup> cell assay showing (**a**) live cells in green and (**b**) 367 dead cells in red. (c) Cell viability derived from the fluorescent microscopy images. (d, e) Cell 368 counts over OECT gates, source/channel/drain region (denoted as S/Ch./D), and spaces 369 elsewhere (denoted as empty spaces) for cells supplemented with 20 mM lactate, and (d) 370 without or (e) with 40 mM fumarate. (f, g) Optical Density at 600 nm (OD<sub>600</sub>) was measured from

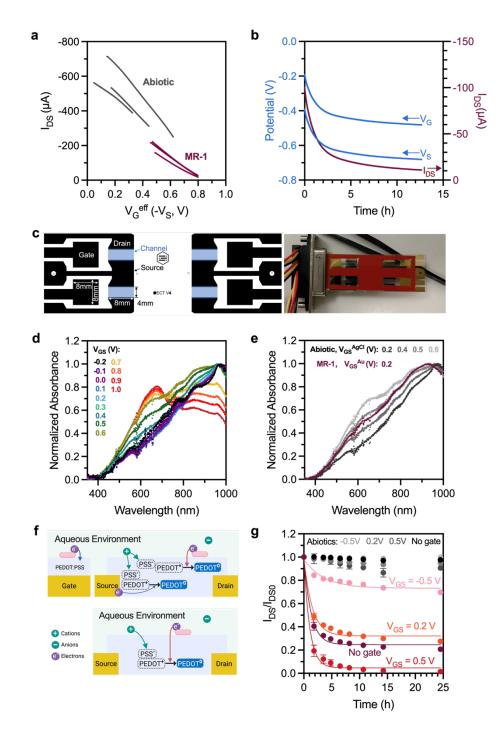
OECTs biased at  $V_{DS} = -0.05V$  and  $V_{GS} = 0.2V$ , cells supplemented with 20 mM lactate, and (f) without or (g) with 40 mM fumarate. The shaded region indicates the range of standard deviation. (h) Colony forming units (CFUs) were counted 24 hours after OECTs operation with constant  $V_{DS} = -0.05 V$  and  $V_{GS}$  biased at -0.5 V, -0.2 V, 0.0 V, or 0.2 V. CFU/mL *p* value for the indicated pair *p* = 0.0042. Cells were supplemented with 20 mM lactate and 40 mM fumarate. Data show the mean ± SD of 3 biological replicates, unpaired two-tailed Student's t-tests were performed without adjustments for multiple comparisons, n.s. represents *p* > 0.05.





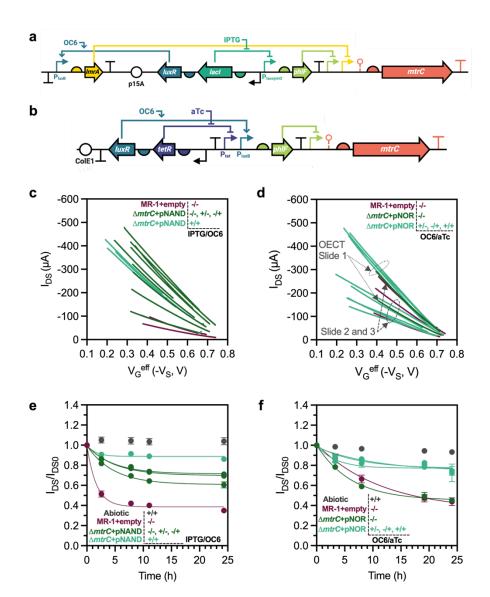
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380 Figure S2. OECT channel current I<sub>DS</sub> reduction in abiotic and different cell metabolic states. The I<sub>DS</sub>/I<sub>DS0</sub> curves during (a) anaerobic stabilization without bacteria cells and (b) 48-hour 381 operation followed by exposure to the oxygen in the ambient environment. The I<sub>DS</sub>/I<sub>DS0</sub> curves of 382 383 S. oneidensis MR-1 inoculated OECTs with (c) varying inoculation  $OD_{600}$  numbers or (d) 384 supplemented with either 20 mM sodium lactate (Lac.), 20 mM sodium pyruvate (Pyr.), or 20 mM 385 sodium acetate (Ace.) as the electron donors. No carbon source was supplied to the starved cells 386 (Stv.) (e) The I<sub>DS</sub>/I<sub>DS0</sub> curves for living, lysed, and heat-killed S. oneidensis MR-1 cells, as well as 387 living E. coli MG1655 cells. (f) The I<sub>DS</sub> decay rate constants for living S. oneidensis and E. coli 388 cells, as well as S. oneidensis supernatant. Fitted rate constant p values for the indicated pair p = 3.8 × 10<sup>-5</sup>. The lysed S. oneidensis curves were fitted using a linear regression model. Cells 389 390 were supplemented with 20 mM lactate for S. oneidensis MR-1 and 20 mM glucose for E. coli 391 MG1655, no electron acceptor was added. Initial inocula were adjusted to OD<sub>600</sub> of 0.01. Data 392 show the mean ± SD of 3 biological replicates, unpaired two-tailed Student's t-tests were 393 performed without adjustments for multiple comparisons, n.s. represents p > 0.05. 394



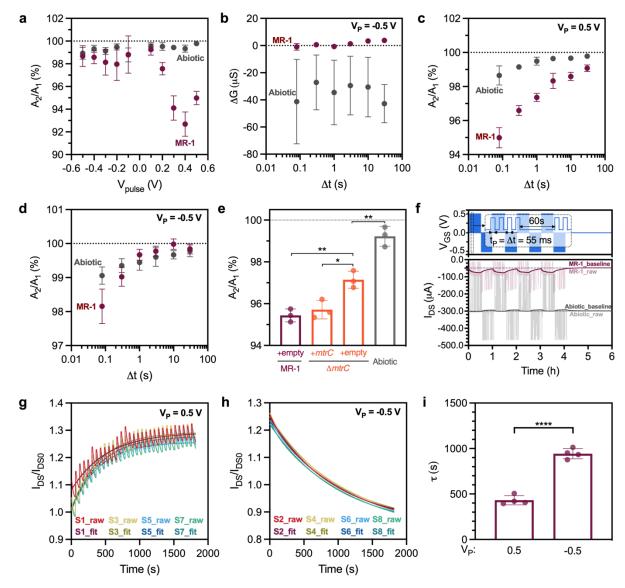
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- **Figure S3**. OECT de-doping investigated using electrochemical and spectroscopy methods. (a) Transfer curves plotted using effective gate voltage ( $V_G^{eff}$ ) values with respect to Ag/AgCl pellet reference electrodes. (b) Gate ( $V_G$ )and source ( $V_S$ ) potentials with respect to the Ag/AgCl pellet reference electrode plotted with the channel current  $I_{DS}$  for an OECT inoculated with *S*. *oneidensis* MR-1. (c, left) Schematic and (c, right) photo-image of the large channel OECT used exclusively for the UV-Vis instrument. UV-Vis spectra were collected for (d) abiotic PEDOT:PSS channel under different Ag/AgCl pellet gate bias voltages, and (e) for *S. oneidensis* MR-1
- 403 inoculated channel overlayed with abiotic channels. (f) Cartoon illustrations comparing the

404 cellular extracellular electron transfer (EET) de-doping process of OECT and the 2-electrode 405 device. (**g**) The channel current  $I_{DS}/I_{DS0}$  curves of the 2-electrode devices and OECTs under 406 constant gate bias conditions. *S. oneidensis* MR-1 was used for inoculum with an inoculation 407  $OD_{600}$  of 0.01. Data in panel (g) show the mean ± SD of 3 biological replicates. Figure (f) 408 created with BioRender.com.



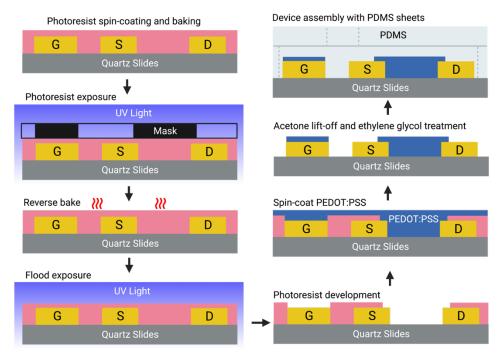


411 **Figure S4**. OECT responses to *AmtrC* strains carrying NAND and NOR Boolean gate plasmids. Cartoon illustrations of the plasmid architecture of the (a) NAND and (b) NOR Boolean gates 412 413 expressing *mtrC*. The  $\triangle mtrC$  mutants carrying the corresponding Boolean gate plasmids were 414 brought to steady-state MtrC expression with combinations of 500 µM IPTG, 200 nM OC6, and 415 10 nM aTc inducers. Transfer curves of the induced (c) NAND and (d) NOR gates samples. The 416 channel current I<sub>DS</sub>/I<sub>DS0</sub> curves for (e) NAND and (f) NOR gates samples with different inducer 417 combinations. Data in panels (e) and (f) show the mean ± SD of 3 biological replicates. Figure (a) 418 and (b) created with BioRender.com.



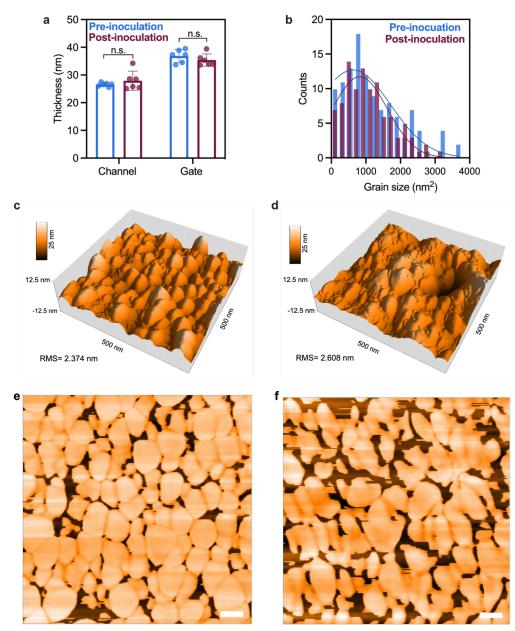


420 Figure S5. Modulation of the OECT synaptic behavior with varying pulse conditions and strains. 421 (a) A2/A1 index plotted with varying pules voltage V<sub>P</sub>, while pules duration t<sub>P</sub> and pules interval 422  $\Delta t$  were fixed at 80 ms. (b) OECT channel conductance changes  $\Delta G$  with varying pulse interval 423  $\Delta t$ , fixed t<sub>P</sub> = 80 ms and V<sub>P</sub> = -0.5 V. A2/A1 index for OECTs with varying pulse interval  $\Delta t$ , fixed 424  $t_P$  = 80 ms and  $V_P$  of (c) 0.5 V or (d) -0.5 V. (e) A2/A1 index of  $\triangle mtrC$  strain carrying mtrC Buffer 425 gate (+mtrC) or empty vector plasmid (+empty) under steady-state protein expression. A2/A1 426 index p values for pairs indicated from top to bottom p = 0.0046, p = 0.0045, and p = 0.0149. (f) 427 Channel current  $I_{DS}$  responding to the continuous voltage pulses with  $V_P$  of 0.5 V or -0.5 V. Faded 428 lines represent the raw I<sub>DS</sub>, while the bolded lines represent I<sub>DS</sub> baselines after filtering out the 429 pulses. (g, h) One-phase exponential fitting of the I<sub>DS</sub> baselines for each continuous 4-pulse 430 session, with  $V_P$  equal to (g) 0.5 V or (h) -0.5 V. (i) The corresponding time constants of the fitting results. Time constant p values for the indicated pair  $p = 1.0 \times 10^{-5}$ . In panel (a) to (e), data show 431 432 the mean ± SD of 3 biological replicates. In panel (e) and (i), unpaired two-tailed Student's t-tests 433 were performed without adjustments for multiple comparisons, n.s. represents p > 0.05.



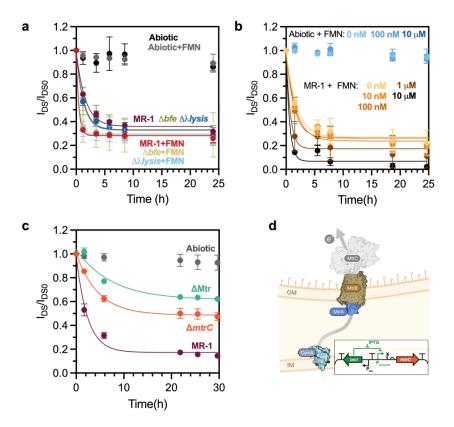
**Figure S6**. Cartoon illustration of the major OECT channel fabrication and assembly steps.

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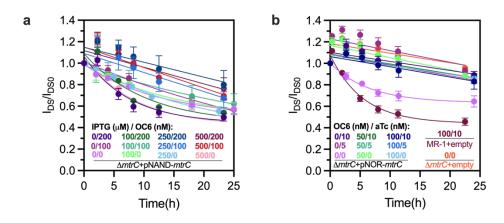
438 Figure S7. Morphological characterization of PEDOT:PSS channel and gate-tip coating.

(a) Thickness of PEDOT:PSS films derived from AFM scans. OECTs were inoculated with S. 439 oneidensis and operated with constant bias voltages at the gate  $V_{GS}$  = 0.2 V and drain  $V_{DS}$  = -440 441 0.05 V for 48 hours. (b) Histogram and Gaussian fit (lines) of PEDOT grain size in the channel 442 region based on (e) and (f). Representative surface topologies of the PEDOT:PSS channel (c) 443 prior to and (d) after the S. oneidensis incubation. Representative phase images of the 444 PEDOT:PSS channel (e) prior to and (f) after the incubation, scale bars represent 50 nm. AFM 445 measurements were performed on 6 independent samples. Panel (c) to (f) presents a zoomed-in 446 morphology graph extracted from 2 representative samples. In panel (a), unpaired two-tailed 447 Student's t-tests were performed without adjustments for multiple comparisons, n.s. represents 448 p > 0.05.



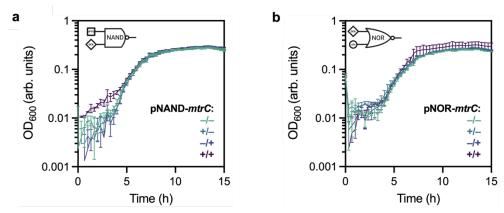
**Figure S8**. OECT response to different extracellular electron transfer (EET) mechanisms.

(a) The I<sub>DS</sub>/I<sub>DS0</sub> curves of knockout strains with and without the addition of exogenous flavin mononucleotide (FMN) (1µM), and (b) S. oneidensis MR-1 cells with varying exogenous FMN concentrations. Initial inocula were adjusted to OD<sub>600</sub> of 0.05. (c) The  $I_{DS}/I_{DS0}$  curves of  $\triangle mtrC$ ,  $\triangle$ Mtr, and MR-1 strains with initial inoculation OD<sub>600</sub> at 0.1. (d) Cartoon illustration of the  $\triangle$ *mtrC* strain and (d, insert) diagram of the mtrC Buffer gate controlled by the IPTG inducer. Faded shapes indicate removed proteins with genomic deletion and inhibition of electron transfer through the inner (IM) and outer (OM) membranes. Data show the mean ± SD of 3 biological replicates. Figure (d) created with BioRender.com.



**Figure S9**. Channel current  $I_{DS}/I_{DS0}$  curves of strains carrying the Boolean logic gates plasmids. (a)  $\triangle mtrC$  strain carrying NAND Boolean mtrC plasmids. (b)  $\triangle mtrC$  strain carrying NOR Boolean *mtrC* plasmids. MR-1 strain and  $\triangle mtrC$  strain carrying empty vectors were used as positive and negative controls, respectively. Inducible mutants were brought to steady-state expression under various inducer combinations and concentrations before inoculation. Data show the mean ± SD

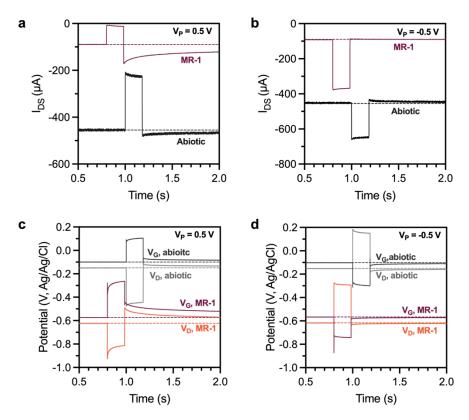
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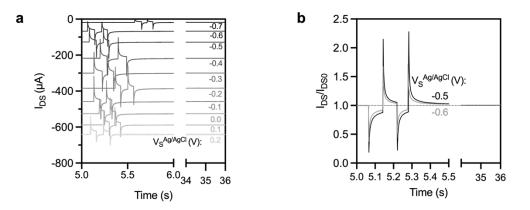
473 **Figure S10**. Growth curves of mutant strains carrying the Boolean logic gates plasmids.

(a) NAND or (b) NOR Boolean *mtrC* plasmids under different inducer (IPTG, OC6, and aTc)
 combinations. Data show the mean ± SD of 3 biological replicates. Created with BioRender.com





**Figure S11**. Measured channel currents  $I_{DS}$  and electrode potentials responding to gate pulses. Single gate pulses with pulse duration  $t_p = 1.5$  s, pulse voltage  $V_P$  equal to (**a**, **c**) 0.5 V and (**b**, **d**) -0.5 V were applied between gate and source electrodes while constant drain voltage  $V_{DS} = -0.05$ V was applied. Gate potential  $V_G$  and drain potential  $V_D$  were measured against Ag/AgCl pseudoreference electrodes.

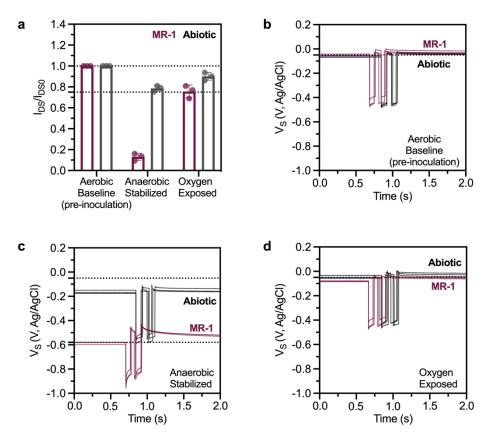


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Figure S12. Paired-pulse responses of abiotic OECTs at varying electrochemical doping states. The source electrode potential ( $V_s$ ) was constantly biased at the specified voltages against the

486 The source electrode potential ( $V_S$ ) was constantly biased at the specified voltages against the 487 Ag/AgCl pseudo-reference electrode. (**a**) Channel current  $I_{DS}$  of OECT with  $V_S$  biased from 0.2 V

- 488 to -0.7 V in increment of 0.1 V. (b) Normalized  $I_{DS}$  for V<sub>S</sub> biased at -0.5 V and -0.6 V.
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491 **Figure S13**. Electrochemical response of the hybrid OECTs under varying oxygen conditions.

(a) Channel current  $I_{DS}$  were normalized to the pre-inoculation aerobic baseline values, with gate voltage  $V_{GS} = 0$  V and drain voltage  $V_{DS} = -0.05$  V. Paired pulse responses of pulse voltage  $V_P =$ 0.5 V and drain voltage  $V_{DS} = -0.05$ V were shown with the measured source electrode potentials against Ag/AgCl pseudo-reference electrode during (**b**) aerobic pre-inoculation condition, (**c**) anaerobic stabilization post-inoculation, and (**d**) subsequent re-oxygenation to ambient conditions. Data in panel (a) show the mean ± SD of 3 biological replicates.

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