

Science  
AAAS

2016 Science/ AAAS  
使用手冊



智泉智泉國際事業有限公司  
VI Services, Ltd.

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## AAAS 簡介

American Association for the Advancement of Science 美國科學促進聯會，簡稱 AAAS (<http://www.aaas.org/>)，於 1848 年創立，旨在成為科學所有學科的代表機構。AAAS 歷史悠久，而且堪稱是全球最大的科學學會，目前著力支持科學交流，也支持科學與社會議題的討論。

## Science 全球首頁簡介 ([www.sciencemag.org](http://www.sciencemag.org))

為使讀者不論是經由電腦或手機，都能更方便地閱讀與分享文章，自 2016 年 1 月 12 日起，AAAS 發布了新設計的網站。AAAS 全新的網站設計著重於更易取得文章內容，並提供一鍵式 (one-click) 的文章引用、共享工具，加強的資訊計量數據等等，意欲提供讀者最佳瀏覽及互動經驗。

The screenshot shows the Science magazine website homepage. At the top, the 'Science' logo is prominent, with 'AAAS' in smaller text below it. Navigation links include 'Home', 'News', 'Journals', 'Topics', and 'Careers'. A search bar is located on the right. Below the navigation is a red banner with 'Recent Videos', 'Latest Podcasts', 'Photo Galleries', and 'Data Stories Contest'. The main content area features a large image of a DNA double helix. The headline reads 'Viral 'fossils' in our DNA may help us fight infection' with a sub-headline 'Latest News | Mar. 3, 2016'. Below this are sections for 'Latest News', 'Career News', 'Journals', and an 'Advertisement' for a webinar.

Science 全球首頁除了提供可連結到 AAAS 所出版的期刊內容，還匯集許多豐富的輔助及多媒體資源，您可利用綠色方框內之不同標籤來選擇不同專區。下面將針對不同專區逐一說明。



## 首頁專區 (Home)

在 Science 全球首頁專區之下，有數個固定項目：

### Recent Videos

提供科學研究的有關影片資源。

### Latest Podcasts

每週 podcast 提供您了解科研領域一個導引工具，可直接透過電腦線上收聽、下載 MP3 檔案或是從 iTunes 中訂閱收聽。

### Photo Galleries

科研領域也有卓越的非文字形式之多媒體資源，如投影片檔，Science 也一次彙整相關資訊於此。

### 其他最新消息發布

在 Science 全球首頁上，出版社會因有最新消息發布，而不定期地可能有暫時性的網頁加入。

## 新聞專區 (News)

每天 Science 雜誌的新聞組都會為線上使用者提供幾篇有關科研成果或科學政策的每日最新消息；另外也會提供每週出版之 Science 雜誌收錄的新資訊。這些消息簡潔扼要，使讀者以些許的時間，就能即時瞭解世界各地各科研領域的最新進展。

### Latest News

每日均有數篇最新科學/科技相關新聞，由 Science 本身的特約作者或是外部特約作者寫作發表，讓讀者能了解最新趨勢發展。

### Scienceinsider

提供使用者世界各地有關科學政策的重大消息與分析。



## ScienceShots

Science 新聞組為線上使用者提供的重大每日新聞精選，並附有令人驚嘆的影像或影片。

## Sifter

編輯部從各網站中精選出科學相關的故事。

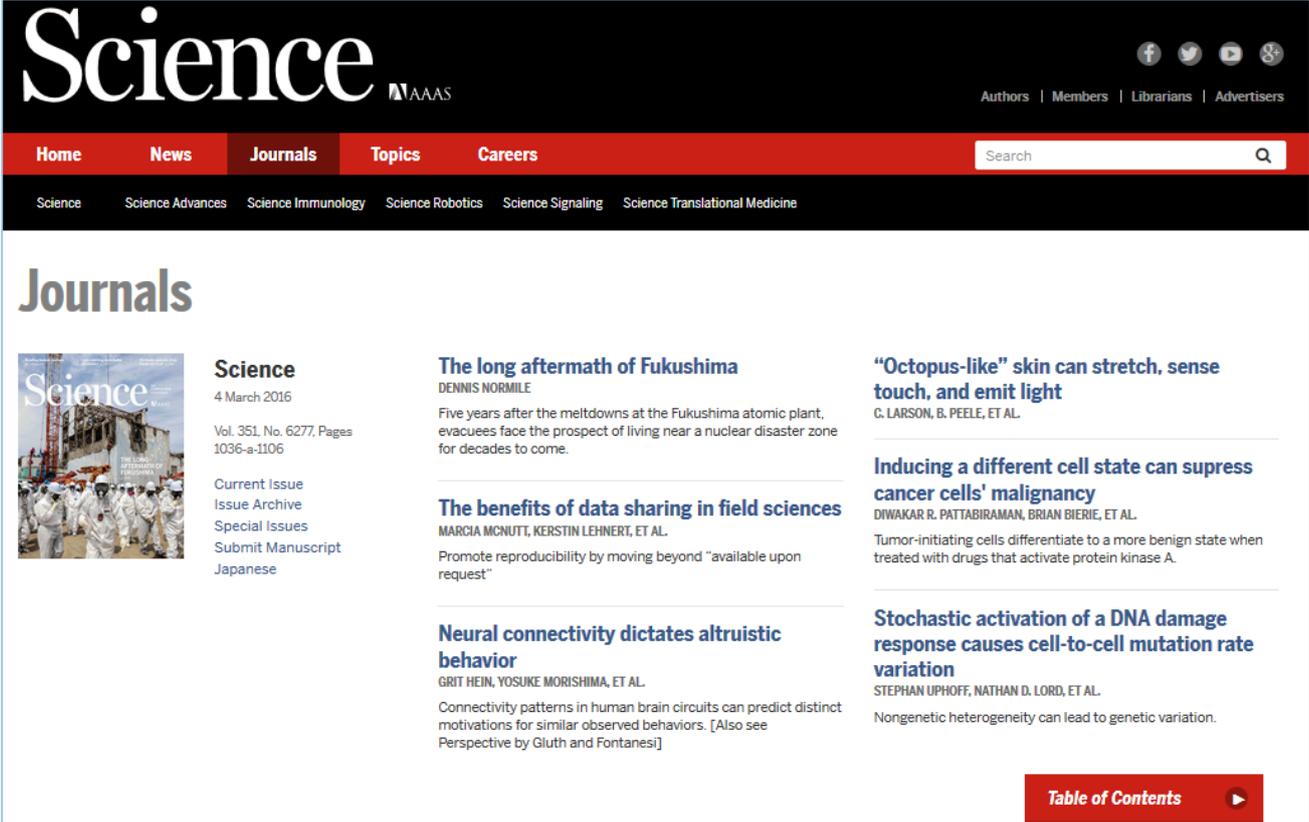
## Quizzes

此處提供 Science 自製的一些科學相關的問答題目。

## 期刊專區 (Journals)

目前 AAAS 除了出版頗富盛名的 *Science* 線上版之外，更提供其他的刊物，為某個專門領域的研究人員提供豐富、互聯且可檢索的資源，包括 *Science Signaling*、*Science Translational Medicine* (以上兩刊均須另外訂閱)，以及 *Science Advances* (Open Access 刊物)。

另外，AAAS 另有兩本新刊 —— *Science Immunology* 與 *Science Robotics* 正在籌備出版中。



**Science** AAAS

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Science Science Advances Science Immunology Science Robotics Science Signaling Science Translational Medicine

### Journals



**Science**  
4 March 2016  
Vol. 351, No. 6277, Pages 1036-a-1106  
Current Issue  
Issue Archive  
Special Issues  
Submit Manuscript  
Japanese

**The long aftermath of Fukushima**  
DENNIS NORMILE  
Five years after the meltdowns at the Fukushima atomic plant, evacuees face the prospect of living near a nuclear disaster zone for decades to come.

**The benefits of data sharing in field sciences**  
MARCIA MCNUTT, KERSTIN LEHNERT, ET AL.  
Promote reproducibility by moving beyond "available upon request"

**Neural connectivity dictates altruistic behavior**  
GRIT HEIN, YOSUKE MORISHIMA, ET AL.  
Connectivity patterns in human brain circuits can predict distinct motivations for similar observed behaviors. [Also see Perspective by Gluth and Fontanesi]

**"Octopus-like" skin can stretch, sense touch, and emit light**  
C. LARSON, B. PEELE, ET AL.

**Inducing a different cell state can suppress cancer cells' malignancy**  
DIWAKAR R. PATTABIRAMAN, BRIAN BIERIE, ET AL.  
Tumor-initiating cells differentiate to a more benign state when treated with drugs that activate protein kinase A.

**Stochastic activation of a DNA damage response causes cell-to-cell mutation rate variation**  
STEPHAN UPHOFF, NATHAN D. LORD, ET AL.  
Nongenetic heterogeneity can lead to genetic variation.

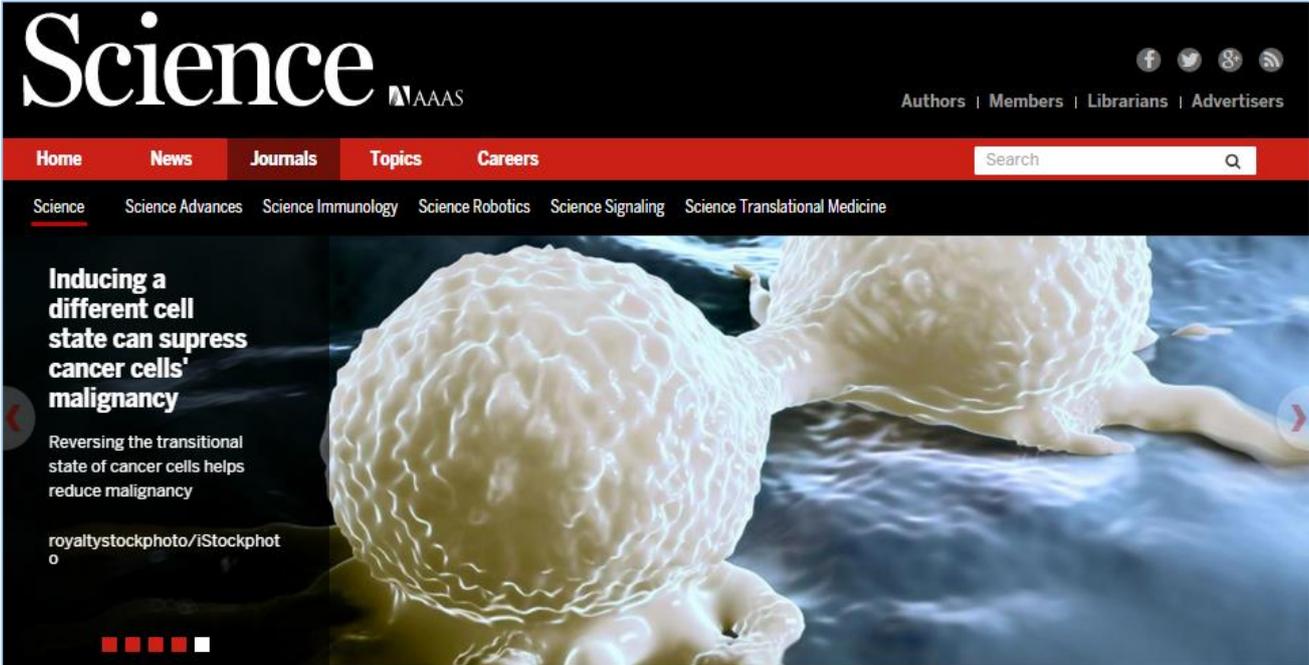
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## Science ([science.sciencemag.org](http://science.sciencemag.org))

*Science* 在美國發明家 Thomas Edison 的支援下於 1880 年創刊，如今在尖端研究、科學新聞及評論方面，已發展成為世界領先的刊物，其付費發行情量在同儕評鑑的一般科學期刊中居於首位。線上版 *Science* 雜誌可為您提供最新發佈的獲獎新聞內容，以及具高影響力的同儕評鑑研究資訊。*Science* 的內容依年代區分為下列三個產品(需個別訂閱才可使用)：

- ☑ **Science Online (1997 - 迄今)**： *Science* 的最新出版內容，每週五出刊。
- ☑ **Science Classic (1880 – 1996)**： *Science* 的過刊內容。
- ☑ **First Release (原為 Science Express)**：為 *Science* 的出版前服務。提供最新收錄的精選研究文章。通常這些文章會比紙本提早出版四至六週。待文章正式出版於 *Science* 之後，將由此處移除。



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**Inducing a different cell state can suppress cancer cells' malignancy**

Reversing the transitional state of cancer cells helps reduce malignancy

royalystockphoto/iStockphoto

**Contents** 04 MARCH 2016 VOL 351, ISSUE 6277

**THIS WEEK IN SCIENCE**

Research in *Science* journals.

SCIENCE | 04 MAR 2016 : 1038 | 🔒

Full Text

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- Collections
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  - Mission and Scope
  - Editors and Advisory Boards

## Science Advances ([advances.sciencemag.org](http://advances.sciencemag.org))

*Science Advance* 是新加入 *Science* 家族的同儕評論(peer-reviewed)期刊，於 2015 年 2 月份初次



出版，採投稿者付費而使用者免費的機制出版(gold open access)。 *Science Advances* 的主題範圍與 *Science* 大致相同，但收錄的文章涵蓋面更廣泛。對於那些因內容過長（受限於紙本空間大小）或者超出 *Science* 主題範圍，而無法收錄於 *Science* 的高品質論文，都會在 *Science Advances* 上出版。

**Science Advances** Authors | Members | Librarians | Advertisers

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**"MyShake": A smartphone app for earthquake warning**  
 Android app harnesses accelerometers in users' smartphones  
 Allanswart/istockphoto.com

**Latest Online** Archives

**A universal cooperative assembly-directed method for coating of mesoporous TiO<sub>2</sub> nanoshells with enhanced lithium storage properties**  
 BY BU YUAN GUAN, LE YU, JU LI, XIONG WEN (DAVID) LOU  
 SCIENCE ADVANCES | 04 MAR 2016 : E1501554 | CCBY-NC

A universal cooperative assembly method is developed for growing mesostructured TiO<sub>2</sub> shells on diverse functional particles.

Abstract Full Text PDF

**MORE FROM SCIENCE ADVANCES**

- Aims & scope
- Editorial board
- Author information
- Editorial policies
- Article licensing
- Article processing charges
- Frequently asked questions

## Science Signaling ([stke.sciencemag.org](http://stke.sciencemag.org))

*Science Signaling* 為研究員提供有關動態細胞訊號傳導領域中的突破性研究與論述的最新資源。從基礎科學到治療學的設計，以及從分子到網路與系統設計，*Science Signaling* 讓您的研究員、教師及學生保持領先地位。*Science Signaling* 能讓您立即獲取引領訊號傳導方面突破的理念和方法。



**Science Signaling**

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**Finding the active site**  
Crystal structure reveals a nonconsensus active site for an atypical phosphatase  
Irani *et al.*/Science Signaling

**Contents** 01 MARCH 2016  
VOL 9, ISSUE 417

**FOCUS**  
**Properly dividing with YAP**  
BY GERD P. PFEIFER  
SCI. SIGNAL. | 01 MAR 2016 : FS3 |

YAP helps to define where cells undergoing mitosis separate.  
Abstract | Full Text | PDF

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## Science Translational Medicine ([stm.sciencemag.org](http://stm.sciencemag.org))

*Science Translational Medicine* 為來自所有與成熟及新興學科相關的基礎、轉譯及臨床研究人員及見習生，提供交流和交換想法的討論區，將基礎科學家與臨床研究者聯繫起來，進而提高世界各地的病人護理水準。*Science Translational Medicine* 有助於加快基礎研究影響臨床研究，以及臨床研究影響基本研究的多向過程。如今，源自實驗室的研究，可在患者治療方面發揮有利作用。





**Science Translational Medicine** AAAS Authors | Members | Librarians | Advertisers

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**Gliomas' Achilles' heel**  
Loss of a chromatin remodeler enhances glioma sensitivity to DNA-damaging therapies  
Koschmann et al. / *Science Translational Medicine*

**Contents** 02 MARCH 2016  
VOL 8, ISSUE 328

**RESEARCH ARTICLES**  
**Dendritic cell vaccines based on immunogenic cell death elicit danger signals and T cell-driven rejection of high-grade glioma**  
BY ABHISHEK D. GARG, LIEN VANDENBERK, CAROLIEN KOKS, TINA VERSCHUERE, LOUIS BOON, STEFAAN W. VAN GOOL, PATRIZIA AGOSTINIS  
*SCIENCE TRANSLATIONAL MEDICINE* | 02 MAR 2016 : 328RA27 | PDF

Combining an immunogenic cell death inducer with dendritic cell immunotherapy treats high-grade glioma in a preclinical model.  
Editor's Summary | Abstract | Full Text | PDF

**MORE FROM SCIENCE TRANSLATIONAL MEDICINE**

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## Science Immunology

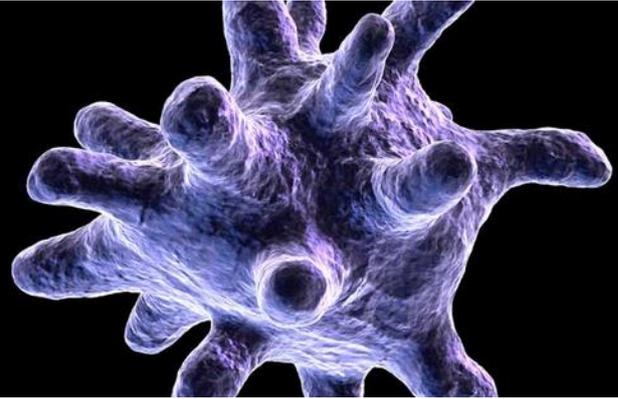
目前免疫學正處於一個前所未有的擴張與進展時期。免疫系統對健康與疾病在各個不同方面的廣泛影響力，正開始被醫界與學界理解與重視。*Science Immunology* 將在這個不斷增長的領域中，為最激動人心的發現提供一個交流平台。

*Science Immunology* 將出版具原創性、經同儕評審、以科學為基礎、包含重要新工具和技術，促進免疫學各個領域的關鍵進展的文章。由編輯群及科學家組成的國際諮詢小組，將保證 *Science Immunology* 所出版的文章，擁有與 *Science* 系列期刊同樣的高品質水準。(本刊 2016 年春季將開始接受投稿，投稿相關資訊請參考：

<http://www.sciencemag.org/journals/immunology/information-for-authors>)



Now accepting  
papers!



## Science Robotics

無論是完全獨立或與人類緊密合作，機器人逐漸變得無所不在。無論是在太空和深海勘探發掘、執行手術或駕駛汽車，機器人的影響力與日俱增。*Science Robotics* 為最新的技術進展，以及被機器人圍繞所產生的社會、倫理和政策關鍵問題提供了一個急需的論壇。

*Science Robotics* 同時迎合滿足研究人員和一般利益相關者，它的多學科內容，涵蓋了機器人的傳統學科，以及新出現的趨勢，例如先進材料和仿生設計；它也涵蓋了所有尺度，從非常大的系統到微/奈米機器人；它的適用範圍很廣，同時兼顧理論的進展與實際應用。

*Science Robotics* 將出版具原創性、經同儕評審、以科學或工程為基礎、包含機器人領域重要進展的文章。由學術編輯群組成的國際小組，將保證 *Science Robotics* 所出版的文章，擁有與 *Science* 系列期刊同樣的高品質水準。(本刊 2016 年春季將開始接受投稿，投稿相關資訊請參考：<http://www.sciencemag.org/journals/robotics/information-for-authors>)

Coming  
soon!



## 主題專區 (Topics)

### Hot Topics

提供收錄在 News 專區及 Science Online 中相關於該熱門主題的新聞報導與研究文章。

### Curated Topics

提供收錄在 News 專區及 Science Online 中相關於該專門主題的聞報導與研究文章。

### Special Issues

提供 Science 所出版的特刊連結，以及相關該特刊主題的新聞報導以及其他 Science Online 中的文章。

### Custom Publishing

提供委由 AAAS Custom Publishing Office 出版，由某些機構單位所贊助的客製化主題單冊，內容多半包含隸屬該贊助機構之作者的相關主題文章（不一定是曾出版在 Science 的文章）。

## 職涯專區 (Careers)

Science 的職涯專區致力於成為世界首要，不論是工業、學術或是政府機關中能夠給予科學家最佳的資訊資源環境。其宗旨在提供科學家們相關的需求資源，相對於也給予雇主一個最有效率的資源環境。

### Articles

提供相關職涯建立與抉擇的相關文章。

### Find Jobs ([jobs.sciencecareers.org](http://jobs.sciencecareers.org))

目前已有超過三千種工作需求，並且每日更新。

### Career Resources

提供如何準備與撰寫履歷表 (How-To 系列專輯)、最新職涯趨勢報導 (免費小冊供下載)、ScienceCareen 臉書連結……等等相關資訊。



## Forum

提供對於科學職涯疑問的 Q&A 討論論壇。

## For Employers

提供雇主關於在 Science 紙本期刊中刊登廣告，或是在 ScienceCareer 中刊登職務需求的相關資訊……等等。

## Graduate Programs

提供國際間大學或實驗機構所提供的碩/博士課程計畫的相關資訊。

## Careers Features

提供由 AAAS Custom Publishing Office 所撰寫的職涯特點文章。



## Science 期刊網站使用介紹

此處以 Science Online 網站之使用為例，當連結到期刊首頁之後，首頁上方會以跑馬燈形式展示當期 Science 內容精選五篇，點選圖片即可連結至該篇文章。

The screenshot shows the Science journal website homepage. The main header features the 'Science' logo and the AAAS logo. Below the logo is a navigation bar with links for Home, News, Journals, Topics, and Careers. A search bar is located on the right side of the navigation bar. The main content area features a large image of an octopus-like skin with the headline '“Octopus-like” skin can stretch, sense touch, and emit light'. Below the headline is a sub-headline: 'A newly developed artificial skin demonstrates multifunctionality'. A callout box points to a row of five small red squares, indicating '五篇精選的當期文章' (Five selected articles of the current issue). Another callout box points to the search bar, labeled '快速檢索框' (Quick search box). A third callout box points to the date and issue information: '04 MARCH 2016 VOL 351, ISSUE 6277', labeled '當期 Science 的出版日期與卷期編號' (Publication date and volume/issue number of the current Science). A fourth callout box points to the social media and RSS links in the top right corner, labeled '連結到 Science 的 facebook, Twitter, google+ 網頁，以及訂閱 Email 通知及 RSS feed。' (Link to Science's facebook, Twitter, google+ website, and subscribe to Email notifications and RSS feed). A fifth callout box points to the 'Contents' section, labeled '當期 Science 的出版日期與卷期編號' (Publication date and volume/issue number of the current Science).

利用網頁右側捲軸向下拉，即可瀏覽當期的每一篇文章的篇名。除摘要(Abstract) / 概要(Summary)之外，每篇文章視情況會提供 HTML 或 PDF 格式的全文 (或兩種格式皆提供)。在每篇文章篇名之下，有一個小鎖的符號，鎖頭若是呈現打開狀，則表示可下載閱讀 (機構已付費訂閱或該篇文章可能為 OA 文章)；若小鎖呈現封閉狀，則表示需訂閱才可下載閱讀。從文章的右側的連結區 (MORE FROM SCIENCE) 可連結至其餘相關的網頁。



**THIS WEEK IN SCIENCE**  
**Research in *Science* journals.**  
*SCIENCE* | 04 MAR 2016 : 1038 | 🔒  
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**EDITORIAL**  
**#IAmAResearchParasite**  
 BY MARCIA MCNUTT  
*SCIENCE* | 04 MAR 2016 : 1005 | 🔒  
 Summary | Full Text | PDF

**Data sharing**  
 BY MARCIA MCNUTT  
*SCIENCE* | 04 MAR 2016 : 1007 | 🔒  
 Summary | Full Text | PDF

**EDITORS' CHOICE**  
**This week in other journals.**  
*SCIENCE* | 04 MAR 2016 : 1039 | 🔒  
 Full Text

**PRODUCTS & MATERIALS**  
**New Products**  
*SCIENCE* | 04 MAR 2016 : 1101 | 🔒  
 A weekly roundup of information on newly offered instrumentation, apparatus, and laboratory materials of potential interest to researchers.  
 Summary | Full Text | PDF

**IN BRIEF**  
**News at a glance**  
*SCIENCE* | 04 MAR 2016 : 1009-1011 | 🔒

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**About The Cover**

## Current Table of Contents

瀏覽當期刊期內容

## First Release Science Papers

瀏覽最新收錄的精選研究文章(First Release 原為 Science Express)。First Release Science Papers 為 Science 的出版前服務，文章挑選標準包括研究的即時性及重要性。通常這些精選文章會比 Science 紙本提早出版四至六週。待文章正式出版於 Science 之後，將由此處移除。此處所提供的文章版本，可能因之後編輯群的編修，而會與正式出版於 Science 時有所不同。

## Archive

此處提供 Science 自 1880 年首次發行至迄今所有的刊期索引，使用者可以依照年份卷期瀏覽過期刊的目次，進一步取得摘要或全文。請留意 1880 年至 1996 年的刊期屬於 Science Classic 的內容，需另外訂閱才可取得全文。



# Archives

JULY 03, 1880 - MARCH 04, 2016

« 1990s 2000s **2010s** »

2010 2011 2012 2013 2014 2015 **2016** - - -

## 2016

 <b>Jan 01</b> Vol 351, Iss 6268	 <b>Jan 08</b> Vol 351, Iss 6269	 <b>Jan 15</b> Vol 351, Iss 6270	 <b>Jan 22</b> Vol 351, Iss 6271	 <b>Jan 29</b> Vol 351, Iss 6272	 <b>Feb 05</b> Vol 351, Iss 6273	 <b>Feb 12</b> Vol 351, Iss 6274	 <b>Feb 19</b> Vol 351, Iss 6275
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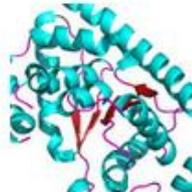
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Q

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Research journals
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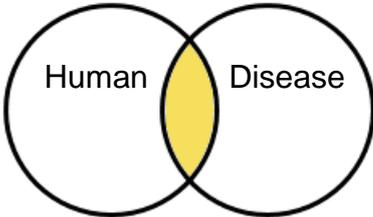
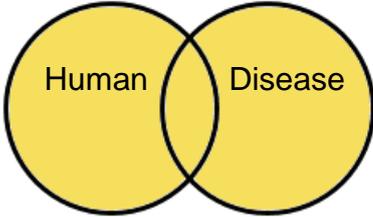
在 Search Term 欄位中採用的檢索定義的是字彙檢索(word)，因此如果檢索的是一個詞(phrase)或片語，就必須使用引號 ""。例如，輸入"Genome Sequence"，檢索結果只會查詢到 Genome Sequence；如果輸入的是 Genome Sequence，沒有引號，檢索結果則別包括 genome、sequence 和 genome sequence。

### 2. 切截符號

可使用 "\*" 作為切截符號，如"bio\*"則可檢索出 biology、biophysics、biological、bio-chemistry ……等等結果。

### 3. 布林邏輯

可使用布林邏輯 AND、OR、NOT，可用於同一欄位中。

布林邏輯	解釋與範例
<p>AND</p> 	<p>兩個或兩個以上概念的交集。例如：我想查詢 Human 和 Diseases 的資料，請輸入 "Human AND Diseases"</p>
<p>OR</p> 	<p>兩個或兩個以上概念的聯集。例如：我想查詢 Human 或 Diseases 的資料，請輸入 "Human OR Diseases"</p>





## 檢索結果說明 (此處以 Ebola virus 為例)

在摘要或作者部份出現的檢索關鍵字會以粗體字標示。

# Search Results

JOURNALS CAREERS DAILY NEWS

預設檢索結果為僅檢索 Science 系列期刊，若要檢索新聞的內容，則請再點選“DAILY NEWS”連結。

**Your search for "term "ebola virus"" returned 375 results.**

Items/Page

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**Isolation of potent neutralizing antibodies from a survivor of the 2014 Ebola virus outbreak**

BY ZACHARY A. BORNHOLDT, HANNAH L. TURNER, CHARLES D. MURIN, WEN LI, DEVIN SOK, COLBY A. SOUDERS, ASHLEY E. PIPER, ARTHUR GOFF, JOSHUA D. SHAMBLIN, SUZANNE E. WOLLEN, THOMAS R. SPRAGUE, MARNIE L. FUSCO, KATHLEEN B. J. POMMERT, LISA A. CAVACINI, HEIDI L. SMITH, MARK KLEMPNER, KEITH A. REIMANN, ERIC KRAULAND, TILLMAN U. GERNGROSS, KARL D. WITTRUP, ERICA OLLMANN SAPHIRE, DENNIS R. BURTON, PAMELA J. GLASS, ANDREW B. WARD, LAURA M. WALKER  
*SCIENCE* | 04 MAR 2016 : 1078-1083 |

...antibodies from a survivor of the 2014 **Ebola virus** outbreak 10.1126/science...USA.  
 Antibodies targeting the **Ebola virus** surface glycoprotein (EBOV G) in recent years, **Ebola virus** (EBOV) outbreaks have increased in

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- **Full Text:** HTML 格式的文章全文
- **PDF:** PDF 格式的文章全文
- **Editor's Summary:** 編輯所寫的文章概要

**Productive Replication of Ebola Virus Is Regulated by Tyrosine Kinase**

BY MAYRA GARCÍA, ARIK COOPER, WEI SHI, WILLIAM BORNHANN, RICARDO CARRION, DANIEL KALMAN, GARY J. NABEL  
*SCIENCE TRANSLATIONAL MEDICINE* | 29 FEB 2012 : 123RA24 |

...Articles Productive Replication of **Ebola Virus** Is Regulated by the c-Abl1 Tyrosine...University, Atlanta, GA 30322, USA. **Ebola virus** growth is regulated by a c-Abl...c-Abl antagonists. Disabling **Ebola virus Ebola virus** causes severe, often...

Abstract

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...Protective monotherapy against lethal **Ebola virus** infection by a potentially neutralizing...Diseases, Bethesda, MD 20892, USA. **Ebola virus** disease in humans is highly lethal...human survivor of the 1995 Kikwit **Ebola virus** disease outbreak is still circulating...

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**An Ebola whole-virus vaccine is protective in nonhuman primates**  
BY ANDREA MARZI, PETER HALFMANN, LINDSAY HILL-BATORSKI, FRIEDRIKE FELDMANN, W. LESLEY SHUPERT, GABRIELE NEUMANN, HEINZ FELDMANN, YOSHIHIRO KAWAOKA  
*SCIENCE* | 24 APR 2015 : 439-442 |

...Previously, we showed that a whole **Ebola virus** (EBOV) vaccine is protective in nonhuman primates...defective...protective immune responses. The **Ebola virus** vaccine is protective in nonhuman primates...Africa...One countermeasure to mitigate **Ebola virus** infections is vaccination. Several...

Abstract | Full Text | PDF | Editor's Summary

**Epitopes Involved in Antibody-Mediated Protection from Ebola Virus**  
BY JULIE A. WILSON, MICHAEL HEVEY, RUSSELL BAKKEN, SHAWN GUEST, MIKE BRAY, ALAN L. SCHMALJOHN, MARY KATE HART  
*SCIENCE* | 03 MAR 2000 : 1664-1666 |

...Antibody-Mediated Protection from **Ebola Virus** 10.1126/science.1011111  
provide protection from **Ebola viruses**, monoclonal antibodies (mAbs...epitopes is conserved among all **Ebola viruses** that are known to be pathogenic...

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- Science Translational Medicine (25)
- Science Signaling (8)
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Isolation of potent neutralizing antibodies from a survivor of the 2014 Ebola virus outbreak

Zachary A. VanDevanter<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Charles D. Murin<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Shua D. Shamblin<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Suzanne E. Wollen<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Thomas Pommert<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Lisa A. Cavacini<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Heidi L. Smith<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Michael A. Pauland<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Tillman U. Gerngross<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Karl D. Wittrup<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Erica Ollmann Saphire<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Dennis R. Burton<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Pamela J. Glass<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Andrew B. Ward<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>, Laura M. Walker<sup>1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29,30,31,32,33,34,35,36,37,38,39,40,41,42,43,44,45,46,47,48,49,50,51,52,53,54,55,56,57,58,59,60,61,62,63,64,65,66,67,68,69,70,71,72,73,74,75,76,77,78,79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94,95,96,97,98,99,100</sup>

+ Author Affiliations

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Science 04 Mar 2016:  
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DOI: 10.1126/science.1257888

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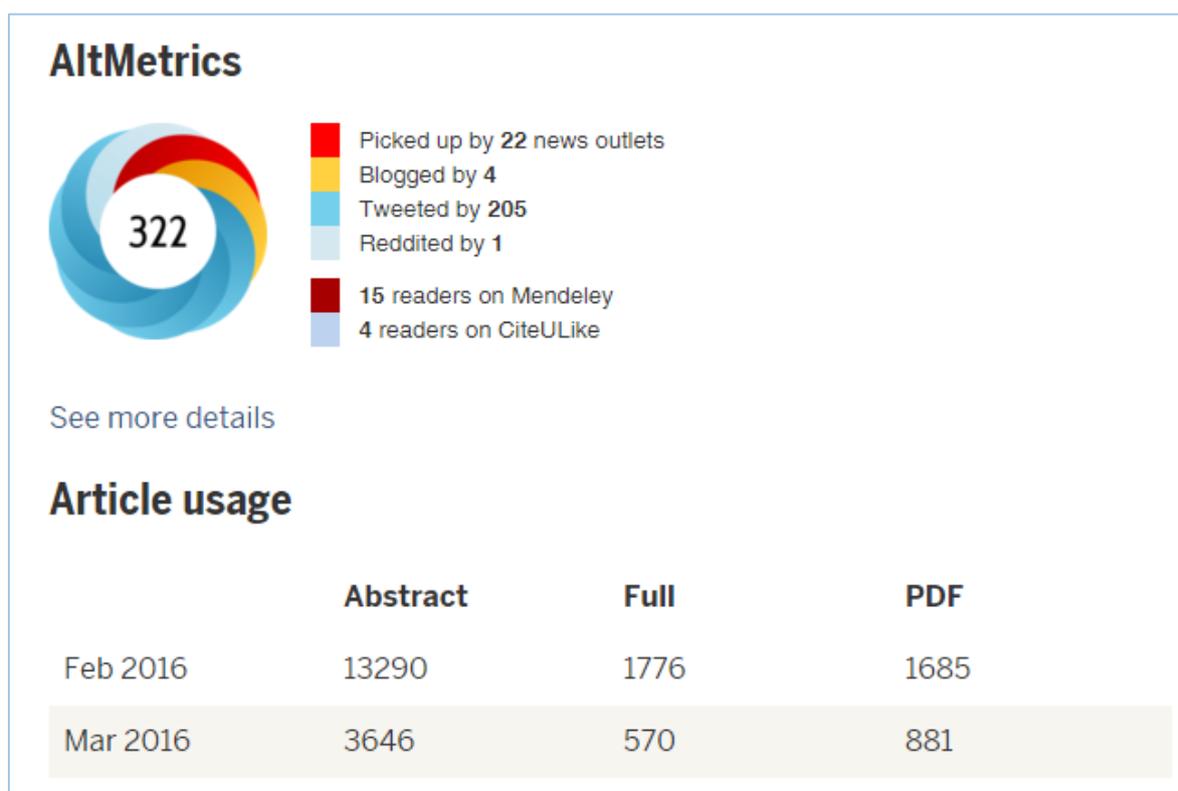
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mAbs exhibit unprecedented potency. Structures of selected mAbs in complex with GP reveal a site of vulnerability located in the GP stalk region proximal to the viral membrane. Neutralizing antibodies targeting this site show potent therapeutic efficacy against lethal EBOV challenge in mice. The results provide a framework for the design of new EBOV vaccine candidates and immunotherapies.

In recent years, Ebola virus (EBOV) has caused outbreaks in West Africa, with duration, and geographical spread, underscoring the need for pre- and post-exposure treatments (1). The membrane-anchored EBOV glycoprotein (GP) trimer is the sole known target for protective antibodies and is currently the primary target for antiviral vaccines and therapies (2, 3). A small number of protective monoclonal antibodies (mAbs) to GP have been isolated from immunized mice, and recent structures of these antibodies in complex with GP have illuminated key sites of vulnerability on the EBOV glycoprotein (3-7). However, only a small number of mAbs to GP have been isolated, and the characteristics of the human response to GP remains undefined.

In this study, we aimed to compare the protective efficacy of a panel of mAbs against EBOV GP by cloning an extensive library of mAbs from a convalescent donor (subject 1) and testing them in mice. Three months after primary infection, we isolated mAbs with high neutralizing activity, suggesting that this donor had a strong and durable neutralizing (Nab) response to GP by this time.

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lane 5). Notably, transfection of c-Ab11 siRNA decreased the quantity of VLPs by ~5-fold or by ~2.5-fold as measured by NP or VP40 protein levels, respectively, after immunoprecipitation with GP (Fig. 1A, lane 11). No effect was observed on intracellular protein levels of GP (Fig. 1A, lanes 7 to 9). The effect was specific; similar results were observed for a control siRNA (Fig. 1A, lanes 10 and 12), and c-Ab11 siRNA had no effect on the expression of the protein, eIF4E (eukaryotic initiation factor 4E) (Fig. 1B, compare lane 13 with lanes 14 to 16). Moreover,

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Fig. 2. (A) Percentages of selected K252 competitors that cross-reacted with GP core that can cap (GPCL). (B) Percentage of selected K252 competitors that cross-reacted with GP core that can cap (GPCL). (C) ELISA results showing the binding of selected mAbs to GP core that can cap (GPCL). (D) Summary of the antigenic sites targeted by the mAbs to GP. All data are representative of two or more independent experiments.

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