

Ion channels of cell membranes



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Nobel Prize in Chemistry 2003

to Peter Agre "for the discovery of water channels" and Roderick MacKinnon "for structural and mechanistic studies of ion channels."





channels	of radii
$a \approx$	$6 \mathring{A}$
and length	

 $L \approx 120 \mathring{A}$

at scale

 $\xi \approx a \sqrt{\epsilon_{\text{water}}/\epsilon_{\text{lipid}}} \ln(\epsilon_{\text{water}}/\epsilon_{\text{lipid}})$ $\approx 140 \text{\AA}$

electric field stays inside channel

Cylindrical Cow

[Kamenev Zheng Larkin Shklovskii]

Ion channels in cell membranes



 $U(x_1 - x_2) \approx eE_0|x_1 - x_2|$

discontinuity $E_0 = 2e/a^2 \epsilon_{\text{water}}^{-}$

Energy to move an ion thru channel $U(L/4) \approx 4k_B T_{room}$ typically impedes ion transport If screened by then barrier drops to $k_B T_{room}$ which is enough to pass an ion



boundary charge (order parameter)

Q

ID Coulomb Gas

Potential Energy U

Grand canonical

partition function

$$\begin{aligned} & = -\frac{eE_0}{2} \sum_{i,j} \sigma_i \sigma_j |x_i - x_j| & \text{ions} \quad n_1 e \quad -n_2 e \\ & \mathcal{Z}_L = \sum_{N_1, N_2 = 0}^{\infty} \frac{f_1^{N_1} f_2^{N_2}}{N_1! N_2!} \prod_{i=1}^{N_1} \int_0^L dx_i \prod_{j=1}^{N_2} \int_0^L dx_j \ e^{-U/k_B T} \end{aligned}$$

We want to rewrite the partition function as quantum mechanical amplitude between two boundary states of charge q

density $\delta[\rho(x) - \sum_{j} \sigma_{j} \delta(x - x_{j})]$ elevate into $\exp \int [d\theta] \exp(i\theta(x)[...])$ Resum x integrals $\sum_{N} [f \int dx \, e^{i\sigma\theta(x)}]^{N} / N! = \exp\{f \int dx \, e^{i\sigma\theta(x)}\}$

Integrate over coordinates (Inverse ID Laplace) $\exp\{(T/eE_0)\int dx\,\theta\partial_x^2\theta\}$

QM amplitude
$$\mathcal{Z}_L = \left\langle q \left| \mathcal{X} e^{-\frac{eE_0}{k_B T} \int_0^L dx \, \hat{H}} \right| q \right\rangle$$

ID Coulomb Gas as Quantum Mechanics

Amplitude
$$\mathcal{Z}_L = \left\langle q \left| \mathcal{X}e^{-\frac{eE_0}{k_BT} \int_0^L dx \, \hat{H}} \right| q \right\rangle = \sum_m |\langle q|m \rangle|^2 e^{-\frac{eE_0L}{k_BT} \epsilon_m(q)}$$

Hamiltonian $\hat{H} = (i\partial_{\theta} - q)^2 - (\alpha_1 e^{in_1\theta} + \alpha_2 e^{-in_2\theta})$ concentration of ions $\alpha_{1,2} = f_{1,2}k_BT/eE_0$

H acts on Hilbert space of periodic functions $\psi_m(\theta) = \psi_m(\theta + 2\pi)$ scalar product $\langle q|m \rangle = \int_0^{2\pi} d\theta e^{-iq\theta} \psi_m(\theta)$

Thermodynamical Transfer Properties

Boundary charge **q** — **Bloch quasi-momentum**.

 $\langle q|m\rangle = \int_0^{2\pi} d\theta e^{-iq\theta} \psi_m(\theta)$

Momentum is periodic with unit period because integral values of charge can be screened by boundary ions

Pressure proportional to first eigenvalue $P = k_B T \frac{\partial \ln \mathcal{Z}_L}{\partial L} \xrightarrow{L \to \infty} -eE_0 \epsilon_0(q)$

Energy barrier for ion transport proportional to the width of first Bloch band $U_0 = eE_0L\Delta_0$

Thus transfer properties are defined in term of $\epsilon_0, \Delta_0, n_1, n_2$ We shall find those using semiclassical methods

Symmetries

[<u>T. Gulden</u>, <u>M. Janas</u>, <u>A. Kamenev</u>, <u>P. Koroteev</u>]

$$\hat{H} = (i\partial_{\theta} - q)^2 - (\alpha_1 e^{in_1\theta} + \alpha_2 e^{-in_2\theta})$$

- PT symmetry. P: $\theta \rightarrow -\theta$ T: complex conjugation
- Isospectrality. $\theta \rightarrow \theta + \theta_0$

$$\alpha_1^{n_2}\alpha_2^{n_1} = \text{const}$$

after scaling

$$\hat{H} = \alpha \left[\hat{p}^2 - \left(\frac{1}{n_1} e^{in_1\theta} + \frac{1}{n_2} e^{-in_2\theta} \right) \right]$$

quantization

 $\hat{p} = \alpha^{-1/2} (-i\partial_{\theta} + q); \qquad [\theta, \hat{p}] = i\alpha^{-1/2}$

concentration = Planck constant!



Band Structure

u-plane







Analytic Solution

First consider $n_1 = n_2 = 1$

Action form Symplectic form $\ d\lambda = dp \wedge d\theta$

Energy equation $2u = p^2 - 2\cos\theta$ $\lambda = pd\theta$



Double cover of complex plane $p^2 = 0$ at $z_+ = -u \pm i\sqrt{1-u^2}$ and $p \sim z^{1/2}$, $(z \sim \infty)$ $p \sim (z - z_+)^{1/2},$ $(z \sim z_+)$





Cosine Potential

Energy equation (complexified)





 $S_j(u) = \oint_{\gamma_j} \lambda$



 $\lambda(u) = p(\theta) \, d\theta = p(z) \frac{dz}{iz} = \frac{(z^2 + 2uz + 1)^{1/2}}{iz^{3/2}} \, dz$



 $p \sim z^{-1/2},$ $(z \sim 0)$ $p \sim z^{1/2},$ $(z \sim \infty)$ $p \sim (z - z_{\pm})^{1/2},$ $(z \sim z_{\pm})$

$$z_{\pm} = -u \pm i\sqrt{1 - u^2}$$

A bit of Geometry

Homology $H_1(T^2; \mathbb{R}) = \mathbb{R}^2$ $\{\delta_0, \delta_1\}$



Cohomology $H^{1}(T^{2};\mathbb{R}) = \mathbb{R}^{2}$ $\{\lambda, \lambda'\}$

Second derivative must be linearly dependent on λ, λ' This results in **Picard-Fuchs** equation on period integrals

$$(u^2-1)S_j''(u)+rac{1}{4}S_j(u)=0$$

3 regular singularities: $u=\infty,\pm 1$

Two solutions

$$S(u) = A F_0(u^2) + B u F_1(u^2)$$

$$F_0(u^2) = {}_2F_1\left(-\frac{1}{4}, -\frac{1}{4}; \frac{1}{2}; u^2\right)$$
$$F_1(u^2) = {}_2F_1\left(+\frac{1}{4}, +\frac{1}{4}; \frac{3}{2}; u^2\right)$$

Period Integrals

Identify period integrals from boundary conditions

 $S_{0}(u) = C_{00}F_{0}(u^{2}) + C_{01}uF_{1}(u^{2}) \qquad C_{00} = e^{-i\pi/2}C_{10} = 8\pi^{-1/2}\Gamma(3/4)^{2}$ $S_{1}(u) = C_{10}F_{0}(u^{2}) + C_{11}uF_{1}(u^{2}) \qquad C_{01} = e^{+i\pi/2}C_{11} = \pi^{-1/2}\Gamma(1/4)^{2}.$

Near
$$u = -1$$
 $S_0(u) \propto (1+u)$
 $S_1(u) \propto \operatorname{const} + (1+u) \ln(1+u)$

Monodromy $(u+1) \rightarrow (u+1)e^{2\pi i}$

$$\begin{pmatrix} S_0(u) \\ S_1(u) \end{pmatrix} \to \begin{pmatrix} 1 & 0 \\ -2 & 1 \end{pmatrix} \begin{pmatrix} S_0(u) \\ S_1(u) \end{pmatrix} = M_{-1} \begin{pmatrix} S_0(u) \\ S_1(u) \end{pmatrix}$$

resurgence

 $S_1(u) = Q_1(u) + \frac{i}{\pi} S_0(u) \ln(1+u)$



Monodromy

Near
$$u = -1$$
 $M_{-1} = \begin{pmatrix} 1 & 0 \\ -2 & 1 \end{pmatrix}$
Near $u = 1$ $M_1 = \begin{pmatrix} 1 & 2 \\ 0 & 1 \end{pmatrix}$

$$S_0(u) = Q_0(u) - iS_1(u)\ln(1-u)/\pi$$

Near $u = \infty$ (classical regime)

$$S_i(u) = u^{1/2} \left[V_i(u) + W_i(u) \ln u \right]$$

with little work

$$M_{\infty} = \begin{pmatrix} 1 & 2\\ -2 & -3 \end{pmatrix}$$

 $V_0(u) = i\pi W_1(u) \mp V_1(u),$ $W_0(u) = \mp W_1(u),$ $V_1(u) = 4i\sqrt{2} \left[\ln \left(e^2/8 \right) + 2/u \right]$ $W_1(u) = -4i\sqrt{2} \left[1 - (4u)^{-2} \right],$

Full monodromy is trivial $M_\infty = M_{-1}M_1$



WKB quantization
Near
$$u = -1$$
 $S_0(u_m) = 2\pi \alpha^{-1/2}(m + 1/2)$

Energy levels $\epsilon_m = 2\alpha u_m$ **Pressure** $P = -eE_0\epsilon_0 = 2k_BTf - \sqrt{k_BTeE_0f}$. Debye-Huckel

Bandwidth via instanton action $(\Delta u)_m = \frac{\omega}{\pi \sqrt{\alpha}} e^{i\alpha^{1/2}S_1(u_m)/2}$

$$S_1(u_m) = 16i + \frac{2i}{\alpha^{1/2}} \left(m + \frac{1}{2}\right) \ln\left(\frac{m + 1/2}{32e\alpha^{1/2}}\right)$$

gives

$$\Delta \epsilon_m = \frac{4}{\pi} \left(\frac{32e}{m+1/2} \right)^{m+1/2} e^{-8\alpha^{1/2} + (m/2+3/4)\ln\alpha}$$

(cf. solutions of Mathieu equation, can do the same for Lame)

Band Spectrum of Mathieu Equation [© Dunne Unsal]



(2, I) Gas



(3,I) Gas

Energy equation describes genus 2 surface

$$\frac{4}{3}u = p^2 - \left(\frac{z^3}{3} + \frac{1}{z}\right)$$



Solutions of Picard Fuchs $(u^4 - 1)S^{(4)} + 8u^3S^{(3)} + \frac{217}{18}u^2S'' + uS' + \frac{65}{144}S = 0$ give four independent cycles





For higher charges of ions
(4,1):
$$\frac{5}{4}u = p^2 - \left(\frac{z^4}{4} + \frac{1}{z}\right),$$

(3,2): $\frac{5}{6}u = p^2 - \left(\frac{z^3}{3} + \frac{1}{2z^2}\right)$

degrees of PF grow

$$(4,1): \quad (u^5+1)S^{(4)}(u) + \frac{9u^5-1}{u}S^{(3)}(u) + \frac{235}{16}u^3S''(u) + \frac{5}{4}u^2S'(u) + \frac{39}{64}uS(u) = 0,$$

$$(3,2): \quad (u^5+1)S^{(4)}(u) + \frac{9u^5-1}{u}S^{(3)}(u) + \frac{140}{9}u^3S''(u) + \frac{5}{4}u^2S'(u) + \frac{119}{144}uS(u) = 0.$$

all solutions have similar semiclassical behavior

$$S(u) \sim \sqrt{u}, \sqrt{u} \log u$$

Resurgence

In a nutshell resurgence is a manifestation of an intimate connection between perturbative and nonperturbative phenomena in QFT/String theory

We have seen it in our QM examples $S_0(u) = e^{-i\pi/2}S_1(e^{i\pi}u)$

The classical action [0] was related to the action on instanton-antiinstanton trajectory $[I,\bar{I}]$

Remarkably, this continues further...

 $[0] \sim [I, \overline{I}] \sim [II\overline{I}\overline{I}] \sim \dots$

The reason of this is that perturbation series in QFT are **asymptotic**

Resurgence — 0+0d example

[A. Cherman, P. Koroteev, M. Unsal]

Integral

$$\mathcal{Z}(g) = \frac{1}{\sqrt{g}} \int_{-\pi/2}^{\pi/2} dx \, e^{-\frac{1}{2g} \sin(x)^2}$$

can be done to get a Bessel function

$$\mathcal{Z}(g) = \frac{\pi}{\sqrt{g}} e^{-\frac{1}{4g}} I_0\left(\frac{1}{4g}\right)$$

two saddles: pert at x = 0, S=0 and non-pert at x = pi/2, S = 1/2g

Taylor expand near x =0, integrate...

$$\mathcal{Z}(g) \stackrel{?}{\simeq} \sqrt{2\pi} \left[1 + \frac{g}{2} + \frac{9g^2}{8} + \frac{75g^3}{16} + \frac{3675g^4}{128} + \frac{59535g^5}{256} + \frac{2401245g^6}{1024} + \cdots \right]$$

coefficients grow factorially and needs to be resumed!

A well-defined answer should have this form

$$\mathcal{Z}(g,\sigma_0,\sigma_1) = \sigma_0 e^{-S_0} \sum_{k=0}^{\infty} p_{k,0} g^k + \sigma_1 e^{-S_1} \sum_{k=0}^{\infty} p_{k,1} g^k$$

Holomorphic Approach

Rewrite the partition function using inverse coupling as

 $(2\xi)^{-1} = 2g$

change variables $\cos \phi = -x$

$$x \qquad \mathcal{Z}_*(\xi) = \int_{\mathcal{C}} \frac{e^{-\xi x}}{\sqrt{(1+x)(1-x)}} dx$$

We can interpret Z as a contour integral

$$\mathcal{Z}_*(\xi) = \int\limits_A \lambda$$

 $\mathcal{Z}(\xi) = \sqrt{\xi} e^{-\xi} \mathcal{Z}_*(\xi)$

 $\mathcal{Z}_*(\xi) = \int^{+\pi} e^{\xi \cos \phi} d\phi = 2\pi I_0(\xi)$

There must be a B-cycle. Let's try Picard-Fuchs $\xi Z_*'' + Z_*' - \xi Z_* = 0$

which has 2 solutions! $\mathcal{Z}_* = C_1 I_0(\xi) + C_2 K_0(\xi)$

Near $\xi = 0$ $K_0(\xi) = -(f(\xi) + \gamma_E) \log\left(\frac{\xi}{2}\right) + g(\xi)$ $I_0(\xi) = f(\xi)$, $f(\xi) = -1 - \frac{\xi^2}{4} + O(\xi^4)$

PF Monodromy

$$K_{0}(e^{\pi i}\xi) = f(\xi)\log\xi - \pi i f(\xi) = K_{0}(\xi) - \pi i I_{0}(\xi)$$
Define $\mathcal{Z}_{*A}(\xi) = K_{0}(\xi)$, $\mathcal{Z}_{*B} = \frac{1}{\pi i}I_{0}(\xi)$
Partition function $\mathcal{Z}(\xi, c_{1}, c_{2}) = c_{1}\mathcal{Z}_{A}(\xi) + c_{2}\mathcal{Z}_{B}(\xi)$
Monodromy around the origin $M_{0} = \begin{pmatrix} 1 & -2 \\ 0 & 1 \end{pmatrix}$
 $\mathcal{Z}_{A} \to \mathcal{Z}_{A} - 2\mathcal{Z}_{B}$

Perturbation series is divergent, however, if we take the series and subtract from it the same series after taking the monodromy, we'll get the `instanton' part. Factorially-divergent pieces disappear

The same can conclusion can be drawn from Borel resumation

Thanks!