

Box 1 | The Firmicutes and the Bacteroidetes

The Firmicutes and the Bacteroidetes are divisions — or phyla — within the domain Bacteria. The microbiota of the human gut is dominated by their members, most of which are benign, although a few are pathogenic.

The Firmicutes is the largest bacterial phylum. It contains more than 250 genera, including *Lactobacillus*, *Mycoplasma*, *Bacillus* and *Clostridium*. There is considerable variety in the phylum. For example,

the *Clostridium* species are obligate anaerobes (that is, they absolutely require anoxic conditions), whereas members of *Bacillus* form spores and many of them are obligate aerobes.

Streptococcus pyogenes, the well-known cause of 'Strep. throat', is also a member of the Firmicutes.

The Bacteroidetes include about 20 genera. In the human gut, *Bacteroides* is probably the most abundant single genus, and *Bacteroides thetaiotaomicron*

is one of the most abundant organisms. Species of *Bacteroides* are obligate anaerobes that are benign inhabitants of the gut. However, they are opportunistic pathogens that can cause disease if they gain access to the peritoneal cavity, for example following surgery or a perforated ulcer. Members of the Bacteroidetes are found in the intestinal tracts of many warm-blooded animals, but are also abundant in soil and sea water.

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greater than that of mice receiving microbiota from lean mice. Taken together, these data suggest that differences in the efficiency of caloric extraction from food may be determined by the composition of the microbiota, which, in turn, may contribute to differential body weights.

This is a potentially revolutionary idea that could change our views of what causes obesity and how we depend on the bacteria that inhabit our gut. But a great deal remains poorly understood. Most notably, it is not clear whether such small changes in caloric extraction can actually contribute to meaningful differences in body weight. There are few data that substantiate the predicted increased caloric extraction in obese humans. Small but persistent increases in efficiency might potentially cause the accumulation of excess body weight over long periods, but these small differences are not the cause of obesity in leptin-deficient mice. These mice have a single gene mutation that prevents the production of biologically active leptin⁷. The resulting increased caloric intake and reduced caloric expenditure is many times larger than the small difference in extraction that could be produced by differences in the microbiota. In fact, the differences in body fat between mice given the 'obese microbiota' and those given the 'lean microbiota' are so small that they could be accounted for entirely by the tiny differences in food intake, rather than by differences in caloric extraction.

Another unknown is why and how the make-up of the microbiota is shifted by differences in body weight. Given that acquiring food from the environment can be both calorically expensive and potentially dangerous, it would seem to be most adaptive to extract as many calories from every bite of food as possible. Moreover, if caloric extraction does become more efficient, the regulatory system would dictate that the organism responds by reducing its caloric intake. If a host organism had the ability to change its microbiota so as to increase caloric extraction, it would seem most adaptive to do so when facing famine conditions and losing weight. However, the data

indicate just the opposite — the microbiota seems to be more efficient in obese humans who already have the most stored energy, and shifts to being less efficient as the subjects lose weight¹.

There is also the issue of how conditions in the host organism could change the make-up of the microbiota. Low levels of leptin are a signal of starvation that triggers several changes in the neuroendocrine system that work to conserve calories⁶. Consequently, it would make sense that low leptin might also impart a signal to the microbiota to become more efficient at extracting calories from food. This hypothesis would fit the microbiota of the obese, leptin-deficient mice. However, in humans where

obesity is associated with increased leptin, the same trends in microbiota composition are found, making it unlikely that the microbiota is responding to leptin directly. Moreover, it seems that when the bacteria are transferred to a lean mouse in which leptin is normal, the bacteria retain their 'obese' character over a two-week period. Thus, it is not clear how gut bacteria 'know' whether the host is obese or lean.

Gordon and colleagues' results^{1,2} tempt consideration of how we might manipulate the microbiotic environment to treat or prevent obesity. But questions about how and why the composition of gut microbiota is regulated will have to be answered first. As we have discussed, those questions are many and various. The two papers nonetheless open up an intriguing line of scientific enquiry that will ally microbiologists with nutritionists, physiologists and neuroscientists in the fight against obesity. ■

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- Ley, R. E., Turnbaugh, P. J., Klein, S. & Gordon, J. I. *Nature* **444**, 1022–1023 (2006).
- Turnbaugh, P. J. *et al. Nature* **444**, 1027–1031 (2006).
- Farooqi, I. S. & O'Rahilly, S. *Endocr. Rev.* doi:10.1210/er.2006-0040 (2006).
- Hill, J. O. *Endocr. Rev.* doi:10.1210/er.2006-0032 (2006).
- Ahima, R. S. & Flier, J. S. *Nature* **382**, 250–252 (1996).
- Woods, S. C., Benoit, S. C., Clegg, D. J. & Seeley, R. J. *Science* **280**, 1378–1383 (1998).
- Zhang, Y. *et al. Nature* **372**, 425–432 (1994).

ASTROPHYSICS

A burst of new ideas

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Gigantic cosmological γ -ray bursts have fallen into a dichotomy of long and short bursts, each with a very different origin. The discovery of an oddball burst calls for a rethink of that classification.

The events known as γ -ray bursts (GRBs) are the most violent and luminous explosions observed in the Universe. In the early 1990s, it became clear that they come in two distinct flavours: longer-duration bursts, typically longer than 2 seconds, with a spectrum of emitted radiation that peaks at lower ('softer') energy; and shorter-duration bursts with a more energetic, 'harder' spectrum¹. Observations of burst afterglows in the past decade — particularly in the past year^{2–4} — have seemed to show that this division is a clean one, and is firmly rooted in the progenitor of each type of burst. According to this picture, long bursts are associated with a young stellar population, marking the deaths of massive stars whose lifetime is short⁵. Short bursts, on the other hand, are associated with an old stellar population, and are

probably powered by mergers of compact objects such as neutron stars or black holes⁶.

In this issue, four papers^{7–10} blow a hole in this cosy paradigm. They contain observations of a bright γ -ray burst, GRB 060614, that triggered NASA's GRB sentinel, the Swift satellite, at 12:43:48 UT on 14 June 2006. The burst defies pigeonholing within the current scheme.

Gehrels *et al.* (page 1044)⁷ detail the circumstances of this peculiar burst's discovery. It is one of the brightest ever seen, and was soon located precisely not only by Swift's instrumentation, but also by other space- and ground-based telescopes. The burst is situated in the suburbs of a faint and relatively nearby dwarf galaxy⁸. Its duration, recorded by Swift as 102 seconds⁷, characterizes it unambiguously

as a long GRB. According to previous experience, evidence for the death of a star in a stellar explosion — a supernova — should have been spotted in the burst's neighbourhood before too long. But the many optical telescopes around the world trained on the target, waiting for yet another confirmation of the connection between GRBs and supernovae, saw nothing.

Three further papers^{8–10} provide independent reports of the stringent upper limits on the radiation flux from a possible supernova underlying GRB 060614. Gal-Yam and colleagues (page 1053)⁸ made a series of observations with the Hubble Space Telescope in the weeks after the burst trigger. These set an upper limit more than 100 times fainter than the faintest supernova previously associated with a GRB — and indeed considerably fainter than any supernova ever observed. Della Valle *et al.* (page 1050)⁹ report complementary observations from the European Southern Observatory's Very Large Telescope in the Atacama desert in northern Chile. This survey started 15 hours and ended 65 days after the burst, and provides an upper limit on the flux that is about three times higher than that of Gal-Yam and colleagues', but still well below the luminosity of any known supernova over an unprecedentedly long time span. Fynbo *et al.* (page 1047)¹⁰ use a range of telescopes to arrive at a similar result — and also discover a second long burst with no apparent supernova signature.

The absence of a supernova need not in itself be revolutionary. The production of a significant amount of nickel-56, which is a prerequisite for a supernova, is not guaranteed in a collapsing star^{7,8}, and the earliest model to connect GRBs with the massive-star collapses indeed characterized the bursts as 'failed supernovae'⁵. A supernova might also precede its associated GRB¹¹. Nevertheless, the weight of evidence from the past decade is consistent with there being no significant gap between a GRB and its supernova, as well as with the hypothesis that every long GRB has a supernova accompanying it¹².

What makes the story more intriguing is that every property of GRB 060614 places it in the short-burst category — except, that is, for its duration. Gehrels *et al.*⁷ show that the time-lag of softer radiation behind harder parts of the burst's spectrum is small, as it is in a short GRB. Gal-Yam *et al.*⁸ find that the afterglow of the burst has a large offset from the star-forming region of the host galaxy; again, a feature more characteristic of a short GRB. Similarly, the star-forming rate of the host galaxy is relatively small compared with those of normal long GRBs^{8–10}, consistent with an old stellar population more likely to host a short burst.

Even regarding its duration, the seemingly long GRB 060614 can be shortened. Its γ -ray light curve consists of a short, hard early episode lasting around 5 seconds, followed by a long, soft tail⁷. Recent observations also indicate that most 'short' GRBs are not necessarily

| | Type Ia | Type II (Type Ib/c) |
|--------------------|---|--|
| Stellar population | Old | Young |
| Host galaxy | All types of galaxy | Late-type galaxies |
| Progenitor | Binary systems (accretion-induced collapse of white dwarfs) | Single-star systems (core collapse of massive stars) |

Figure 1 | The classification of supernovae¹⁸.

| | Type I (short-hard) | Type II (long-soft) |
|-----------------------------|---|---|
| Duration | Usually short (may have a long tail?) | Usually long |
| Spectrum | Usually hard (tail is soft) | Usually soft |
| Spectral lag | Short | Long |
| Associated supernova | No | Yes |
| Stellar population | Old | Young |
| Host galaxy | All types of galaxies (predominantly in regions of low star formation rate) | Late-type galaxies (predominantly in irregular, dwarf galaxies) |
| Location in the host galaxy | Outskirts | Central |
| Progenitor | Mergers of compact objects in binary systems? | Single-star systems? (Core collapse of massive stars) |

Figure 2 | A classification scheme for γ -ray bursts. The red rows show the analogy with the supernova classification scheme. Blue cells show the properties of GRB 060614 (refs 7–10).

so short, and are usually followed by a softer emission tail lasting around 100 seconds^{3,4}. Using an empirical relation between the spectrum hardness and the total energy budget of GRBs, it is possible to show¹³ that GRB 060614 would be marginally classified as short if only it were around eight times less energetic.

So how can this burst and the existing classification scheme be squared? There are in principle three possibilities⁸. First, GRB 060614 is indeed a long GRB associated with a collapsing star. If so, its progenitor must be very different from those of most other long GRBs because of the anomalous properties detailed above. Second, the burst belongs to the merger-type short GRBs — in which case, these should not carry the name 'short' any more. Third, this is the prototype of a completely different, third category of burst.

Given that the long–short paradigm is no longer adequate to describe the entire GRB phenomenon, a new terminology can be invented. Dividing bursts into Type I and Type II bursts by analogy with the supernova classification scheme might seem to lack imagination, but a comparison shows that such a definition might not be a bad choice (Figs 1, 2)¹³. The progenitors of Type Ia supernovae, like those of the traditional short GRBs, belong to the old stellar population and live in binary systems. Similarly, the progenitors of type II supernovae, like those of traditional long GRBs, are collapsing massive stars that die at a young age.

Comparing the observational properties of GRB 060614 (blue cells)^{7–10} with the multiple criteria in Fig. 2, one can see that this burst merges the properties of both categories. If one prefers to fit this burst into the straitjacket of bimodal classification, as I do, it would seem safer to apportion it to Type I, the traditional

short category. There are great theoretical difficulties in producing extended radiation emission from a merger of compact stars, although various ideas to overcome these problems have been suggested^{14–17}.

To resolve definitively whether GRB 060614 is a peculiar example of a Type I burst, or whether it is a representative of a third class of object, more data are needed. In particular, given that our current information about the make-up of this GRB's host galaxy does not fully rule out a Type II origin, discovering whether or not GRBs with similar properties will be detectable in elliptical host galaxies^{8,13}, which consist entirely of older stellar populations, will hold the key to the final answer. ■

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- Kouveliotou, C. *et al.* *Astrophys. J.* **413**, L101–L104 (1993).
- Gehrels, N. *et al.* *Nature* **437**, 851–854 (2005).
- Fox, D. B. *et al.* *Nature* **437**, 845–851 (2005).
- Barthelmy, S. D. *et al.* *Nature* **438**, 994–996 (2005).
- Woosley, S. E. *Astrophys. J.* **405**, 273–277 (1993).
- Paczynski, B. *Astrophys. J.* **308**, L43–L46 (1986).
- Gehrels, N. *et al.* *Nature* **444**, 1044–1046 (2006).
- Gal-Yam, A. *et al.* *Nature* **444**, 1053–1055 (2006).
- Della Valle, M. *et al.* *Nature* **444**, 1050–1052 (2006).
- Fynbo, J. P. U. *et al.* *Nature* **444**, 1047–1049 (2006).
- Vietri, M. & Stella, L. *Astrophys. J.* **492**, L59–L62 (1998).
- Woosley, S. E. & Bloom, J. S. *Annu. Rev. Astron. Astrophys.* **44**, 507–556 (2006).
- Zhang, B. *et al.* *Astrophys. J. Lett.* (in the press); preprint available at www.arxiv.org/astro-ph/0612238
- Faber, J. A. *et al.* *Astrophys. J.* **641**, L93–L96 (2006).
- Dai, Z. G. *et al.* *Science* **311**, 1127–1129 (2006).
- Rosswog, S. preprint available at www.arxiv.org/astro-ph/0611440 (2006).
- King, A. *et al.* *Mon. Not. R. Astron. Soc.* (in the press); preprint available at www.arxiv.org/astro-ph/0610452
- Filippenko, A. V. *Annu. Rev. Astron. Astrophys.* **35**, 309–355 (1997).