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#### Current issues in relapsing fever

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#### Abstract

Purpose of review: Relapsing fever has the highest incidence of any bacterial disease in Africa and a massive epidemic potential due to current political turmoil in the Horn of Africa. This review focuses on recent advances in diagnostics, molecular biology and host-pathogen interactions.

Recent findings: Complete relapsing fever genomes have recently been published, and the first site-specific genetic knockout complementation has been performed. Relapsing fever has gone from being a neglected disease to garnering interest in aspects such as tissue invasion, membrane biochemistry and complement evasion. Relapsing fever symptoms are variable, and the disease is commonly misdiagnosed as, for example, malaria. Although relapsing fever is considered a transient disease, it persists as a residual infection in the brain, which can be reactivated on immunosuppression. Therefore, single-dose antibiotic treatment should be avoided. Instead, treatment should cover a longer period, similar to the recommended regime for Lyme disease. Relapsing fever is a common cause of pregnancy complications such as intrauterine growth retardation and placental damage with spirochaetes crossing the maternal-foetal barrier, resulting in congenital infection.

Summary: Although relapsing fever remains a big problem, recently described host-pathogen interactions, diagnostics and molecular biology advances such as completed genome sequences and the dawn of genetic tools have brought relapsing fever research into the 21st century.

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#### Introduction

Relapsing fever is caused by spirochaetes belonging to the genus *Borrelia* and is transmitted by ticks or lice. The disease is initially blood borne and



#### Prevention

LBRF is best prevented by maintaining good hygiene, thereby avoiding louse infestation. During epidemics, isolation and delousing of relapsing fever patients and their personal belongings is important. Ornithodoros ticks are always close to their hosts, and tick-borne relapsing fever is best prevented by avoiding sleeping in close proximity to the vector's host animals. In the long run, decimation of the host population might be effective but in a shorter perspective could lead to ticks instead seeking humans for feeding. Tick bites can be avoided by using insect repellent containing *N*,*N*-diethyl-*m*-toluamide and permethrine-treated mosquito bed nets. A bed elevated above the ground is preferable to sleeping on the floor to restrict contact with questing ticks. Human settlements may be sprayed with insecticides, for example, permethrine, to eradicate tick infestation, but ticks may survive or repopulate the hut from external sources. Therefore, treatment has to be repeated, for example, once a year. The best solution, not yet affordable for everyone, is housing made of concrete instead of clay or straw, providing an uninhabitable environment for ticks (Fig. 4).



Figure 4

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#### Treatment

Relapsing fever Borrelia are so sensitive to antibiotics that a single dose is sometimes recommended [42,43], especially for treatment of LBRF. This is often sufficient and may be justified during epidemics when drug availability is low. However, these recommendations were written when relapsing fever was considered a disease solely of the blood. Today, more and more evidence points to silent and residual infections in sites difficult to reach by some antibiotics such as the brain [4,19]. A short antibiotic treatment may eliminate circulating Borrelia, but the brain and perhaps other sites might function as reservoirs, whereby bacteria can re-enter the blood after insufficient treatment. Several observations show that relapsing fever might relapse after 1 or 2 days' treatment. This persistence has been reported to be more common in tick-borne relapsing fever than LBRF [19]. As relapsing fever shows close resemblance to Lyme disease in its ability to cause persistent infection in sites, which might be hard to reach by antibiotics, we suggest therapy similar to that for Lyme disease patients, for example, oral amoxicillin or doxycycline. In more severe cases, intravenous injections with ceftriaxone or cefotaxime are recommended, or as a third alternative, penicillin G (Table 1) [44]. Animal experiments have also shown that ceftriaxone clears a residual B. duttonii infection in the brain [27]. LBRF epidemics often occur as mixed infections with epidemic typhus (Rickettsia prowazekii) [43]. Doxycycline is a suitable antibiotic when there is a risk of misdiagnosis or coinfection, as it is



49 Grosskinsky S, Schott M, Brenner C, et al. Borrelia recurrentis employs a novel multifunctional surface protein with anticomplement, antiopsonic and invasive potential to escape innate immunity. PLoS ONE 2009; 4:e4858. [Context Link]

Keywords: Borrelia; coinfection; diagnostics; relapsing fever; treatment; zoonotic reservoirs



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### Figure 1

Estimated distribution of relapsing fever Borrelia



The inset depicts the geographic distribution of malaria. Note, the almost complete geographic overlap of the two diseases.



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### Figure 3

Erythrocyte rosetting



Some relapsing fever species, for example, *B. duttonii*, *B. crocidurae* and *B. hispanica*, aggregate erythrocytes into so-called rosettes, which obstruct circulation in small blood vessels, causing microthrombosis, microhaemorrhage and hypoxia. The adhesion is weak and spirochaetes can easily swim back and forth within the aggregate. The biochemical nature of this interaction is yet not known.

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### Figure 4

Endemic relapsing fever habitat in a Mouba village in the Savanes region of Northern Togo



The big pots containing grain and corn attract rodents, which together with dogs, chickens and other domestic animals constitute hosts for relapsing fever transmitting ticks. Domestic and wild animals are in close contact here. The open houses made of mud and straw provide habitat for rodents and ticks transmitting disease. Control of reservoir animals and vectors, but most important, improved economical and housing conditions are needed to eradicate relapsing fever.

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#### Current issues in relapsing fever Christer Larsson, Marie Andersson and Sven Bergström

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Current Opinion in Infectious Diseases 2009, 22:443-449

#### Purpose of review

Relapsing fever has the highest incidence of any bacterial disease in Africa and a massive epidemic potential due to current political turmoil in the Horn of Africa. This review focuses on recent advances in diagnostics, molecular biology and host – pathogen interactions.

#### Recent findings

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Your search Search terms used: aspirin acetylsalicylic acid asa aspergum treat treat treatment therapy remedy heart disease heart diseases disorder of heart morbus cordis	<ul> <li>Aspirin-resistant platelet aggregation in a cohort of patients with coronary heart disease.</li> <li>Pamukcu, Burak; Oflaz, Huseyin; Onur, Imran; Oncul, Aytac; Umman, Berrin; Koylan, Nevres; Bugra, Zehra; Meric, Mehmet; Nisanci, Yilmaz</li> <li>Blood Coagulation &amp; Fibrinolysis. 18(5):461-465, July 2007. [ORIGINAL ARTICLES]</li> <li>AN: 00001721-200707000-00011.</li> <li>View Abstract</li> <li>AB Aspirin resistance could be defined as thrombotic and embolic cardiovascular events despite regular aspirin therapy. The study aimed to determine the profile and</li> </ul>	<ul> <li>PDF Full Text</li> <li>Abstract Reference</li> <li>Complete Reference</li> <li>Table of Contents</li> <li>Ovid Full Text</li> <li>Library Holdings</li> <li>Request Permissions</li> <li>Internet Resources</li> </ul>
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Full Text & CAB Abstracts Fulltext					Latest Update		









#### Take a look at the new OvidSP

Try it now!



#### **Resource Selection - OvidSP**

 Opening Ovid or Selecting Ovid Resources link will prompt the user with a popup to select their OvidSP resources

	Search Journals Books My W	orkspace External Links
	Basic Search   Find Citation   Search Fields	Advanced Ovid Search   Multi-Field Search
▼ Selected Resources	View Selected Resources     Select Reso     Enter topic or question     Limits (Click to exp:     All Resources	urce(s) to search:
Universal Search: ScienceDirect, Google, C	<ul> <li>▶ Search History (4 searches) (C</li> <li>□ All</li> <li>□ Add</li> <li>□ Tra</li> <li>□ E8</li> <li>□ E</li></ul>	Armals@Ovid ditional Content nsplant Library April 28, 2009 M Reviews - ACP Journal Club 1991 to March 2009 M Reviews - Cochrane Central Register of Controlled Trials M Reviews - Cochrane Database of Systematic Reviews 1 M Reviews - Cochrane Methodology Register 2nd Quarter M Reviews - Database of Abstracts of Reviews of Effects 2 M Reviews - Health Technology Assessment 2nd Quarter id MEDLINE(R) In-Process & Other Non-Indexed Citations Select Resource(s)



### Main Navigation Bar

- Main Navigation Bar replaces browsing links
- Selected Resources databases, journals, books
- Language, Support and Training links moved
- Personal Account My Account

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OvidSP

• My Workplace - Saved Searches, eTOCs & My Projects

Wolters Kluwer Ovid	My Account   My PayPerView   Ask A Librarian   Help   Logoff
Search Journals Books My Workspace External Links	
Basic Search   Find Citation   Search Fields   Advanced Ovid Search   Multi-Fi • Selected Resources • Ovid Resources: Soil Science - 1973 to April 2009, Plant Genetics and Breeding • Crop Science - 1973 to June 2009	eld Search - 1973 to April 2009,
Universal Search: ScienceDirect, Google, CINAHL, Web of Science, Google Scholar,  The search of the	Amazon.com, Inc, Library of Congress
Limits (Click to expand)     Include Related Terms Universal Se	arch
<ul> <li>Search History (4 searches) (Click to expand)</li> </ul>	View Saved

OvidSP - 多种语言界面						
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▶ 检索历史 (0项检索) ( <i>点嵌展开</i> ) 查看已保存的检索历史						
基本检索   引文检索   字段检索   <b>高级检索</b>   多个字段检索 ▼ 已选资源 Ovid 资源:						
<ul> <li>● 关键词</li> <li>○ 作者</li> <li>○ 期刊</li> <li>○ 检索</li> </ul>						
▶ <b>常用限制</b> ( <i>点击展开</i> )						
English   Français   Deutsch   繁體中交   Españo   简体中文 版帜所有 (c) 2000-2010 Ovid Technologies, Inc. 使用条款   支持和培训   关于我们   联系我们 版本: OvidSP_UI03.00.00_H37.101, SourceID 50829						
Wolters Kluwer OvidSP Search Without Boundaries						

Health

## **Results Management**



### **Results Tools - Options**

- Search Aids are now called Results Tools
  - users may add or remove items from the search tools using the Options menu

otions Select All
View Results Tools:
Select research tools to be displayed on left side of page
Search Information
Filter By
My Projects



## **Results Tools - Search Information**

- Search Information
  - Lists the actual search executed
  - Lists search terms used including related terms
  - Lists possible misspelling
  - Search results numbers returned
  - Sort by, and customize display options





# Results Tools - Filter By

- Filter By
  - Filters are like dynamic limits:
    - Selected Only displays selected results (via checkboxes)
    - Star ranking filter 4 or more, etc.
    - Year filter
    - Subject
    - Author
    - Journal
    - Resource
    - Publication Type
       Filters depend on database(s) selected
  - Add to Search History option adds filters as a search statement



### Output Icons & View Menu

- Results manager has gone now Print, Email, Export, My Projects
  - Select All all results

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- View: Title/Citation/Abstract three quick view options for displaying results
  - Title is title only
  - Citation view is standard Ovid Display
  - Abstract with all abstracts expanded
- Hyperlinked Article Title (to HTML, PDF or Complete Reference)



# My Projects 我的课题



### Access to My Projects

- Logged in to My Account, note Annotations icon
- Existing projects appear in Results Tools/My Projects

	Select All	🎒 Order 🛛 🚔 Print 🔤 Email 🕫	Export 🛛 🔂 Add to My Projects
My Projects	View: Title Cit	ation Abstract 10 Per Page 🗸 Go to #: 1 Go >>	Next Result
Allergic Asthma Project	1 🔲		Ovid Full Text     Complete Reference
• Stroke Rehabilitation	Unique Identifier	19969573	
	Authors	Smith R.	• Find Similar
	<b>Authors Full</b>		• Find Citing Articles
	Name	Smith, Robert.	
	Title	Faux review. A report on the zombie outbreak of 2009: how mathematics can save us (no, really).	• Full Text
	Source	CMAJ Canadian Medical Association Journal. 181(12):E297-300, 2009 Dec 8.	
	Publication Type	Journal Article.	

### My Workspace, My Projects

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 My Workspace stores all the users' research – Projects, Saved Searches/Alerts, eTOCs

. Wolters	Kluwer	Ovid	SP		My Account	My PayPerView	õ <sup>†</sup> Ask a Librar	ian   Help   Logoff
Search Jou	imals	Books M	y Workspace	External Lin	n <b>ks</b> My eTocs   Instal	l Toolbar		
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## Adding to My Projects

- Items that can be added to a project or folder:
  - Uploaded files (上传文件)
  - Ovid Results (Ovid检索結果)
  - Saved Searches (储存的检索历史)
  - Journal Articles PDF or HTML (期刊文章全文 PDF or HTML)
  - Images (图像)
  - Text Snippets (文本剪贴)
  - User-created Citations (客戶自己生成的题录)
  - Book Chapter (电子书章节)
  - Journal Issue List result (期刊目录結果)
  - Auto-Alert results (储存定题通告)

### Add to My Projects

- Add single items using item actions




# Add to My Projects

 Select multiple items and click "Add to projects" button from the list action



## Add to My Projects - Drag and Drop

- Items can be added to a project from the results display in the following ways:
  - Drag and drop results into My Projects

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### Journals@Ovid HTML Full Text view

Neuromodulation         Issue: Volume 9(1), January 2006, p 28-33         Copyright: Copyright (C) 2006 Blackwell Publishing Ltd.         Publication Type: [CASE REPORT]         DDI: 10.1111/j.1525-1403.2006.00039.x         ISSN: 1094-7159         Accession: 2006145036         Keywords: adult, aphasia, arm, article, Barthel index, brain cortex, brain depth stimulation, case report, functional magnetic         ICASE REPORT]         Effects of Bilateral Extradural Cortical Stimulation for Plegic Stroke Rehabilitation         Canavero, Sergio MD (US FMGEMS); Bonicalzi, Vincenzo MD; Intonti, Stefania Rehab. Spec; Crasto, Stefania MD; Castellano, Giancarlo MD            • Author Information         Turin Advanced Neuromodulation Group, Turin, Italy	Neuromodulation Issue: Volume 9(1), January 2006, p 28-33 Copyright: Copyright (C) 2006 Blackwell Publishing Ltd. Publication Type: [CASE REPORT] DOI: 10.1111/j.1525-1403.2006.00039.x ISSN: 1094-7159 Accession: 2006145036 Keywords: adult, aphasia, arm, article, Barthel index, brain cortex, brain depth stimulation, case report, functional magnetic resonance imaging, human, leg, male, medical assessment, single photon emission computer tomography, stroke [CASE REPORT] Effects of Bilateral Extradural Cortical Stimulation for Plegic Stroke Rehabilitation	Article Tools         Article as PDF (151KB)         Complete Reference         Abstract Reference         Print Preview         Email lumestart
Issue: Volume 9(1), January 2006, p 28-33         Copyright: Copyright (C) 2006 Blackwell Publishing Ltd.         Publication Type: [CASE REPORT]         Dob: 10.1111/j.1525-1403.2006.00039.x         ISSN: 1094-7159         Accession: 2006145036         Keywords: adult, aphasia, arm, article, Barthel index, brain cortex, brain depth stimulation, case report, functional magnetic resonance imaging, human, leg, male, medical assessment, single photon emission computer tomography, stroke         [CASE REPORT]         Effects of Bilateral Extradural Cortical Stimulation for Plegic Stroke Rehabilitation         Canavero, Sergio MD (US FMGEMS); Bonicalzi, Vincenzo MD; Intonti, Stefania Rehab. Spec; Crasto, Stefania MD; Castellano, Giancarło MD         * Author Information Turin Advanced Neuromodulation Group, Turin, Italy	Issue: Volume 9(1), January 2006, p 28-33 Copyright: Copyright (C) 2006 Blackwell Publishing Ltd. Publication Type: [CASE REPORT] DOI: 10.1111/j.1525-1403.2006.00039.x ISSN: 1094-7159 Accession: 2006145036 Keywords: adult, aphasia, arm, article, Barthel index, brain cortex, brain depth stimulation, case report, functional magnetic resonance imaging, human, leg, male, medical assessment, single photon emission computer tomography, stroke [CASE REPORT] Effects of Bilateral Extradural Cortical Stimulation for Plegic Stroke Rehabilitation	Article as PDF (151KB)  Complete Reference Abstract Reference Print Preview Fmail lumpstart
[CASE REPORT]         Effects of Bilateral Extradural Cortical Stimulation for Plegic Stroke Rehabilitation         Canavero, Sergio MD (US FMGEMS); Bonicalzi, Vincenzo MD; Intonti, Stefania Rehab. Spec; Crasto, Stefania MD; Castellano,         Giancarlo MD            • Author Information         Turin Advanced Neuromodulation Group, Turin, Italy	[CASE REPORT] Effects of Bilateral Extradural Cortical Stimulation for Plegic Stroke Rehabilitation	Email lumostart
Effects of Bilateral Extradural Cortical Stimulation for Plegic Stroke Rehabilitation Canavero, Sergio MD (US FMGEMS); Bonicalzi, Vincenzo MD; Intonti, Stefania Rehab. Spec; Crasto, Stefania MD; Castellano, Giancarlo MD  Author Information Turin Advanced Neuromodulation Group, Turin, Italy  Email Article Text Add to My Projects Export All Images to PowerPoint	Effects of Bilateral Extradural Cortical Stimulation for Plegic Stroke Rehabilitation	
Canavero, Sergio MD (US FMGEMS); Bonicalzi, Vincenzo MD; Intonti, Stefania Rehab. Spec; Crasto, Stefania MD; Castellano, Giancarlo MD  • Author Information Turin Advanced Neuromodulation Group, Turin, Italy  Export All Images to PowerPoint		Email Article Text
Turin Advanced Neuromodulation Group, Turin, Italy	Canavero, Sergio MD (US FMGEMS); Bonicalzi, Vincenzo MD; Intonti, Stefania Rehab. Spec; Crasto, Stefania MD; Castellano, Giancarlo MD	Add to My Projects
	Turin Advanced Neuromodulation Group, Turin, Italy	Export All Images to PowerPoin



# Adding graphics from Journals@Ovid

Adding graphics from HTML Full Text in Journals@Ovid ullet





close or Esc Key

\*

# Adding text quotes using Snippet Tool

### Adding text using the Snippet Tool

### A report on the zombie outbreak of 2009: how mathematics can save us (no, really)

Smith, Robert MSc PhD?

#### Author Information

From the Department of Mathematics and the Faculty of Medicine, University of Ottawa, Ottawa, Ont.

**Correspondence to:** Dr. Robert Smith?, Department of Mathematics and Faculty of Medicine, University of Ottawa, Ottawa ON K1N 6N5; rsmith43@uottawa.ca

Acknowledgements: The author is supported by an NSERC Discovery Grant, an Early Researcher Award and funding from MITACS. So huge truckloads of taxpayer money were spent on research that involved capturing zombie specimens and travelling to exotic locations to observe zombies in the wild. Numerous lavish lunches were also consumed along the way, but none involved the brains of the living. Honest.

#### Back to Top

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#### Abstract

An outbreak of zombification wreaked havoc recently in Canada and the rest of the world. Mathematical models were created to establish the speed of zombie infection and evaluate potential scenarios for intervention, mainly because mathematicians don't have anything better to do with their time. We review the development of these models and their effect on the undead.

In August 2009, a new disease emerged in North America and quickly made its way around the world. 1 Media reports suggest the outbreak began in Ottawa 2 but rapidly spread across Canada 3,4 and was transported thereafter to the United States 5 and the United Kingdom. 6,7

OvidSP



### **AutoAlerts**

- Adding Items from AutoAlert Results
- allows new results to automatically go into a project

elivery Options
Email 🔽 RSS 🔽 My Projects
Email RSS My Projects
Add Search to:
C An Existing Project
My Cancer Research
C. A. Now Project
O A New Project
Project Name*
Description



# Managing Projects

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- Projects may be Active or Archived or in the Trash
- Projects can be dragged between the archived and active window
- Folders can be dragged and dropped between projects
- Items remain in trash for 30 days before being permanently deleted
- Projects expire after 1 year, but can be renewed by the user

### Actions - Creating New Project/Folder/Citation

Manage Projects			Actions	
Name	Items	Мо	dified 🔻	
🗟 My Cancer Research	3	200	9-may-06	
🔚 Lymphoma	45	200	9-may-06	
🔚 Pancreatic	73	2009-may-02		
🚞 Brain Tumor	147	2009-may-02		
🛅 Oropharyngeal	7	200	9-may-01	
🕥 Managing Burns	346	200	09-apr-23	
Sew Procedures	146	2009-apr-23		
Obstetrics & Gyneco	7	2009-feb-02		
🚞 Other	346	200	09-jan-23	

Archived Projects				
Name	Items	Modified <b>V</b>		
🔤 My Neruology	0	2009-may-06		
📧 Dr. Smith's Paper	346	2009-apr-23		
Nursing advancment	2009-mar-23			
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📴 Pancreatic	73	200	Upload	File	act All
🔚 Brain Tumor	147	200			acc run
Oropharynoeal	7	200	9-may-01		Sort by:

- Actions menu allows you to create projects or folders, to add references manually or upload files
- Projects may be Active, Archived or Deleted



# Uploading a File



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# My Projects Storage

- Note; reference, full text, images, etc. are NOT stored in My Projects when added from OvidSP; the citation is referenced
  - (a link points back to the original reference).

This minimizes storage overhead

and Assures access is based on a current subscription



## OvidSP 3.0 Toolbar

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- The Toolbar is designed to allow the user to collect external resources and save them to My Projects
  - OvidSP will check for the existence of the toolbar, and if it is not installed, we will offer a small icon/link allowing the user to install it.



## OvidSP 3.0 Toolbar

- Clicking "x" will remove the toolbar
- OvidSP will check version, and alert the user when new versions are available
- Toolbar will be visible on all sites, but grey (disabled) on OvidSP

ScienceDirect - NeuroImage : Juicy fruit and creepy cr	awlies: An electrophysiological study of - Windows Internet Explorer	
COO . Ktp://www.sciencedirect.com/science?_ob=4	ArticleURL&_ud=86WNP-43MKMV3-1&_user=10&_rdoc=1& 💌 🕒 😽 🗙 ಶ Uve Search	2
File Edit View Favorites Tools Help	🛪 🚭 Convert 👻 🔂 Select	
🚔 Favorites 🛛 🙀 🏀 Suggested Sites 🔹 🔊 Get More Adr	d-ons -	
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Health	😒 ScienceDirect - Ne 🗙 🖉 OSP2 < Main < Ovid5 🔰 🏠 ・ 🖾 · 🗠 👼	• Page • Safety • Tools • 📦 •
ScienceDirect	You have Guest access to ScienceDirect Find out more	Login: III Register
Home Browse My Settings Alerts Help		
Quick Search All fields	Author	
@ search tips Journal/book title	Volume Issue Page Clear 🐼 Go 🕣	

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# Thank You !



