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Dr. Boris Farber

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 "Apparently, the possibilities for controlling the thinking process are endless. They cannot be exhausted, because Reason, the greatest instrument of knowledge and transformation of the world, is capable of transforming itself. Who can say that there is a limit to the process of humanizing a person?.. As long as a person exists, the management of this force will be improved. We are only at the very beginning of a long journey." G.S. Altshuller, 1979, "Creativity as an exact science"

Dr. Polovinkin Alexander: Fundamentals of Engineering Creativity, Automation of Search Design



Имеется т п-мерных параллелепипедов

 $a_i^l \ll s_i^l \ll b_i^l, \quad i=1,\dots,n_i, \quad j=1,\dots,m_i, \quad (30)$ как с непрерывным, так и с дискретным характером изменения переменных \overline{X}_i^l . Для каждого из параллелепинедов вадана по единому критерию качества целевая функция

$$f = f'(\overline{X}_j), j = 1 ..., m,$$
 (31)

и система ограничений

$$g_r^i(\bar{X}_i) \ge 0$$
, $r = 1, ..., p_i$, $j = 1, ..., m$. (32)

Требуется найти точку \widetilde{X}_{j}^{*} , принадлежащую j^{*} -му параллеления для которой

$$g_r^{j^*}(\overline{X}_{j^*}^*) \geqslant 0, \quad r = 1, \dots, p_l^{r^*};$$

 $f^{j^*}(\overline{X}_{j^*}^*) = \min_{f^l}(\overline{X}_{j}^*);$
 $1 \leqslant j \leqslant m.$

$$(33)$$

Таким образом, задача структурной оптимизации состоит в нахождении глобально-оптимальной структуры и глобально-оптимальных эначений переменных внутри этой структуры, т. е. эту задачу можно назвать также задачей структурно-параметрической оптимизации.

К задачам структурной оптимизации относится задача выбора оптимальной компоновки ТО. При постановке и решении таких задач следует использовать закономерность минимизации компоновоченых затрят (см. п. 3 гл. 5).

Отметим некоторые особенности задач структурной оптимизации. Во-первых, ночти всегда в этих задачах одновременно присутствуют и дискретные, и непрерывные переменные, т. е. задачи структурной оптимизации в общем случае относятся к смещанным задачам математического программирования. Во-вторых, при структурных преобразованиях изменяются число и характер переменных и соответственно функции ограничений и целевые функции. Что касается характера многосвязной области поиска, то отдельные подобласти или имеют различную размерность или (при совпадении размерности) образованы различными наборами переменных.

Алгоритм поиска глобального экстремума. Алгоритм поиска глобально-оптимального решения можно использовать для решения задач как параметрической, так и структурной оптимизации. Укрупненная блок-схема ал-

 Каждая основная точка делает шаг локального поиска, в результате чего точки (34) переходят в новую последовательность

$$\bar{X}_{1}^{t+1}, \bar{X}_{2}^{t+1}, \dots, \bar{X}_{n}^{t+1}$$
. (36)

3. Синтезируется λ_{t+1} дополнительных допустимых точек, каждой из которых разрешается сделать t+1 шагов локального поиска при условии, что после каждого шага с номером τ (0 \ll τ \ll t) ее критерий не хуже, чем соответствующий член последовательности (35). При нарушении этого условия точка исключается и не участвует в дальнейшем поиске глобального экстремума. Таким образом, имеется q (q \ll λ_{t+1}) дополнительных точек, сделавших t+1 шаг локального поиска:

$$\overline{X}_{1}^{t+1}, \overline{X}_{2}^{t+1}, \dots, \overline{X}_{q}^{t+1}$$
 (37)

$$\overline{X}_{1}^{t+1}, \overline{X}_{2}^{t+1}, \dots, \overline{X}_{n}^{t+1},$$
 (38)

которые являются основными на (t+1)-м групповом шаге поиска. Значение худшего критерия точек из последовательности (38) дополняет последовательность (35) числом \mathbf{w}_{t+1}

 Цикл по пп. 2—4 повторяется до нахождения глобального экстремума по заданным условиям прекращения поиска. В качестве условий прекращения поиска могут быть использованы, например, выполнение заданного числа Т групповых шагов.

Считая параметры λ_i независимыми от i, будем иметь только два настраиваемых параметра алгоритма: η — число основных точек и λ — число дополнительных точек.

Проведенные исследования [1, 17] позволяют рекомендовать следующие оптимальные значения этих параметров: $\eta = 2 \div 3$, $\lambda = 12 \div 18$. Для простоты реализации алгоритма можно брать постоянные значения η и λ .

В качестве процедуры ШЛП рекомендуется использовать следующие алгоритмы поиска локального экстремума [1]:

алгоритм случайного поиска в подпространствах (см. п. 2 гл. 13);



Dr. Polovinkin Alexander's Automation of Search Design

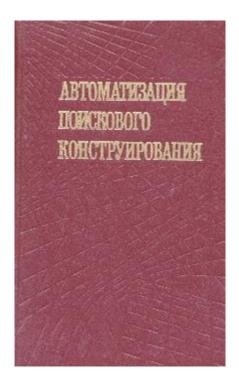


Рис. 38. Главная магистраль

т1, Т2, Т3 — предшествующие массою випускаемые ТО, Т4 — рассматриваемый протогия: с — угол приска без видини ИТР; В — сужекие угла поиска в ориентицией из

развитии ТО:

щих свойств:

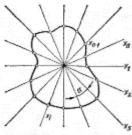
батываемого объекта.

Descriptions of computer Mathematical Modeling methods of exploratory design and construction used to search for improved physical operating principles and technical solutions are

provided.

y x₅₀ y₁₀ y₁₀

Рас. 66. Прямоугольная система компленият



Рис, 67. Полярная системи коор-

зуют либо прямоугольную либо полярную систему координат. Произвольная форма тела описывается в прямоугольной системе (рис. 66) вектором

$$\overline{X} = [(x_1, y_1), (x_2, y_2), \ldots, x_n, y_n)],$$
 (41)

а в полярной системе (рис. 67) вектором

$$\overline{X} = (x_1, x_2, ..., x_n, \alpha), \alpha = 2\pi/n.$$
 (42)

В ряде задач при описании плоскосимметричных и осесимметричных тел более подходит специальная система координат в виде парадлельных осей X_i (рис. 68). При этом высота оптимизируемой формы вдоль оси y делится на отрезки равной ($\Delta y_i \leftarrow$ const) или неравной ($\Delta y_i \neq$ \neq const) длины. Наборы оптимизируемых параметров в этом случае можно описать соответственно векторами

$$\widetilde{X} = (x_1, x_2, \dots, x_m, \Delta y);$$
 (43)

$$\overline{X} := (x_1, x_2, ..., x_n, \Delta y_1, \Delta y_2, ..., \Delta y_n).$$
 (44)

При фиксированной длине отрезков Δy в формуле (43) или Δy_i в формуле (44) набор оптимизируемых переменных описывается вектором (25).

Если требуется иметь однозначное плавное очертание формы.

значное плавиое очертание формы,

дельном случае равны нулю (когда устройства вообще нет, но необходимая функция выполняется).

3. В ИТР время обработки объекта приближается к нулю или равно нулю.

Определение ИТР. Будем считать техническое решение

1. В ИТР размеры ТО приближаются или совпадают

2. В ИТР масса и размеры ТО или его главных функ-

идеальным, если оно имеет одно или несколько из следую-

с размерами обрабатываемого или транспортируемого

объекта, а чистая масса ТО намного меньше массы обра-

циональных элементов приближаются к нулю, а в пре-

Рис. 68, Способ опредния плоскосимистричных и осесимыетрачных тел последовательный локальный спуск каждого решения (впачале грубый, затем более точный) происходит независимо от спуска других решений.

Конкуренция позволяет за счет отсева решений, спускающихся в локальные экстремумы, достаточно быстро находить глобальный экстремум в задачах, для которых значение функционала, осредненное по области пригяжения глобального экстремума, меньше значения функционала, осредненного по всей область приска, а область притяжения глобального экстремума не сдишком мала.

Алгоритм конкурирующих точек — один из наиболее простых и эффективных по сравнению с другими распространенными влюритмами поиска глобального экстремума. Так, например, трудоемкость поиска (заграты машинного времени) по этому алгоритму на порядок меньше по сравнению с алгоритмом случайного перебора локальных экстремумов и на два порядка меньше по сравнению с методом Монте-Карло [1, 17].

Для удобства изложения алгоритма решение будем называть также точкой (в многомерном пространстве поиска) и независимо от того, решается ли задача параметрической оптимизации (25)—(28) или задача структурной оптимизации (29)—(33), будем обозначать его X,

Алгоритм конкурирующих точек в общем виде включает следующие операции [1].

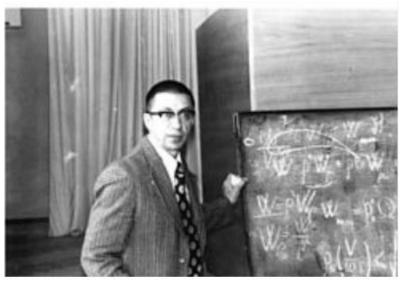
1. По процедуре СДС синтезируется t ($t = \eta + \lambda_0$) точек $\overline{X_j}$ (j = 1, ..., t), в которых определяется значение минимизируемой функции (критерия сравнения). Из этих t точек отбирается η точек, имеющих наилучшие значения критерия, которые в дальнейщем называются основными. Запоминается наихудшее значение критерия основных точек ϕ_0 . При этом считается, что совершен нулевой глобальный (групповой) шаг поиска (t = 0).

Таким образом, на *t*-м групповом шаге поиска имеем основные точки

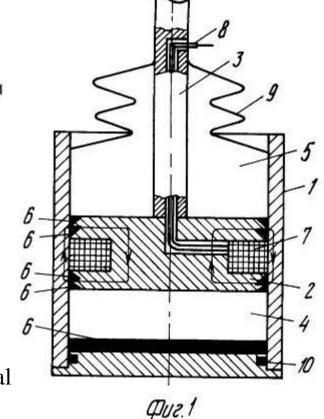
$$\overline{X}_1^t$$
, \overline{X}_2^t , ..., \overline{X}_{0}^t , (34)

и соответственно невозрастающую последовательность чисел Farber B. et al., Method of controlling the power element of the simulator Patent #2081643 (Bio-

controlled FEPOL)



$$B_1 \leftarrow \frac{\Pi_M}{MEX} B_2$$
 $MEX \rightarrow B_2$
 MAM
 $B_1 \leftarrow MEX \rightarrow B_2$
 $MEX \rightarrow B_2$
 $MEX \rightarrow B_2$
 $MAM \rightarrow B_2$
 $MEX \rightarrow B_2$
 $MAM \rightarrow B_2$
 MAM



Vadim Vladimirovich Gogosov-Professor, Doctor of Physical and Mathematical Sciences, Head of the Laboratory of Physical and Chemical Hydrodynamics of the Institute of Mechanics of M.V. Lomonosov Moscow State University



Introducing the magnetic scalar potential ϕ whose negative gradient equals the applied magnetic field, i.e. $\mathbf{H} = -\nabla \phi$, the scalar potential can be given by

$$\phi(x,y) = -\frac{I_0}{2\pi} \left(\tan^{-1} \frac{y+a}{x} + \tan^{-1} \frac{y-a}{x} \right),$$

where I_0 denotes the dipole moment per unit length and a is the distance of the line current from the leading edge. Then, the corresponding field components are given by

$$H_{x} = -\frac{\partial \phi}{\partial x} = -\frac{I_{0}}{2\pi} \left[\frac{y+a}{x^{2} + (y+a)^{2}} + \frac{y-a}{x^{2} + (y-a)^{2}} \right],$$

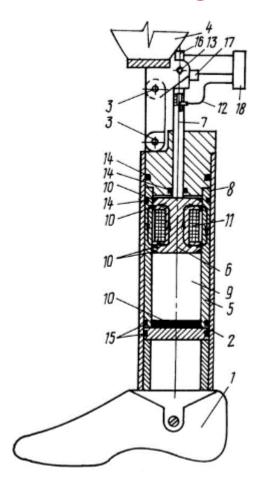
$$H_{y} = -\frac{\partial \phi}{\partial y} = -\frac{I_{0}}{2\pi} \left[\frac{x}{x^{2} + (y+a)^{2}} + \frac{x}{x^{2} + (y-a)^{2}} \right].$$

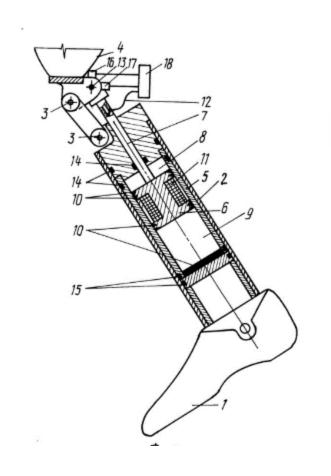
Moreover, the second derivatives are

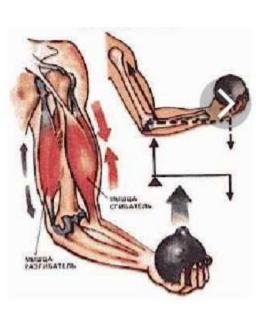
$$\frac{\partial^{2} \phi}{\partial x^{2}} = -\frac{\partial^{2} \phi}{\partial y^{2}} = -\frac{I_{0}}{2\pi} \left[\frac{2x (y+a)}{\left[x^{2} + (y+a)^{2}\right]^{2}} + \frac{2x (y-a)}{\left[x^{2} + (y-a)^{2}\right]^{2}} \right]$$

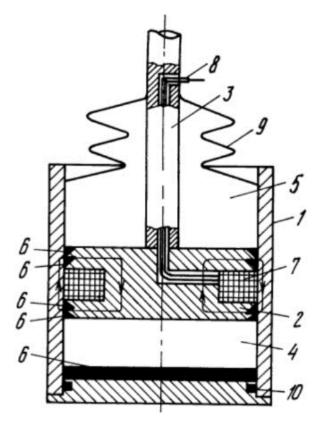
PART 4: Magnetic fluids in medicine

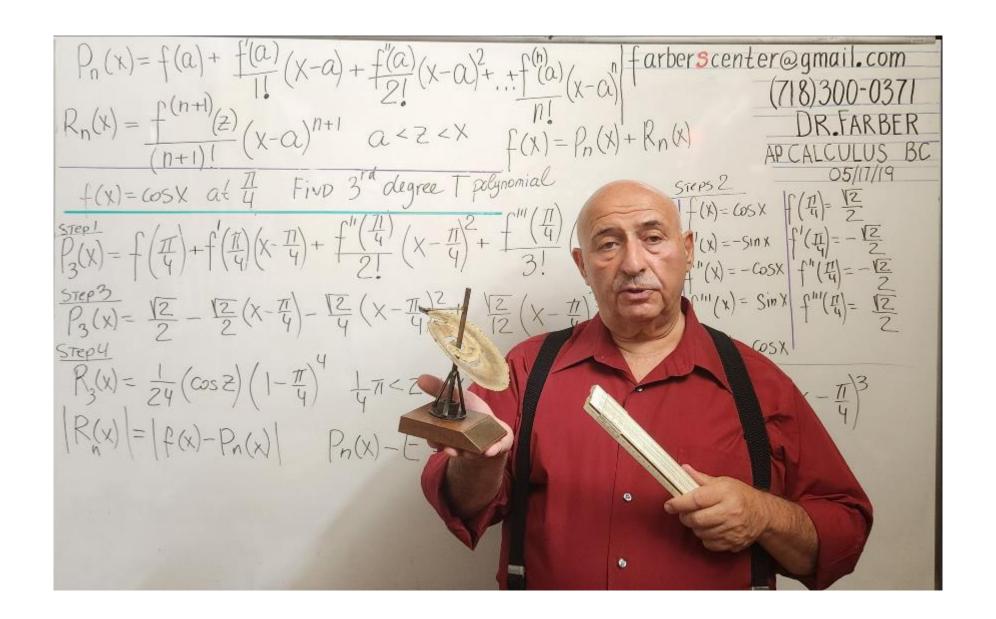
Farber B. et al., Patent 2032434, 1993 Magneto rheological device and method of control Farber B. et al., Rheomagnetic training device Patent 2081643,1993













Journal of Rehabilitation Research and Development Vol. 32 No. 4, November 1995 Pages 325-336

Biomechanical basis of choosing the rational mass and its distribution throughout the lower limb prosthesis segments

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https://www.rehab.research.va.gov/jour/95/32/4/pdf/farber.pdf

Abstract-A solution for finding a rational distribution of mass in lower limb prostheses has been considered based on the formal premise favoring the identification of the movements of a prosthetic and an intact leg. For the purpose of simplicity, an analysis has been carried out for only the swing phase, the data about the properties of moving segments being determined without integrating differential equations of motion. At the formation of equations of motion, an assumption that body segments are absolutely rigid and have constant moments of inertia and locations of the center of

$$rac{d}{dt}\left(rac{\partial L}{\partial \dot{ heta}}
ight) = rac{\partial L}{\partial heta}$$

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It should be noted that the application of the principle of mechanical similarity does not exclude other reasonable assumptions leading to different solutions. Thus, Godunov (2) assumed that the masses of both the prosthetic and sound limb segments must be in the same proportion. This opinion can be recognized as valid only for those prosthetic limb segments which do not involve a residual limb, such as the shin section and artificial foot of an above-knee (AK) prosthesis.

Some research papers provide a priori recommendations on the prostheses mass. Roschin and Delov (3) consider the optimum weight of an AK prosthesis to be within 2.7-3.6 kg and the weight of a below-knee (BK) prosthesis to be 2.3-2.9 kg. Staros (4) stresses in his paper that he considers it incorrect to automatically assume greater energy expenditures for heavier prostheses. He asserts that the work of hip muscles is influenced by the distribution of mass throughout the prosthesis: that is why it is necessary to raise the total center of mass location of a prosthesis by reducing the weight of its distal portion.

METHODS

We shall consider the problem of rational distribu tion of mass by means of the similarity method,

Differential Equations of Motion

Figure 1 presents a dynamic model of human walking in the form of a 9-link biokinematic chain with 11 degrees of freedom. Lower limb segments (hip, shank, fore and hind sections of feet) are presented by four links; trunk, neck, head, and arms (presented as one

Movable links rotate around the axes of a coupling O. with the centers of masses concentrated in the points C. The generalized coordinates are chosen to be X, Z, the horizontal and vertical displacements of the point O_i (the center of the hip joint) and \$\phi_i\$ (the angular displacements of the pelvic and lower limb links from the vertical axis).

While formulating the equations, the following assumptions were admitted: the body segments are absolutely rigid, the distribution of masses within each link is constant and does not depend either on muscular tension or spatial interrelation of the links, and the links have constant inertial moments and the positions of centers of masses Ji=Const., li=Const. The couplings between the links are stationary and the system is

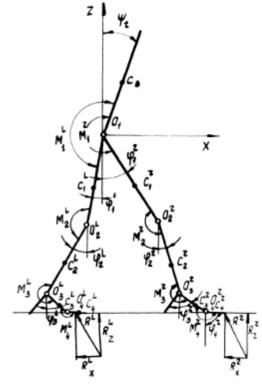


Figure 1. Dynamic model of human gait.

incapable of being integrated. The motion is possible due to the muscular forces, called joint forces, the moments of which are applied in the centers of rotation.

On the basis of the mechanical principle of releasing from couplings, the interaction of lower limbs with the support surface is replaced by the ground reaction, the components of which are marked R, and R. In that case, both R. and R. act as external forces. The differential equations of motion of the system are formulated using Lagrange II equations:

$$\frac{d}{dt}\left(\frac{\partial T}{\partial \dot{q}_i}\right) - \frac{\partial T}{\partial q_i} = Q_i$$

The motions of the suggested system are described by the second-order differential equations and in general can be presented as follows:

$$\begin{split} M_i &= M_{i+1} + \sum_{j=1}^4 \alpha_{ji} \left[\ddot{\varphi}_j Cos(\varphi_j - \varphi_i) + \varphi_j^2 Sin(\varphi_j - \varphi_i) \right] + \\ c_i \left[\ddot{x} Cos\varphi_i + \left(\ddot{z} + g \right) Sin\varphi_i \right] - D_i \left(R_z Sin\varphi_i - R_x Cos\varphi_i \right); \end{split}$$

$$\alpha_{ij} = \alpha_{ji} = \left(\frac{P_j}{g}l_i + \frac{\sum\limits_{k=i+1}^4 P_k}{g}L_j\right)L_{ii} D_i = \begin{vmatrix}L_1\\L_2\\L_R\\X\end{vmatrix};$$
 [1]

$$\alpha_{ii} = \frac{P_i}{g} \rho_i^2 + \frac{\sum_{k=i+1}^4 P_k}{g} L_i^2; c_i = \frac{P_i}{g} I_i + \frac{\sum_{k=i+1}^4 P_k}{g} L_i;$$

Here P_i = masses of segments;

 $L_i = lengths of segments;$

l_i = static radii;

p. = inertial radii of segments;

g = acceleration of the gravity force;

φ_i = generalized coordinates;

R_x, R_Z = horizontal and vertical ground reaction components;

x, y = load point coordinates.

The equations presented above can be considered as the formulae determining the moments of muscular forces, having the data on generalized coordinates, changes, ground reaction, and its load point. The necessary information can be obtained by an experimental method.

The Application of the Mechanical Similarity

Let us assume that each system of equations is recorded twice for two models, one of them imitating the motion of the sound limb (index "c"), the other, that of the prosthetic limb (index "p").

In that case, the main requirement of mechanical similarity (i.e., the requirement of the formal identity of equations of motion for both models will be observed) is met, provided that

$$x_p = x_{ci} z_p = z_{ci}$$

 $\frac{\alpha_{ijp}}{\alpha_{iir}} = \frac{C_{ijp}}{C_{iir}} = \frac{M_{ip}}{M_{ir}}$, [2]

where K=Const, a_{ij} = C_{ij} = M_i are the coefficients of equations, the joint moments,

To prove the identity of equations, it is sufficient to make replacements in the equations for model II

$$\alpha_{iip} = k\alpha_{ik}$$
, $C_{iip} = C_{ik}$, $M_{ip} = M_{ic}$ [3]

to obtain the equations for model I.

The conditions of Equation 1 make it possible to work out a series of independent proportions, which are reduced to the system of three equations with six unknown quantities by means of using biomechanical constants (5).

It should be noted that the coefficients of the system of equations in Equation 1 present the combinations of inertial and geometric characteristics of the human body limb segments.

The results of numerous investigations in the fields of anthropology, anatomy, and biomechanics made it possible to establish regular relations between characteristics of separate segments and the human body as a whole (6.7). It is important to note that at the present time the investigations in this field are carried out to meet the needs of not only prosthetics manufacturing but robotics technology, aviation, and space medicine as well. Therefore, the following parameters are well known: the relative linear dimensions of the human body segments, expressed in relative units "P" (where "P" is 1/56 of human height); the relative masses of segments (the mass of the whole body = 100 percent); the coordinates of mass centers (the human height = 100 percent); and the coordinates of joint centers in percentages of the human height and the proximal joints. The positions of partial mass centers and the value of inertial radii of limb segments measured from a proximal joint are also determined. For the first time the calculations of the mass, the partial mass centers, and the inertial radii of a lower residual limb for different levels of amputation were carried out. It was performed using the approximation method with bodies of rotation recommended by reference books on mechanics (8).

Distribution of Masses Throughout a Prosthesis

The independent proportions obtained from conditions (9) are recorded as the system of equations solved in relation to P_2 , l_2 , p_2 (i.e., the mass, the static radius, and the inertial radius) related, respectively, to the human body mass P and the link length L. Note that the derived system of three equations has six unknown quantities, three of them having been chosen the mean as the result of a considerable number of measurements: 228

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$$K_1 = \frac{P_{sb}}{P} = 0.016; K_2 = \frac{I_{sb}}{L_1} = 0.0598;$$

$$K_3 = \frac{P_{sb}}{L_1} = 0.662,$$
[4]

where P_{sb} , l_{sb} , ρ_{sb} are the mass, the static radius, and the inertial radius of the AK socket, related, respectively, to the human body mass P and the hip length L_1 . These amputations are described by the following equations:

$$p = \left(\frac{l_2}{L_2}\right)^2 = 0.310 \left[\frac{0.010 + K_4 K_5}{3.330 K_4 \left(K_5 - 1.30 K_6^2\right)} + 1\right];$$

$$r = \frac{P_2}{P} = 3.330K_4(K_5 - 1.30K_6^2);$$
 [5]

$$\rho^2 = \left(\frac{\rho_2}{L_2}\right)^2 = 0.731 \frac{l_2}{L_2},$$

where P_2/P , I_2/L_2 , p_2/L_2 are the mass, the static radius, and the inertial radius of the complex chain "shank + foot + footwear" related, respectively, to the human body mass P and the distance "knee-floor" L_2 .

$$K_4 = \frac{P_{bb}}{P}$$
; $K_5 = \frac{l_{bb}}{L_1}$; $K_6 = \frac{\rho_{bb}}{L_1}$, [6]

where P_{kb} , l_{kb} , p_{kb} are the masses, the static radius, and the inertial radius of the AK residual limb, related, respectively, to the human body P and the hip length L_1 .

It should again be stressed that the mean values of the AK socket inertial characteristics replaced in Equation 5 obviously do not depend on the level of amputation, for whatever the length of the residual limb may be, the AK socket depends only on the sound limb hip length, that is, on the human height.

However, it should be taken into account that a purposeful striving for reducing the mass of modern prosthetic parts is not an end in itself, but a method of rational distribution of mass with the help of balancing, for example. Balancing is possible in the case where the actual weight of a complex chain shank + foot proves to be less than the calculated weight.

Balancing is performed in the following way. First the mass of the patient, the level of amputation, and the distance knee-floor L_2 are measured. Then, with the help of a simple device, the scheme of which is given in Figure 2, the actual masses of a complex chain P_F and its static moment M_F are determined. The calculated mass (P_p) , the static radius (l_p) , and the calculated static moment (M_p) are determined with the help of nomograms (Figures 3–5) by the level of amputation, the mass, and the distance "knee-floor" of the patient. The nomograms have been worked out according to the known mathematic rules in logarithmic coordinates on the basis of Tables 1 and 2.

The value of a balancing load and the position in which it is fixed are determined by the following formulae:

$$\Delta P = P_o - P_F;$$
 [7]

$$l_0 = \frac{M_p - M_F}{\Lambda P},$$
 [8]

where the numerator presents the difference between the calculated and the actual static moments and the denominator shows the value of a balancing load. Usually strips of sheet lead 5-6 mm thick are used as a balancing load. A strip of lead plate, the mass of which is equal to the calculated value of a balancing load, should be riveted to the rear surface of a shank tube at the point, located at the distance l_0 from the knee joint axis.

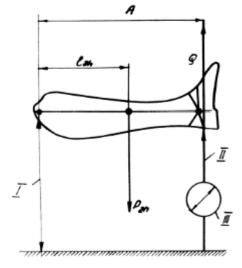
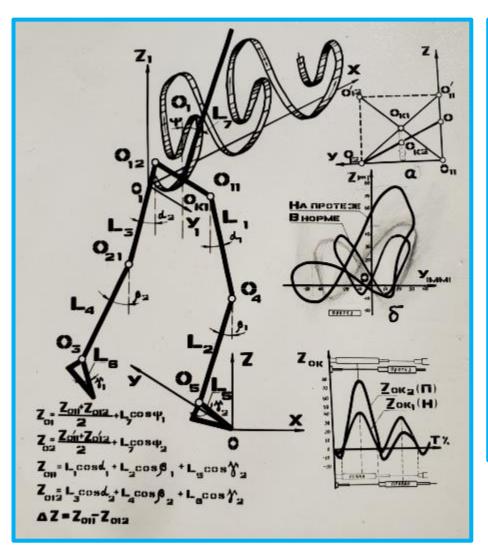
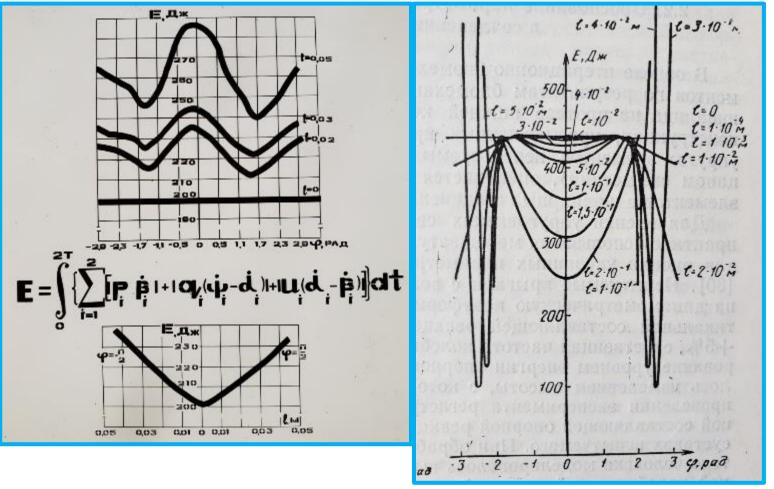


Figure 2. Scheme of a device for determining the position of a center of mass of a chain.





Farber B. et al., Biomechanical basis of choosing the rational mass and its distribution throughout the lower limb prosthesis segments

https://www.rehab.research.va.gov/jour/95/32/4/pdf/farber.pdf

• Journal of Rehabilitation Research and Development Vol. 32 No. 4, November 1995 Pages 325-336

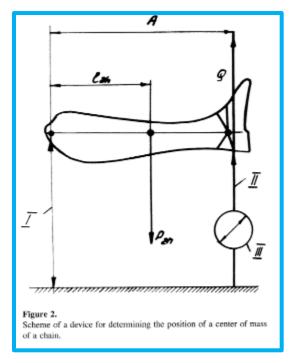


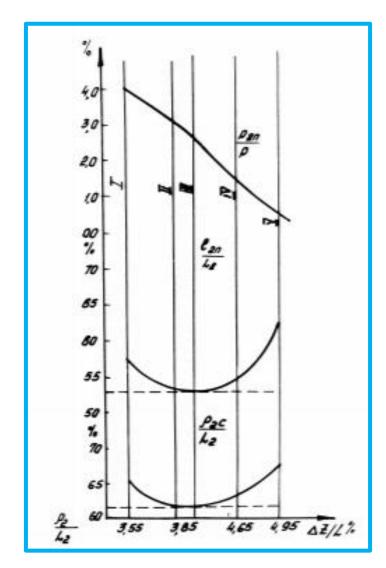
Figure 7.

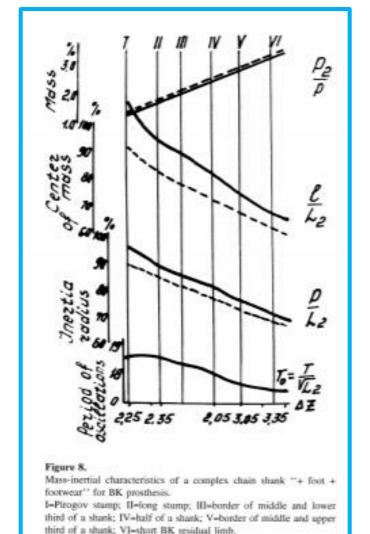
Mass-inertial characteristics of a complex chain shank "+ foot + footwear" for AK prosthesis.

P, is the mass as a percentage of the human body mass; static radius

P₂ is the mass as a percentage of the human body mass; static radius 12 and inertial radius are expressed as a percentage of the distance "knee-floor" L2. On the X-coordinate is the frontal displacement of the total center of masses (TCM) related to the man's height.

I-Gritti stump; II-long stump; III-border of lower and middle third of a hip; IV-half of a hip; V-border of middle and upper third of a hip.





S- curve for Prothesis Knee Units

АКАДЕМИЯ НАУК СССР СИБИРСКОЕ ОТДЕЛЕНИЕ

Институт истории, филологии и философии СО АН СССР

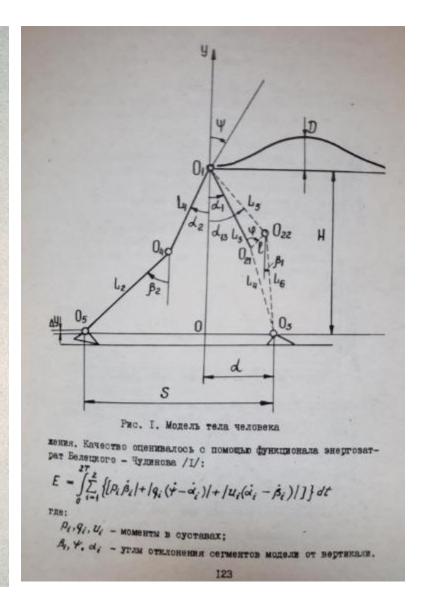
> Философское общество СССР Западно-Сибирское отделение

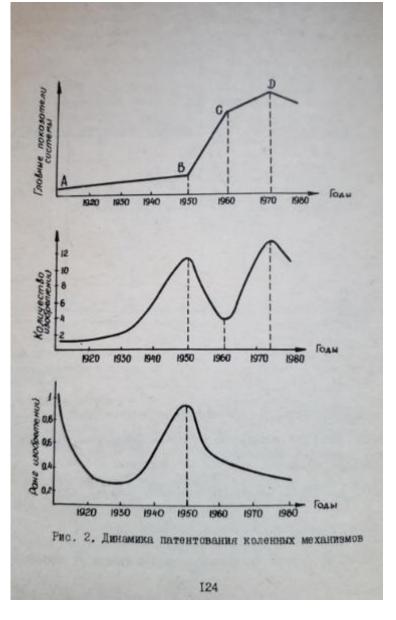
Новосибирский государственный университет им Ленинского комсомола

МЕТОДОЛОГИЯ И МЕТОДЫ ТЕХНИЧЕСКОГО ТВОРЧЕСТВА

Тезисы докладов и сообщений к научно-практической конференции 30 июня — 2 июля 1984 г.

Новосибирск 1984





Член-корреспондент РАН, Профессор, Др Владимир Белецкий



World-known Scientist in the field of celestial mechanics, dynamics, and astronautics, author of works on the theory of rotational motions of artificial and natural celestial bodies. Corresponding Member of the Russian Academy of Sciences, full member of the International Academy of Cosmonautics and the Russian Academy of Cosmonautics. K. E. Tsiolkovsky Institute of Applied Mathematics. Keldysh RAS, Moscow State University

Official opponents: Doctor of Physical and Mathematical Sciences, Professor.

V. Beletsky

Doctor of Medical Sciences, Professor X. A. Janson

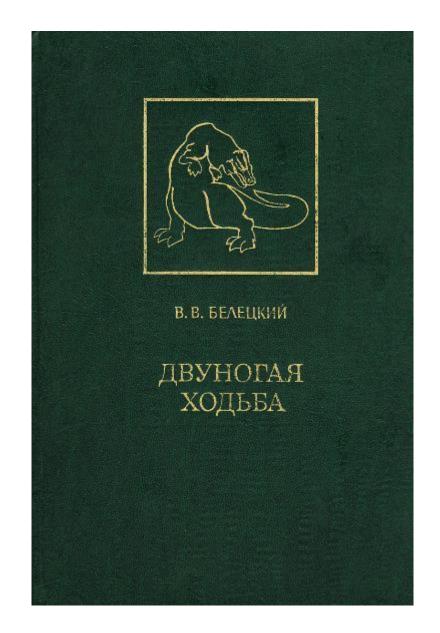
Doctor of Biological Sciences, Professor A. V. Zinkovsky

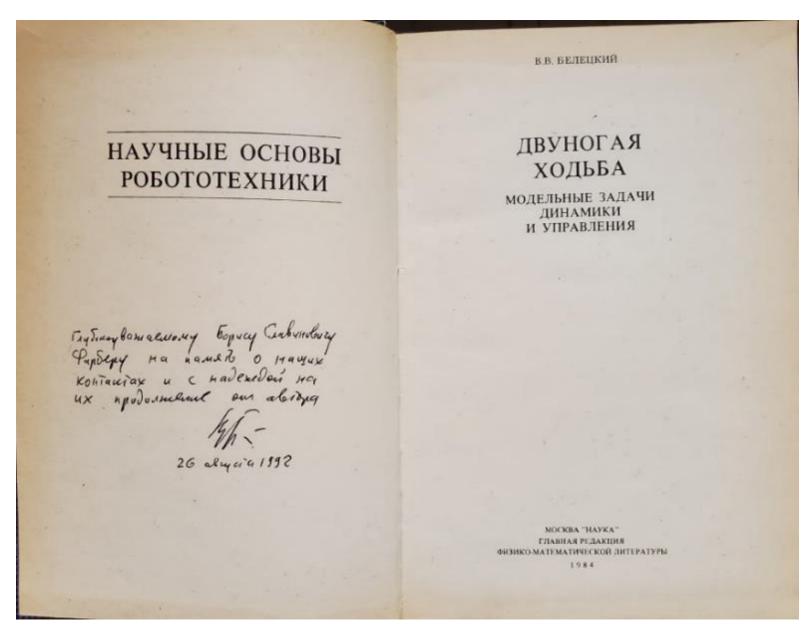






Dr. Professor V. Beletsky Bipedal Walking



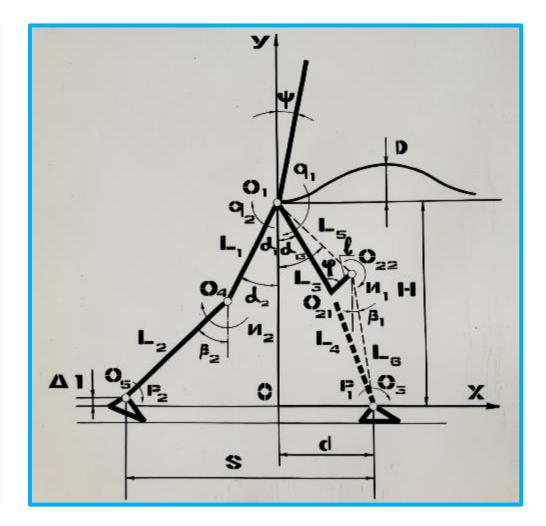






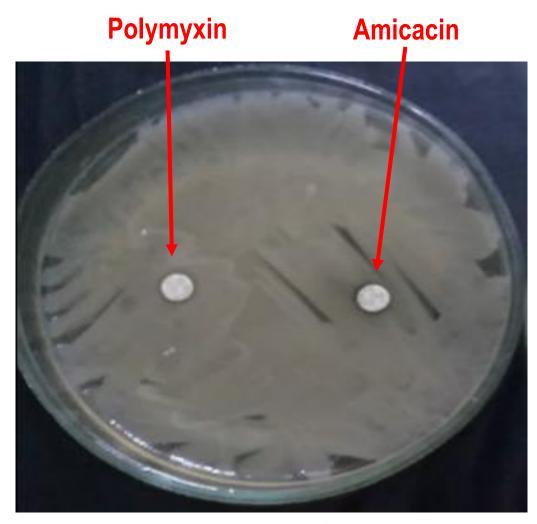
Dr. Farber's Dynamic Mathematical Model

 $M\ddot{x} + (m_1 + 2m_2)L_1'(\ddot{d}_{22}\cos d_2 - \dot{d}_2^2\sin d_2) + (m_5 + 2m_6)L_5'(\ddot{d}_{13}\cos d_{13} - \dot{d}_{13}\sin d_{13}) +$ $+ m_2 L_2' (\ddot{\beta}_2 \cos \beta_2 - \dot{\beta}_2^2 \sin \beta_2) + m_6 L_6' (\ddot{\beta}_1 \cos \beta_1 - \dot{\beta}_1^2 \sin \beta_1) = R_{1x};$ $M\ddot{y} + (m_1 + 2m_2)L'_1(\ddot{d}_2\sin d_2 + \dot{d}_2\cos d_2) + (m_5 + 2m_6)L'_5(\ddot{d}_{13}\sin d_{13} + \dot{d}_{13}\cos d_{13}) +$ + $m_6L_6(\ddot{\beta}_1\sin\beta_1+\dot{\beta}_1^2\cos\beta_1)=R_{19}-Mg;$ $(J_1 + 4m_2L_1^2) \ddot{d}_2 + 2m_2L_1L_2 \ddot{\beta}_2 \cos(d_2 - \beta_2) + (m_1 + 2m_2)L_1 (\ddot{x} \cos d_2 + \ddot{y} \sin d_2) +$ $+2m_2L_1L_2\beta_2^2\sin(d_2-\beta_2)+(m_1+2m_2)gL_1\sin d_2=q_2-u_2;$ $(J_5 + 4m_6 L_5)d_{13} + 2m_6 L_5 L_6 \ddot{\beta}_1 \cos(d_{13} - \beta_1) + (m_5 + 2m_6)L_5 (\ddot{x} \cos d_{13} + \ddot{y} \sin d_{13}) +$ $+2m_6L_1L_2\dot{\beta}_1^2\sin(d_{13}-\beta_1)+(m_5+2m_6)gL_5^2\sin d_{13}=q_1-u_1+(R_{1x}\cos d_{13}+R_{1y}\sin d_{13})2L_5^2$ $J_2 \ddot{\beta}_2 + 2m_2 L_1 L_2 \dot{\alpha}_2 \cos(\dot{\alpha}_2 - \beta_2) + m_2 L_2 (\ddot{x} \cos \beta_2 + \ddot{y} \sin \beta_2) - 2m_2 L_1 L_2 \dot{\alpha}_2 \sin(\dot{\alpha}_2 - \beta_2) +$ + $m_2 g L_2 \sin \beta_2 = u_2$; $J_{6}\ddot{\beta}_{1}+2m_{6}L_{6}\dot{L}_{5}\dot{d}_{13}\cos(d_{13}-\beta_{1})+m_{6}L_{6}(\ddot{x}\cos\beta_{1}+\ddot{y}\sin\beta_{1})-2m_{6}L_{6}L_{5}\dot{d}_{13}\sin(d_{2}-\beta_{2})+$ $+ m_6 g L_6 \sin \beta_1 = u_1 - p_1 + 2 L_6 (R_{1x} \cos \beta_1 + R_{1y} \sin \beta_1);$ $J\ddot{\psi}-M_7L_7(\ddot{x}\cos\psi+\ddot{y}\sin\psi)-M_7gL_7\sin\psi=-q_1-q_2$,

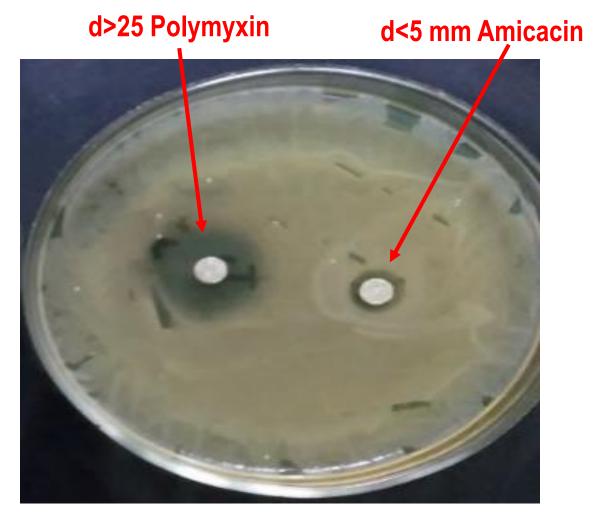


Application 4 : In molecular biology

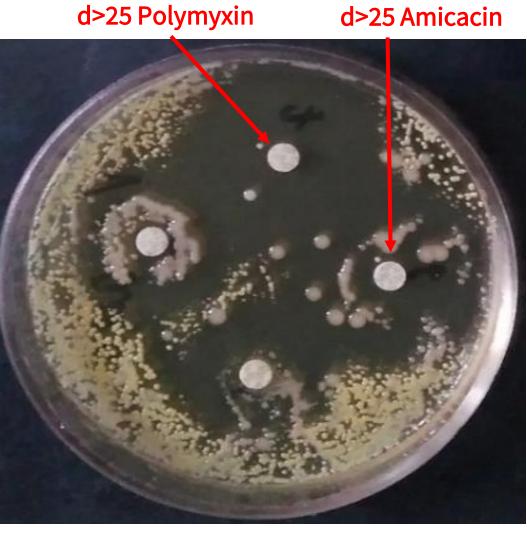
- Differential equations are of basic importance in molecular biology because many biological laws and relations appear mathematically in the form of a differential equation.
- The vast majority of quantitative models in cell and molecular biology are formulated in terms of ordinary differential equations.
- Mathematical cell biology is a very active and fast growing interdisciplinary area in which mathematical concepts, techniques, and models are applied to a variety of problems in developmental medicine and bioengineering.



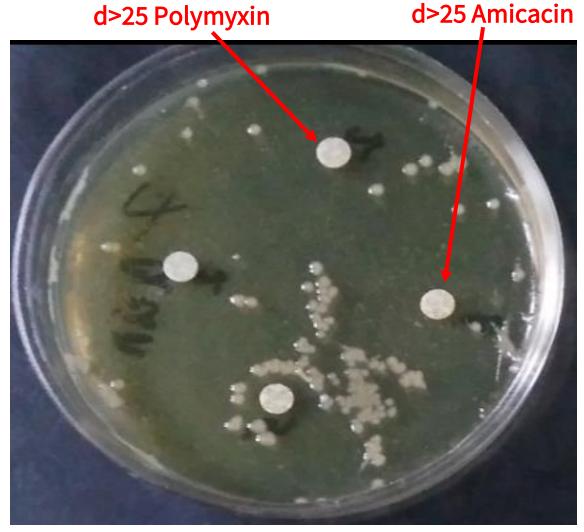
MDR *A.baumannii* growth without NGL025: 6 day growth, 2nd passage.



MDR *A.baumannii* growth with NGL025: 6 day growth, 2nd passage.



MDR *A.baumannii* growth with NGL025: 9 day growth, 3rd passage.



MDR *A.baumannii* growth with NGL025: 12 day growth, 4th passage.

The Key to NOIGEL's breakthrough

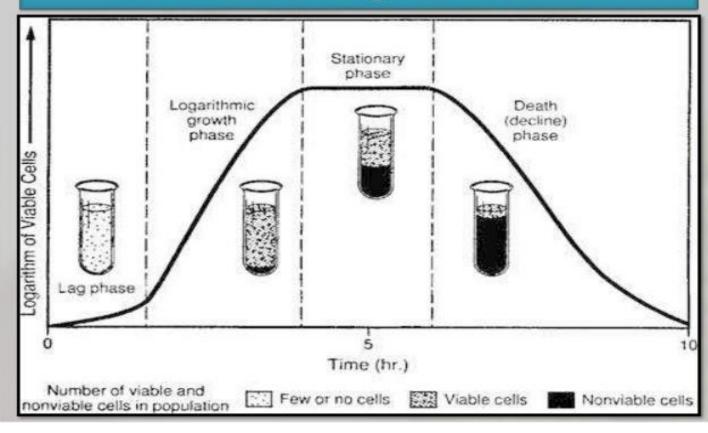
Logarithmic growth-

bacteria in the absence of competition with each other "dump" the majority of virulence factors and toxin formation (including factors of acquired antibiotic resistance)

Log phase -

The most perspective phase as a target by antimicrobials.

The microbial growth curve

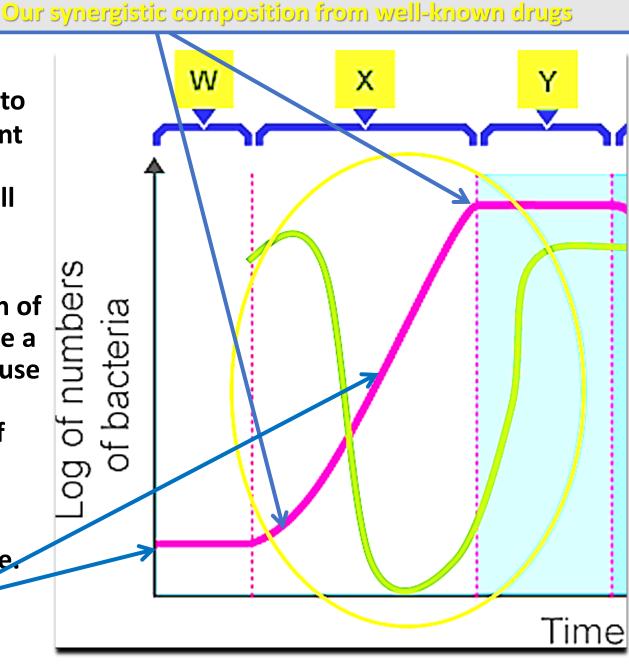


López, S., Prieto, M., Dijkstra, J., Dhanoa, M. S., & France, J. (2004). Statistical evaluation of mathematical models for microbial growth. International Journal of Food Microbiology, 96(3), 289-300

What ARE STRATEGIES?

- Our strategy is to"fool" bacteria in order to eliminate biofilms and to destroy multiresistant bacteria by sending false signals making them think that there is an absence of danger. It will bring to initiation of logarithmic phase of bacterial growth.
- It is well known fact that the intensive growth of the bacterial mass in Log-phase does not cause a release of more toxins, virulence factors or cause a formation of biofilm.
 - Based on this phenomenon, a combination of well known medications and compositions expressed in synergistic effect to stimulate growth of bacteria we were able to find a method to stop bacterial toxicity and virulence.

long-known classical antibacterial



Application 1 : Exponential Growth - Population

- One of the most basic examples of differential equations is the Malthusian Law of population growth $\frac{dp}{dt}=rp$ shows how the population (p) changes with respect to time. The constant r will change depending on the species.
- More complicated differential equations can be used to model the relationship between predators and prey. For example, as predators increase then prey decrease as more get eaten. But then the predators will have less to eat and start to die out, which allows more prey to survive. The interactions between the two populations are connected by differential equations.

Population Dynamics of a Continuous **Propagator for Microorganisms**

R. K. FINN and R. E. WILSON¹ University of Illinois, Urbana, Ill.

As continuous fermentation offers economic advantages over the usual batchwise process, the present work was undertaken to provide a better understanding of the characteristics of a continuous propagator. Laboratory apparatus was developed which allowed sterile propagation of aerobic microorganisms in a stirred-tank reactor of the overflow type. Using Ps. fluorescens, B. linens, and a strain of S. carlsbergensis a theoretical relation was shown between the mean retention time of the cells in the propagator and their growth rate. By proper adjustment of the flow rates, steady populations were attained. In incompletely buffered media, a steady cycling in yeast population was observed, which was traced to steady fluctuations in the pH which were 90 degrees out of phase with fluctuations in population. The phenomenon appears to arise from the inherent feedback in the system coupled with a metabolic lag. Fermentation of the medium was not complete, as only the propagation of cells was of interest. In a practical process additional holding tanks would be provided, so that end products could be obtained in high yield.

MOST STUDIES OF CONTINUOUS FERwith producing consistently high yields of alcohol, yeast, or other specific materials (3, 77, 79). It is appropriate that academic research in the field of bioengineering should undergird these applied studies with a more detailed inquiry into the behavior of living cells held in continuous culture. The purpose of this paper is to begin such an inquiry.

When fermentation is carried out in a cascade of two or more stirred tanks connected in series, the first tank constitutes a propagator. Fresh nutrient is added to the propagator at constant rate and a constant level of liquid is maintained in the propagator by arranging an overflow at the desired height. The rate of withdrawal is at all times equal to the feed rate and is generally so rapid that fermentation within the propagator is not complete, and all nutrients remain in excess of the cell requirements. Under such conditions, the cell population is limited solely by the washout of cells.

Novick and Szilard (74, 15) utilized a continuous propagator to study microbial genetics, especially mutation rates. Their apparatus, which they chose to call a chemostat, was operated in such a manner that growth was limited by

deficiency of a particular nutrient, rather than by the washout rate.

The present work follows more closely that of Adams and Hungate (7). Using a continuous yeast propagator these investigators showed how the flow rate could be predicted from the growth curve of the organism. Several points raised in their paper, however, seemed to warrant further study. In the first place, Adams and Hungate did not observe any appreciable constant-rate phase of growth for yeast in their media. With organisms that show logarithmic growth, the estimation of flow rates could be simplified.

Furthermore, steady populations were not always obtained in the yeast propagator. Although Adams and Hungate (1) did not call attention to the possibility of cyclic fluctuations, their data suggested such an occurrence. More recently Maxon and Johnson (10) have confirmed the cycling phenomenon in a yeast propagator.

The present research concerns the propagation in the logarithmic phase of two aerobic bacteria, Bacterium linens and Pseudomonas fluorescens; cycling was also investigated, using a strain of Saccharomyces carlsbergensis.

Theory of Continuous Propagation With Logarithmic Growth

In batchwise cultivation single-celled organisms usually exhibit a phase of logarithmic growth which can be characterized by the equation

$$\frac{dN}{d\theta} = kN \qquad (1)$$

in which

N =the total number of cells

= a characteristic growth constant for the organism, which depends also on the environmenttemperature, nature of medium, etc. Units are reciprocal time.

The rate constant, k, is conveniently measured by the slope of the growth curve when plotted on semilogarithmic coordinates.

$$k = \frac{\text{(slope)}}{2.303}$$
(2)

To achieve constant population in a flow system, the retention time within the propagator must equal 1/k. Most investigators have been content simply to state this fact, but Monod (12) has provided rigorous proof based on a differential material balance.

By a material balance on the total cells, making use of Equation 1,

$$(In + growth) - out = accumulation$$

$$0 + kNd\theta \sim q \frac{N}{V} d\theta = dN$$
 (3)

in which V is the volume of liquid in the propagator and o is the volumetric flow rate.

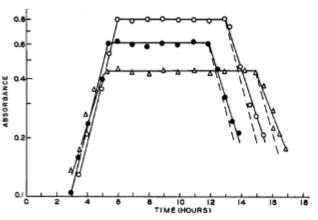


Figure 1. Verification of Equation 4 by substituting taxic reagents for nutrients being fed

Three separate tests are shown. Substitution of phenol begun at 11.5 hours, of peroxide and sulfuric acid at 12.5 hours.

- Ps. fluorescens. 6,. 1.5 hours. Temperature, 23° C. O Saturated phenol
- 30% hydrogen peroxide
- △ Concentrated sulfuric acid
- ... Theoretical washout rates, Equation 4

From Equation 3 the rate of change of population is

$$\frac{dN}{d\theta} = N(k - q/V) \qquad ($$

An alternate form of Equation 4 is

$$\frac{dN}{d\theta} = N \left(\frac{\ln 2}{\theta_*} - \frac{7}{\theta_*} \right) \qquad (4a)$$

where θ_s is the familiar generation time of the organism, and g, is the mean retention time in the apparatus.

For the population to remain constant in a continuous propagator, the flow rate must be set so that

$$\theta_r = \frac{1}{k} = \frac{\theta_{\ell}}{\ln 2}$$
(5)

If this condition is not met, the level of growth will change with time in accord with Equation 4.

It is assumed in the above analysis that flow rates are truly constant, and that mixing within the propagator is instantaneous and complete. The last assumption is generally not so severe a restriction as might be supposed (9).

Experimental

The propagator itself was not of unusual design. It consisted of a 500-cc.

gas-scrubbing bottle appropriately modified to allow for inoculation and the subsequent feeding of nutrients. The overflow was of capillary glass tubing. The entire propagator was immersed in a water bath held at constant temperature, and there was provision for blowing sterile air at measured rates through a sintered-glass disk within the propagator. Dissolved oxygen, measured polarographically, was always in excess of the critical concentration for cellular uptake. No auxiliary agitation was required because of the high air rates used.

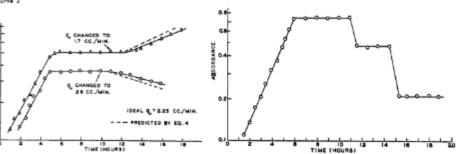
Of more interest to those wishing to study continuous fermentation was the use of a Sigmamotor pump (E and M Enterprises, Middleport, N. Y.) to provide constant flow rates of sterile nutrient. This pump has a cam arrangement, which moves fingerlike pieces of metal so as to press fluids through rubber or Tygon tubing, which passes between these fingers and a metal plate within the pump. Positive pumping at flow rates as low as 2 cc. per minute is easily possible. Changes in the flow rate may be made either by using rubber tubes of different size, or by modifying the speed of the pump. Freedom from contamination and ease of control recommend the use of such a pump over any variations of the Mariotte bottle as a means of attaining constant

Growth was followed by measurements of absorbance (optical density), using a Coleman Nephocolorimeter (red filter No. 8-215). The linear response of the instrument was checked by plate counts, and when necessary dilutions were made before taking absorbance readings. The pH measurements were made with a Beckman line-operated meter. Sugar and carbon dioxide were measured by standard analytical methods.

The operating procedure was simple. To start the propagator, growth was allowed to proceed batchwise until, at an appropriate level of population,

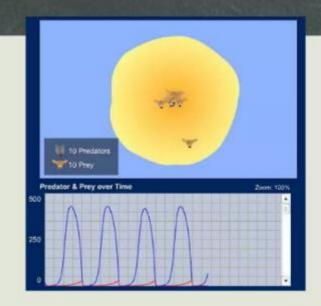
Figure 2. Verification of Equation 4 by altering flow Figure 3. Verification of Equation 4 by altering population level rate in two tests 8. litenz. Temperature 26° C. R., 2.4 hours

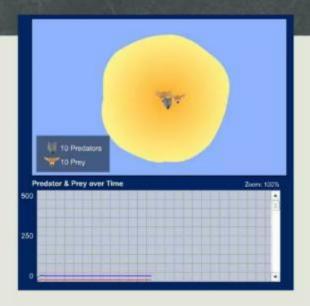
B. liners. Temperature 26° C. △ Curve 1 O Curve 2



³ Present address, Corn Products Refining Co., Argo, Ill.

Graph of Growth Population



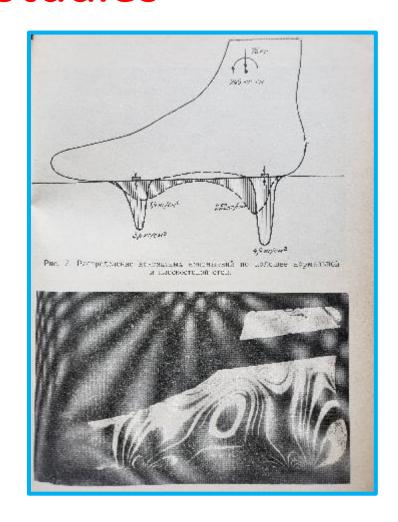


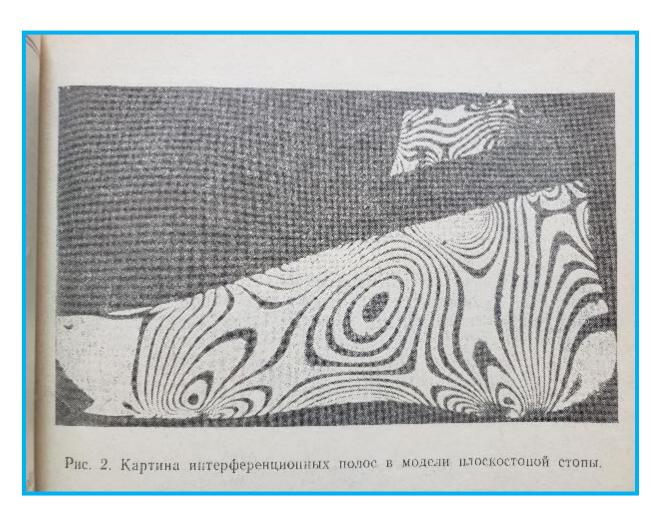
- First Graph shows when the predator is both very aggressive it will attack the prey and rapidly eat the prey population, growing rapidly before it runs out of prey to eat and then there will be no other food, thus dying off again.
- Then predators will be less aggressive and it will lead to both populations in a stable position (Graph two).

Some Other Applications of Differential Equations

- Some Other Applications of Differential Equations are,
 - 1) In medicine for modelling cancer growth or the spread of disease
 - 2) In engineering for describing the movement of electricity
 - 3) In chemistry for modelling chemical reactions
 - 4) In economics to find optimum investment strategies
 - 5) In physics to describe the motion of waves, pendulums or chaotic systems.

Farber B. et al., Photo elasticity for Foot Studies





Grand Idea **LINE - 2**

NOVEL Drugs with dynamic structures (Dynamic drugs) based on TRIZ





To survive, rate of Pathogen Microorganisms "innovations" faster than rate of new classical drugs development. This is time to find another, Dynamic way to fight Patogen Microorganisms

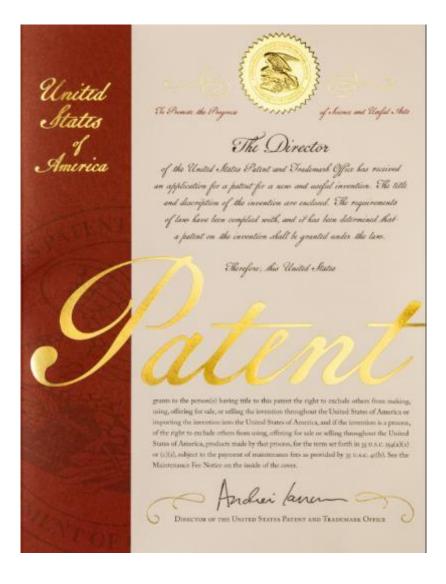
Illustration

Instead of one "key" for one "lock" (the principle of a classic drug with a conservative structure), we propose a selection of "skeleton keys": a group of many similar molecules that "open" many "locks" and adapt to the target. This facilitates a practically 100% effectiveness rate and a maximally wide spectrum of drug activity

Вместо одного «ключа» на один «замок» (принцип классического препарата с консервативной структурой) мы предлагаем набор «скелетных ключей»: группу из множества одинаковых молекул, которые «открывают» множество «замков» и адаптируются к цели. Это обеспечивает практически 100% эффективность и максимально широкий спектр действия препарата.

https://zenodo.org/record/2547580#.XGVsUIVKiWE

https://zenodo.org/record/2639505#.XLMwluj7SUk



Annals of Mechnikov Institute, N 4, 2018 www.imiamn.org.ua/journal.htm 15

CREATION OF NEW MEDICAL DRUGS BASED ON TRIZ AND COMPUTER MATHEMATICAL MODELING

Farber B.S., Martynov A.V., Kleyn I.R

Noigel, LLC; TRIZ Biopharma International, LLC New York; Mechnikov Institute of Microbiology and Immunology (Kharkov, Ukraine).

> In memory of Genrich Altshuller, creator of TRIZ philosophy and a Teacher, with whom we discussed the most significant of our trends and inventions.

TRIZ (the theory of inventive problem solving) is a new philosophy of thinking, created by Genrich Altshuller and further developed by his followers [1]. One of the authors area which affects almost every family, and each and every one of us. Namely, the development of new effective drugs, [12].

To use TRIZ principles in drug development necessitates broad knowledge in several areas rooted in the molecular modeling method. This method includes the application of the laws of quantum physics and quantum chemistry. Additionally, it requires knowledge of the behavior of molecules in various situations and their interaction with each other at different temperatures, as well as in the presence of salts and other compounds.

Truly effective drugs can be developed only on the basis of a systematic approach and in-depth knowledge of the fields of medicine; pharmaceutical chemistry, medical chemistry, physical chemistry, analytical chemistry, pharmacognosy, chemistry of natural compounds; plant medicine technology; biochemistry, molecular biology; pharmacology; and many other disciplines.

Over the last 100 years the pharmaceutical sciences

Annals of Mechnikov Institute, N 1, 2019

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APPLICATION OF SYNERGETIC SET OF TRIZ PRICIPLES FOR DEVELOPING cAMP -ACCUMULATION ACTIVATORS AND THEIR INFLUENCE ON MULTI-DRUG RESISTANCE MICROORGANISMS

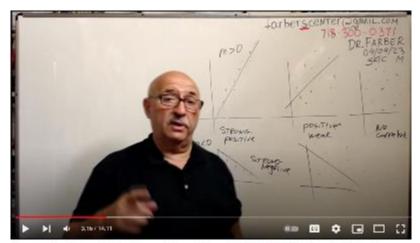
Farber B ²., Martynov A.^{1,2}, Osolodchenko T¹., Kleyn I². 47

out the reverse action. For example, a burn can be attained not only from extreme heat, but also from extreme cold, and expansion process can occur not only by heating, but also by freezing water. Overcoming psychological inertia allowing you to use the opposite action sometimes allows you to find novel solutions. In our case this would mean that instead of killing bacteria we should enhance them.

TRIZ principle of "Preliminary anti-action." (#9).This means that when you know that an undesirable situation

The connection between SAT problem and Cancer Diagnostics Blood Cells Morphometric Method

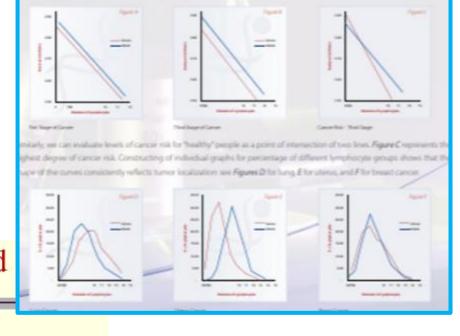
https://www.youtube.com/watch?v=XelQsJHz49g&t=634s&ab_channel=Farber%27sCenter

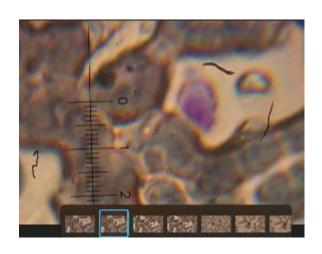


The connection between SAT problem and Cancer Diagnostics Blood Cells Morphometric Method

Blood Cell Morphometric Method

Dr. Valentin Govallo, Dr. Evgenia Skobeltzin, Dr. Boris Farber







Ph.D., D.SCI. Member of the Russian Academy of Medicine









METHOD FOR DIAGNOSTICS OF HUMAN TUMOR DISEASES Blood Cell Morphometric Method (BCMM) 2005 PCT WO 2007/021211

(12) МЕЖДУНАРОДНАЯ ЗАЯВКА, ОПУБЛИКОВАННАЯ В СООТВЕТСТВИИ С ДОГОВОРОМ О ПАТЕНТНОЙ КООПЕРАЦИИ (РСТ)

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- (51) Международная патентная классификация: G01N 33/49 (2006.01)
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- (22) Дата международной подачи:

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Русский

- (71) Заявители и
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- (81) Указанные государства (если не уназано иначе, для намедого вида национальной охранье): AE, AG, AL,

- AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.
- (84) Указанные государства (если не указано иначе, для кажедого вида регномальной охраны): ARIPO (ВW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW), евразийский (АМ, АZ, ВУ, КG, КZ, MD, RU, ТЈ, ТМ), европейский питент (АТ, ВЕ, ВG, СН, СУ, СZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR), OAPI (ВF, ВЈ, СF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Лекларания в соответствии с правилом 4.17:

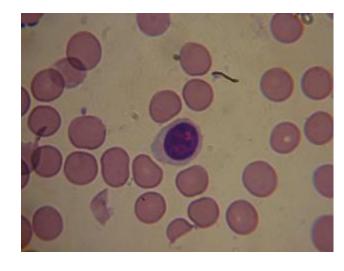
об авторстве изобретения (правило 4.17 (iv))

Опубликована:

с отчётом о международном поиске

В отношении двубуквенных кодов, кодов языков и других сокращений см. "Повснения к кодам и сокращеники", публикуемые в начале каждого очередного выпуска Бюзлетеня РСТ.



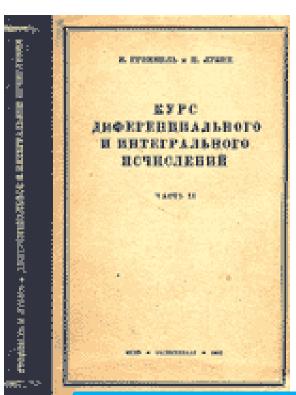


Grenville, W.A., Luzin, N.N. Course of differential and integral calculus ("LUSITANIA").





LUZIN Nikolai Nikolaevich (1883-1950) - Russian mathematician, founder of the scientific school of function theory, academician of the USSR Academy of Sciences





АКАДЕМИК КОНСТАНТИН ФРОЛОВвибротехнические и биомеханические системы



Советский и российский ученый в области машиностроения. Вицепрезидент Российской академии наук, Академии наук СССР.



Security (security) property. Managery received resemble desired

АКАДЕМИКУ К.В. ФРОЛОВУ

75 **JET**

Academic Accommunity Secretaries Species - melanassics premai c napoних ежиси, высовный упурявлени и об-ASCERN SPURIORIST & PROMODERAN, MORRO-Activate moleculated a first automorphism. paralament mayrour representativements a sername Proces, evacuoscoprare year детральных вархом у направлений, марко изировые и присменных и лежей стрике и м дебення. Он возниться опинежники of the contraction of the second of the seco parameter IN seco. use AA Sucception, P.M. donolaywasi, & O. Eccusion C.S. Dydeson a smore discus-



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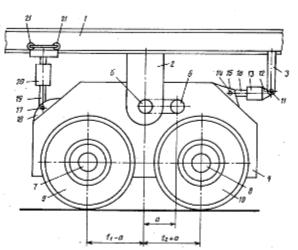
Enterprise and the property of the second se that the set I from the Principle and the second control of the second s The company of the co control tradema primari forti characte. A spinore interest which there are no perform a force property in the conchapter recent a common males entrace. I T.S. Storma reports from a reconstruction

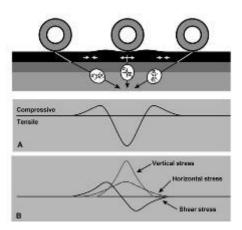
TRIZ Principle 15. Dynamicity

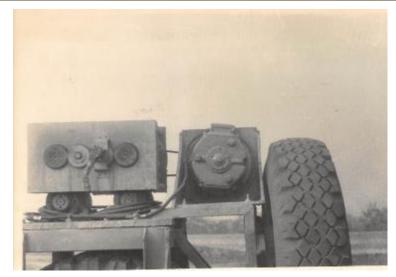
Farber B. et al., Multi axes Dynamic Vehicle Patent #1199885 Farber B. et al., Unbalanced

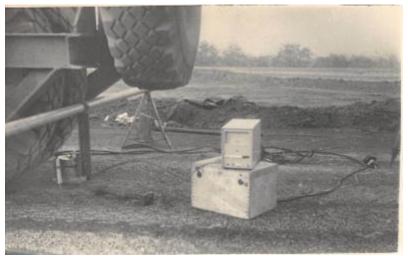
Vibrator Patent # 649478

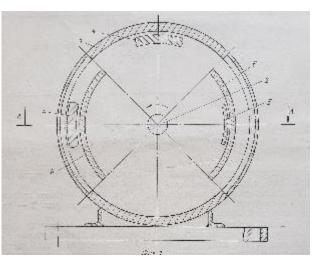






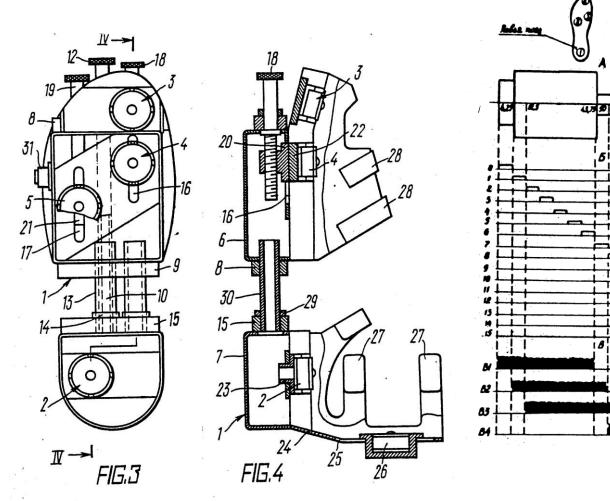






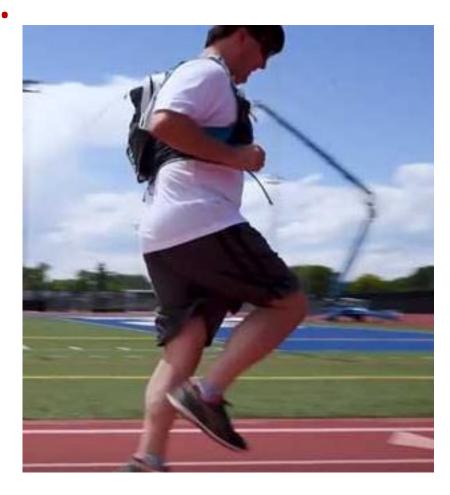
Противоречие: чтобы быстрее восстановиться, пациенту следует выполнять легкие упражнения, в том числе ходить, но он не может ходить. Как ходить, не ходя? «Умная обувь» -Вибросканер // Временная диаграмма управления вибратором в Вибросканере.



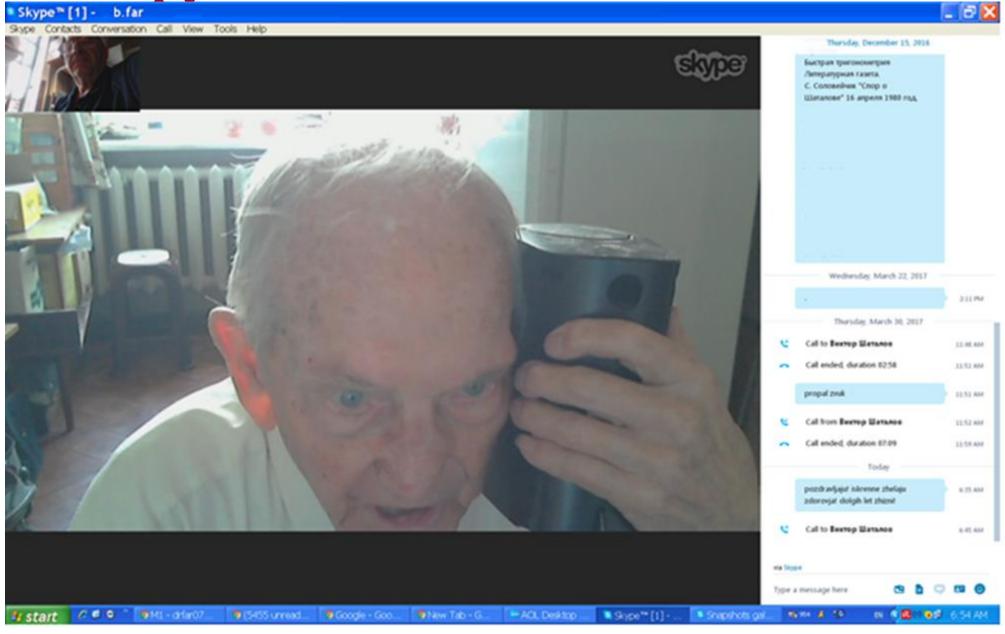


Принцип ТРИЗ №23. Обратная свьязь №18. Механическая вибрация Бегущие слепые-немыеглухие и наше «вибро» решение.





49 лет дружбы «С 90-летием»!

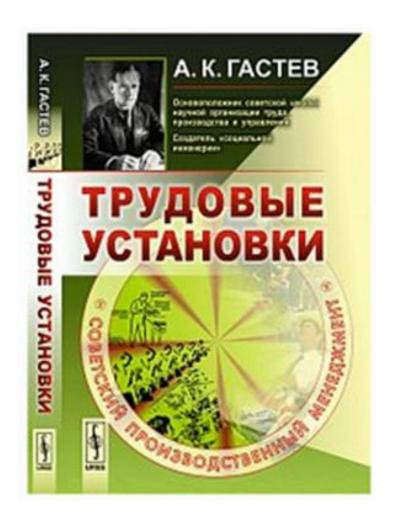


Наследие Алексея Гастева

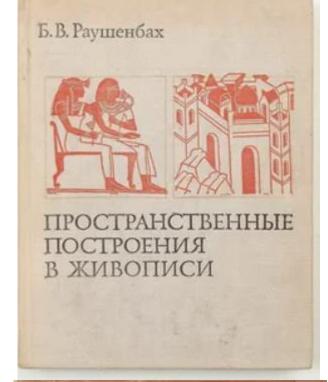


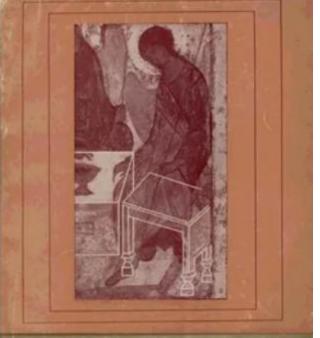


Алексей Гастев - один из пионеров научного менеджмента - теории менеджмента, которая анализирует и синтезирует рабочие процессы. Его основная цель - повышение производительности труда.





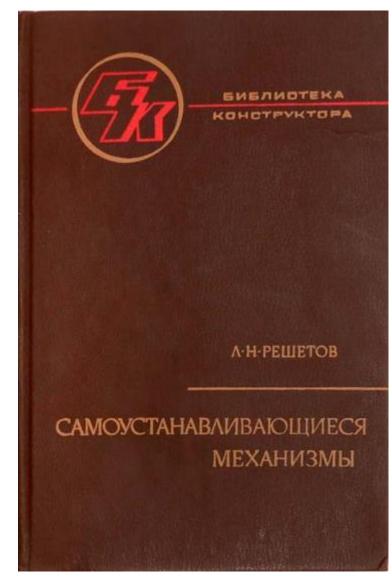


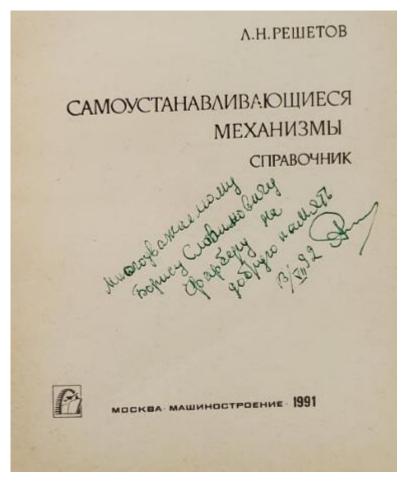


Д-р Леонид Решетов - самостоятельно настроенные системы

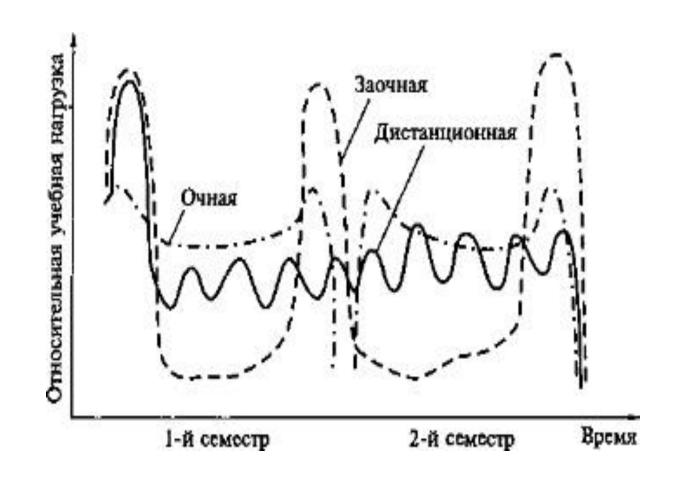


Л.Н. Решетов 1906-1998





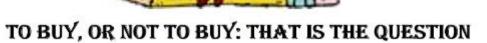
Изменение учебной нагрузки студента в течение года по различным формам. по критериям формирования знаний, умений и навыков, а также вследствие более равномерного распределения учебной нагрузки студента в течение года технология дистанционного обучения сопоставима с очной формой обучения и значительно превосходит по качественным параметрам заочную форму обучения.

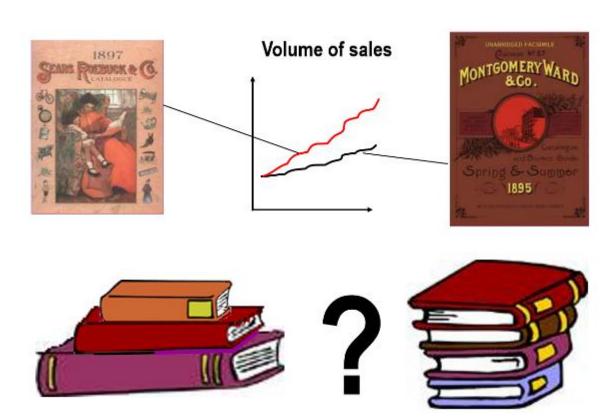




Geometry and Business (Morphometry)

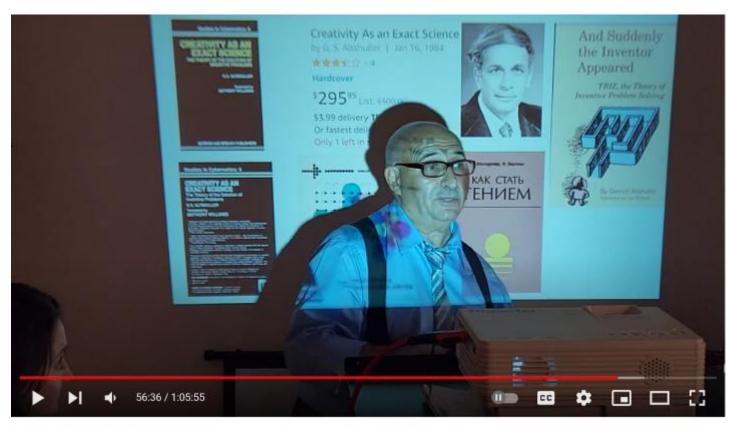






Internship Introductory Meeting (Part 1 & Part2) **Dr. Farber, Dr. Kleyn 07-17-2023**

https://www.voutube.com/watch?v=msWhJmG1nso&ab_channel=Farber%27sCenter





Internship Introductory Meeting (Part 2) Dr.Farber, Dr. Kleyn 07-17-2023



Internship Introductory Meeting (Part 1) Dr.Farber, Dr. Kleyn 07-17-2023

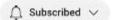
Internship at Dr. Farbers' corporations:

Farber's Center for Academic Success, Inc. Noigel, LLC, TRIZ Biopharma International, LLC



MASTERCLASS AND CAREER GUIDANCE WHEN CHOOSING A PROFESSION (INTERNSHIP AT THE FARBER'S CENTER

















Q&A SESSION





TRIZ SUMMIT 2023

THANK YOU!







