A solitary figure walks through a somber urban landscape. The permeating tone of sadness in this painting gives an impression of life with major depressive disorder, whereas the light punctuating the scene hints at hope. Because depression has become a serious problem in societies around the world, it is crucial to investigate the basic mechanisms underlying this disease and to develop effective prevention and therapies. See the special section beginning on page 67. For the story behind the cover, go to http://scim.ag/cov6103.

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