early (soon after puberty) and very late (approaching menopause) pregnancies are common in humans.
Ming-Hseng Wang, Frederick S. vom Saal
Division of Biological Sciences, University of
Missouri, Columbia, Missouri 65211, USA
e-mail: vomsaalf@missouri.edu

1. vom Saal, F. S. J. Anim. Sci. 67, 1824-1840 (1989).
2. Howdeshell, K. L., Hotchkiss, A. K., Thayer, K. A., Vandenbergh, J. G. \& vom Saal, F. S. Nature 401, 763-764 (1999)
3. Clutton-Brock, T. H. (ed.) Reproductive Success: Studies of Individual Variation in Contrasting Breeding Systems (Univ.

## Intelligence

## Maze-solving by an amoeboid organism

The plasmodium of the slime mould Physarum polycephalum is a large amoeba-like cell consisting of a dendritic network of tube-like structures (pseudopodia). It changes its shape as it crawls over a plain agar gel and, if food is placed at two different points, it will put out pseudopodia that connect the two food sources. Here we show that this simple organism has the ability to find the mini-

Chicago Press, Chicago, 1988).
4. vom Saal, F. S., Finch, C. E. \& Nelson, J. F. in Physiology of Reproduction Vol. 2 (eds Knobil, E., Neill, J. \& Pfaff, D.) 1213-1314 (Raven, New York, 1994).
5. Finch, C. E. \& Kirkwood, T. B. L. Chance, Development, and Aging (Oxford Univ. Press, 2000).
6. Matt, D. W., Lee, J., Sarver, P. L., Judd, H. L. \& Lu, J. K. H. Biol. Reprod. 34, 478-487 (1986).
7. Bernstein, L. et al. J. Natl Cancer Inst. 76, 1035-1039 (1986).
8. Bernstein, L. et al. Br. J. Cancer 58, 379-381 (1988).
9. Key, T. et al. Br. J. Cancer 73, 698-701 (1996).
10.Holliday, R. Science 238, 163-170 (1987).
11.vom Saal, F. S. et al. Proc. Natl Acad. Sci. USA 94, 2056-2061 (1997).
mum-length solution between two points in a labyrinth.

We took a growing tip of an appropriate size from a large plasmodium in a $25 \times 35$ cm culture trough and divided it into small pieces. We then positioned these in a maze created by cutting a plastic film and placing it on an agar surface. The plasmodial pieces spread and coalesced to form a single organism that filled the maze (Fig. 1a), avoiding the dry surface of the plastic film. At the start and end points of the maze, we placed $0.5 \times 1 \times 2 \mathrm{~cm}$ agar blocks containing nutrient $\left(0.1 \mathrm{mg} \mathrm{g}^{-1}\right.$ of ground oat flakes). There were four possible routes ( $\alpha 1$,

c

d

|  | None | $\beta 1$ | $\beta 2$ | $\beta 1$, <br> $\beta 2$ |
| :---: | :---: | :---: | :---: | :---: |
| None | 2 | 0 | 0 | 0 |
| $\alpha 1$ | 0 | 0 | 0 | 0 |
| $\alpha 2$ | 0 | 5 | 6 | 3 |
| $\alpha 1$, <br> $\alpha 2$ | 0 | 0 | 0 | 3 |

Figure 1 Maze-solving by Physarum polycephalum. a, Structure of the organism before finding the shortest path. Blue lines indicate the shortest paths between two agar blocks containing nutrients: $\alpha 1(41 \pm 1 \mathrm{~mm})$; $\alpha 2(33 \pm 1 \mathrm{~mm})$; $\beta 1(44 \pm 1 \mathrm{~mm})$; and $\beta 2(45 \pm 1 \mathrm{~mm})$. b, Four hours after the setting of the agar blocks (AG), the dead ends of the plasmodium shrink and the pseudopodia explore all possible connections. c, Four hours later, the shortest path has been selected. Plasmodium wet weight, $90 \pm 10 \mathrm{mg}$. Yellow, plasmodium; black, 'walls' of the maze; scale bar, 1 cm . d, Path selection. Numbers indicate the frequency with which each pathway was selected. 'None', no pseudopodia (tubes) were put out. See Supplementary Information for an animated version of a-c.
$\alpha 2, \beta 1, \beta 2)$ between the start and end points (Fig. 1a).

The plasmodium pseudopodia reaching dead ends in the labyrinth shrank (Fig. 1b), resulting in the formation of a single thick pseudopodium spanning the minimum length between the nutrient-containing agar blocks (Fig. 1c). The exact position and length of the pseudopodium was different in each experiment, but the path through $\alpha 2$ - which was about $22 \%$ shorter than that through $\alpha 1$ - was always selected (Fig. 1d). About the same number of tubes formed through $\beta 1$ and $\beta 2$ as the difference (about $2 \%$ ) in their path lengths is lost in the meandering of the tube trajectory and is within experimental error.

The addition of food leads to a local increase in the plasmodium's contraction frequency, initiating waves propagating towards regions of lower frequency ${ }^{1-5}$, in accordance with the theory of phase dynamics ${ }^{6}$. The plasmodial tube is reinforced or decays when it lies parallel or perpendicular, respectively, to the direction of local periodic contraction ${ }^{7}$; the final tube, following the wave propagation, will therefore link food sites by the shortest path.

To maximize its foraging efficiency, and therefore its chances of survival, the plasmodium changes its shape in the maze to form one thick tube covering the shortest distance between the food sources. This remarkable process of cellular computation implies that cellular materials can show a primitive intelligence ${ }^{8-10}$.
Toshiyuki Nakagaki ${ } \dagger$,
Hiroyasu Yamada* ${ }^{*} \ddagger$, Ágota Tóth§
*Bio-Mimetic Control Research Center, RIKEN, Shimoshidami, Moriyama, Nagoya 463-0003, Japan
e-mail: nakagaki@postman.riken.go.jp
$\dagger$ Local Spatio-Temporal Functions Laboratory, RIKEN, Wako 351-0198, Japan
$\ddagger$ Research Institute for Electronic Science,
Hokkaido University, Sapporo 060-0812, Japan \$Department of Physical Chemistry, University of Szeged, PO Box 105, Szeged H-6701, Hungary

1. Durham, A. C. \& Ridgeway, E. B. J. Cell Biol. 69, 218-223 (1976)
2. Matsumoto, K., Ueda, T. \& Kobatake, Y. J. Theor. Biol. 122, 339-345 (1986).
3. Miyake, Y., Tada, H., Yano, M. \& Shimizu, H. Cell Struct. Funct. 19, 363-370 (1994).
4. Nakagaki, T., Yamada, H. \& Ito, M. J. Theor. Biol. 197, 497-506 (1999).
5. Yamada, H., Nakagaki, T. \& Ito, M. Phys. Rev. E 59, 1009-1014 (1999).
6. Kuramoto, Y. in Chemical Oscillations, Waves and Turbulence (Springer, Berlin, 1984).
7. Nakagaki, T., Yamada, H. \& Ueda, T. Biophys. Chem. 84, 195-204 (2000).
8. Sepulchre, J. A., Babloyantz, A. \& Steels, L. in Proc. Int. Conf. on Artificial Neural Networks (eds Kohonen, T. et al.) 1265-1268 (Elsevier, Amsterdam, 1991)
9. Sepulchre, J. A. \& Babloyantz, A. Phys. Rev. E 48, 187-195 (1993)
10. Steinbock, O., Tóth, Á. \& Showalter, K. Science 267, 868-871 (1995).

Supplementary information is available on Nature's World-Wide Web site (http://www.nature.com).

