## PROTEGEN

Hilda Delgadillo and Mitchell Petredis Frank R. Seaver College of Science and Engineering Loyola Marymount University October 1, 2013

## OUTLINE

- Purpose
- What is Protegen?
- Biological Information
- Species in the Database
- Biological Questions that can be Answered
- Type of Database

- Maintenance and Updates
- Funding
- Accessibility
- External Links
- Downloads
- "User Friendliness"

## PURPOSE

• Protegen contains curated information about protective antigens

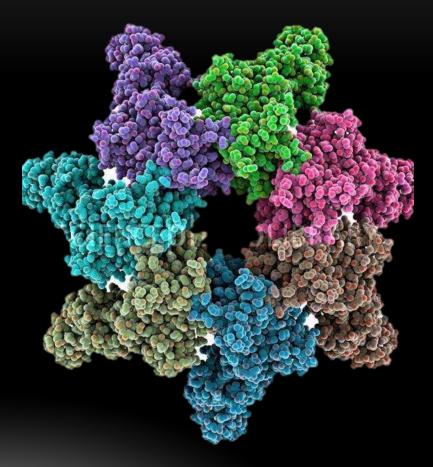
## PROTEGEN

- One of the only databases to curate and analyze protective antigens that can be used for vaccine development
- This database has allowed for the "storage, annotation, comparison, and analysis of 854 protective antigens for 200 pathogens, including more than 50 protective antigens for cancer and allergy" (Yang, Sayers, Xiang, He, 2010).



## **BIOLOGICAL INFORMATION**

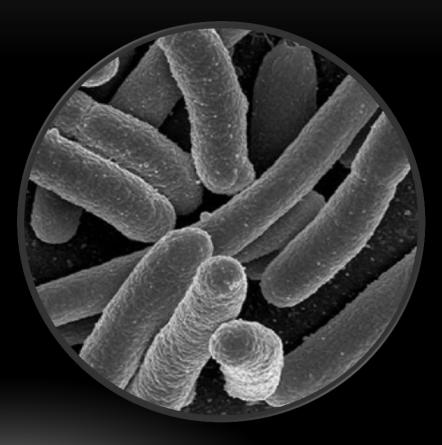
- 854 Protective Antigens for 200 pathogens
  - Storage
  - Annotation
  - Comparison
  - Analysis
    - 50+ Protective Antigens are used for cancer and allergy



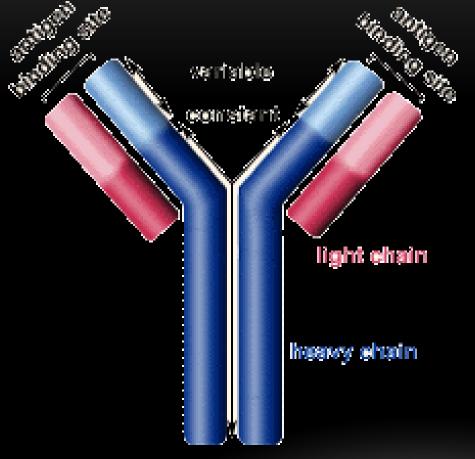
Anthrax Protective Antigen Molecule Credit: Laguna Design/Science Photo Library

## SPECIES

- Protective antigens
- Pathogens
  - Bacteria
  - Viruses
  - Parasites
  - Fungus



## BIOLOGICAL QUESTIONS THAT CAN BE ANSWERED



- What are protective antigens ?
- Importance of identification of protective antigens
- Protective Antigens as biological markers

## TYPE OF DATABASE

- "Protegen stores manually curated protective antigens and associated information."
  - Manual curation includes peer-reviewed publications from PubMed.
- Protegen is an integrated program of the VIOLIN
  - VIOLIN: vaccine investigation and online information network. vaccine database and analysis

## MAINTENANCE

- University of Michigan Medical School, Ann, Arbor, MI.
  - Webmaster: Zuoshuang Xiang
- Database is SUPPOSED to update every quarter
  - Most recent update was on November 11, 2011



## FUNDING

- National Institutes of Health-National Institute of Allergy and Infectious Diseases grant
  - (R01Al81062)



## ACCESSIBILITY

# 

## EXTERNAL LINKS

- PubMed
  - Provides extensive information about a protegen
- Protein Data Bank
  - 3D Models can be found here



## DOWNLOADS

- OWL Format: "Obfuscated Weird Language"
- FASTA: Common for gene sequences
- EXCEL
- HTML: easily accessible by any computer.

FASTA format of the Protegen sequences can be downloaded below.

- protegen-all-2.0-2011-11-17.fna (fasta) (html)
- protegen-all-2.0-2011-11-17.faa (fasta) (html)
- protegen-bacterium-2.0-2011-11-17.fna (fasta) (html)
- protegen-bacterium-2.0-2011-11-17.faa (fasta) (html)
- protegen-virus-2.0-2011-11-17.fna (fasta) (html)
- protegen-virus-2.0-2011-11-17.faa (fasta) (html)
- protegen-parasite-2.0-2011-11-17.fna (fasta) (html)
- protegen-parasite-2.0-2011-11-17.faa (fasta) (html)
- protegen-fungus-2.0-2011-11-17.fna (fasta) (html)
- protegen-fungus-2.0-2011-11-17.faa (fasta) (html)
- protegen-allergy\_cancer-2.0-2011-11-17.fna (fasta) (html)
- protegen-allergy\_cancer-2.0-2011-11-17.faa (fasta) (html)

#### Archieves:

#### Version 1.0.

For other methods of data exchange, please visit data exchange.

## **"USER-FRIENDLINESS"**

¢ for

- Very well organized and visually appealing
- Tutorial available right next to search bar
- Yields categorized searches



GO Clear Help

#### Protegen: Protective Antigens

Protective antigens are specifically targeted by the acquired immune response of the host and are able to induce protection in the host against infectious and non-infectious diseases. Protective antigens play important roles in development of vaccines and biological markers for disease diagnosis and analysis of fundamental host immunity against diseases. Protegen is a web-based central database and analysis system that curates, stores, and analyzes protective antigens.

Search Field	Search Parameter				
Pathogen or Disease	Any pathogen  \$				
Gene Name	(e.g., sodC)				
Locus Tag	(e.g., BAB2_0535)				
Database ID	NCBI Gene GI ‡				
COG category	Any Amino acid transport and metabolism Carbohydrate transport and metabolism Cell cycle control, cell division, chromosome partitioning				
Subcellular Localization	Any Localization Cellwall Cytoplasmic Cytoplasmic Membrane				
Maximum Number of Transmembrane Helices	1 (Note: check box to include)				
Minimum Adhesin Probability (0-1.0)	0.51				
No Similarity to Human Proteins					
No Similarity to Mouse Proteins					
No Similarity to Mouse Proteins					

Search: Vaxquery

#### Statistics: Protegen includes 854 protective antigens for now.

Provenance: The data in Progeten originates from our literature curation and boinformatics analyses



#### Protegen Home

Intr	od	ucti	on

```
Statistics
```

News and Updates

Protegen Query

Protegen BLAST

Selected Bacteria Brucella spp. (25) B. anthracis (13) E. coli (17) M. tuberculosis (26) N. meningitidis (21)

Selected Viruses Ebola virus (15) HIV (41) Influenza virus (49)

Selected Parasites Plasmodium (33)

Data Submission

Data Exchange Data Download

Documentation

FAQs Disclaimer

Contact Us

## EXAMPLE SEARCH

- **RB51** 
  - 4 Categories  $\bigcirc$ 
    - Vaccine
    - Pathogen
    - Vaccine-Related
    - Vaccine-Related Literature
  - Offers links to PubMed or  $\bigcirc$ Protein Data Bank when available.

A abortizs DNA vaccine encoding RpiL and Omp16     A abortizs train 19     A abortizs train 18     A abortizs train 18B51     Licensed (Calle) Brucelit     A molifensis bp20 and TF Nasal Vaccine     A molifensis strain VTR51     Brucelit     Bruce	his searc		arch	curated vaccine d	ata in VIOLIN.				iced Se h Help
A aboritus DNA vaccine encoding RpiL and Omp16     A aboritus DNA vaccine encoding RpiL and Omp16     A aboritus strain 15     A aboritus strain 17851     A aboritus strain VTR51     Brucelit     Brucelit aboritus exsA mutant vaccine     Brucelit     Compare 2. Pathogen Search:	ound 14 v ame for d	accine(s etail. To	s) whose de compare v			es and		mpare	<b>-</b> .
B. abortus strain 19     B. abortus strain 19     B. abortus strain 19     B. abortus sceine strain R851     Licensed (Catlle) drucelli,     B. mellitensis train TRM1     B. mellitensis train VTRM1     B. mellitensis     B. mellitensis train V	Selection	Vaccine	Name			Status		Target	Pathog
B. abortis vaccine strain R851     B. meitleonis bp26 and Tr Hasai Vaccine     B. meitleonis strain VTRM1     B. meitleonis strain VTRM1     B. meitleonis strain VTRM1     Brucelis abortis strain VTRM1     Brucelis abortis strain VTRM1     Divalent DNA 8: abortis ptDNA3:1-17L12-Omp16     Divalent DNA 8: abortis R851SOD     Recombinant 8: abortis RB51SOD     Recombinant 8: abortis RB51SOD     Recombinant 8: abortis RB51SOD     Recombinant 8: abortis RB51SOD     Compare 2. Pathogen Search:		B. abort	B. abortus DNA vaccine encoding RpIL and Omp16 Research						a spp.
B. meillensis bp26 and TF Nessi Vaccine     B. meillensis bp26 and TF Nessi Vaccine     B. meillensis bp26 and TF Nessi Vaccine     B. meillensis stain VTRM1     B. suis strain VTRM1     B. s		B. abortus strain 19 Licensed (Cattle)						Brucella spp.	
B. mellitensis strain VTEM1     B. mellitensis strain VTEM1     B. sub strain VTEM1     Brucelita abortus spk mutant vaccine     Brucelita abortus spk mutant vaccine     Brucelita abortus spk mutant vaccine     Divalent DNA B. abortus vaccine pcDNA3.1-L7L12-Omp16     Live attenuated B. abortus with deletion of znuA     Recombinant B. abortus RB51V0boA     Recombinant B. abortus RB51V0bA     Recombinatt B. abor		B. abortus vaccine strain RB51 Licensed (Cattle)							a spp.
B. suis strain YTR51     Brucella abortus exsA mutant vaccine     Brucella abortus exsA mutant vaccine     Brucella abortus pix mutant vaccine     Divalent DNA B. abortus Yaccine pcDNA3.1-L7/L12-Om16     Live attenuate B. abortus with deletion of 2nuA     Raboral     Recombinant B. abortus RB51SOD     Recombinant B. abortus RB51WboA     Recombinat		B. melitensis bp26 and TF Nasal Vaccine							a spp.
Brucelika abortus exsA mutant vaccine     Brucelika abortus pots Mutant vaccine     Brucelika abortus pots Mutant vaccine     Brucelika abortus pots Mutant vaccine     Divalent DNA 8: abortus potSNA3.1-17/L12-Omp16     Licensed 08: abortus vilh deletion of znuA     Recombinant 8: abortus RE51SOD     Recombinant 8: abortus RE51SVDbA     Recombinant 0: anthropi 49237SOD     Compare 2. Pathogen Search:		B. melite	B. melitensis strain VTRM1						
Brueelka abortus pgk mutant vaccine     Brueelka abortus yaccine pcDNA3.1L7/L12-0m16     Divalent DNA & abortus Vaccine pcDNA3.1L7/L12-0m16     Live attenuated B. abortus with deletion of znuA     Recombinant B. abortus RB51SOD     Recombinant B. abortus RB51WboA     Recombinant B. abortus RB51WboA     Recombinant D. anthropi 49237SOD     Compare  2. Pathogen Search:		B. suis s	B. suis strain VTRS1						a spp.
Divalent DNA 6. abortus Vaccine pcDNA3.1-L7/L12-Omp16     Divalent DNA 6. abortus Valocine pcDNA3.1-L7/L12-Omp16     Recombinant 6. abortus vitil deletion of znuA     Recombinant 6. abortus RES1SOD     Recombinant 6. abortus RES1SOD     Recombinant 6. abortus RES1V90A     Recombinant 0. anthropi 49237SOD     Compare 2. Pathogen Search:		Brucella	Brucella abortus exsA mutant vaccine Research					Brucel	la spp.
Live attenuated <i>B. abortus</i> with deletion of znuA     Raboral     Raboral     Recombinant <i>B. abortus</i> RB51SOD     Recombinant <i>B. abortus</i> RB51Wb6A     Recombinant <i>B. abortus</i> RB51Wb6A     Recombinant 0. anthropi 49237SOD     Compare  2. Pathogen Search:		Brucella abortus pgk mutant vaccine				Resear	ch	Brucel	a spp.
Raboral     Recombinant <i>B. abortus</i> RB51SOD     Recombinant <i>B. abortus</i> RB51WbbA     Recombinant <i>B. abortus</i> RB51WbbA     Recombinant O. anthropi 4923YSOD     Brucell     Compare  2. Pathogen Search:		Divalent	t DNA B. abo	rtus Vaccine pcDNA3	.1-L7/L12-Omp16			Brucel	a spp.
Recombinant B. abortus RB51SOD     Brucelli     Recombinant B. abortus RB51WboA     Brucelli     Recombinant O. anthropi 49237SOD     Brucelli     Compare 2. Pathogen Search:		Live atte	anuated B. at	ortus with deletion of	znuA			Brucel	a spp.
Recombinant <i>B. abortus</i> RB51WboA Brucell     Recombinant O. anthropi 49237SOD Brucell     Compare 2. Pathogen Search:		Raboral	1			License	ed (Dog)	Rabies	virus
Recombinant O. anthropi 49237SOD     Brucelle     Compare 2. Pathogen Search:		Recomt	ainant B. abo	tus RB51SOD				Brucel	a spp.
Compare 2. Pathogen Search:		Recombinant B. abortus RB51WboA					Brucel	a spp.	
2. Pathogen Search:		Recomb	pinant O. anth	ropi 49237SOD				Brucel	a spp.
				Con	npare				
Found 2 pathogen(s) related to the query keywords:		an Sear	ch:						
	Pathog				ds:				
# Name Name Vacalass related related	-	ithogen(	s) related to	the query keyword					
	Pathe Nam	ogen e	Disease Name	# of Associated Vaccines	# of vaccine- related pathogen ge	rel	lated st genes		
Brucella spp. Brucellosis 60 34 6	Path Nam Rabi	ogen e es virus	Disease Name Rabies	# of Associated Vaccines 27	# of vaccine- related pathogen ge	rel	lated st genes 0		eferend 16

#### 3. Vaccine-related Gene Search

#### Found 3 vaccine-related gene(s)

About Introduction Statistics VIOLIN News Your VIOLIN Register or Login Submission Tutorial

accine & Component Vaxquery Vaxgen VBLAST Protegen VirmugenDB DNAVaxDB Vaxjo Vaxvec Vevax Huvax Vaccine Mechanisms Vaxis Vaccine Literature VO-SciMine Litesearch Vaxmesh Vaxlert Vaccine Design Vaxio

Community Efforts Vaccine Ontology

ICoVax 2012 Advisory Com vaccine Society Vaxperts VaxPub VaxCom VaxLaw VaxMedia VaxMeet VaxFund VaxCareet

Data Exchange VIOLINM Help & Documents Publications Documents FAQs Links Acknowledg

Disclaimer Contact Us

Š́∕-

Gene ID	Gene Name	Sequence Strain (Species/Organism)	VO ID	NCBI Gene ID	NCBI Nucleotide Gl	3D structure: PDB ID	Related Vaccines(s)	Details
934	exsA	Brucella melitensis biovar Abortus 2308		3787220			<ul> <li>Brucella abortus exsA mutant vaccine</li> </ul>	More
1677	L7/L12	<i>Brucella abortus</i> RB51-AHVLA					<ul> <li>Divalent DNA B. abortus Vaccine pcDNA3.1-L7/L12- Omp16</li> </ul>	More
1678	omp16	<i>Brucella abortus</i> RB51-AHVLA					<ul> <li>Divalent DNA B. abortus Vaccine pcDNA3.1-L7/L12- Omp16</li> </ul>	More

NOTE: Go to Vaxgen for advanced vaccine gene search

#### 4. Vaccine-related Literature:

#### Found 22 vaccine-related article(s)

- 1. Trant CG, Lacerda TL, Carvalho NB, Azevedo V, Rosinha GM, Salcedo SP, Gorvel JP, Oliveira SC induces protection etent mice.. Infection or to that of vaccine strain 19 in immunocompromised a munity. 2010; 78(5); 2283-2291. [PubMed: 20194591]. and im
- 2. Martins H, Garin-Bastuji B, Lima F, Flor L, Pina Fonseca A, Boinas F Eradication of bovine bro rtugal-Outcome of a 5-year programme (2002-2007) based on test-and-slaughter ation.. Preventive veterinary medicine. 2009; 90(1-2); 80-89. [PubMed: 19439382].
- Nol P, Olsen SC, Rhyan JC Experimental inferictardsonii) with attenuated and virulent stra 2009; 45(1): 189-195. [PubMed: 19204348]. ental infection of Richardson's ground squirrels (Spermophil ilent strains of Brucella abortus., Journal of wildlife diseases.
- Yang X, Walters N, Robison A, Trunkle T, Pascual DW Nasal immunization with recombinant *Bru* melitensis bp26 and trigger factor with cholera toxin reduces *B. melitensis* colonization... *Vaccine*. melitensis bp26 and trigger factor with cholera to 2007; 25(12); 2261-2268. [PubMed: 17239499].
- Olsen SC, Waters WR, Stoffregen WS An aerosolized Brucella spp. challenge mod animals.. Zoonoses and public health. 2007; 54(8); 281-285. [PubMed: 17894637]. lel for laboratory
- Yang X, Becker T, Walters N, Pascual DW Deletion of znuA virulence factor attenuates Brucella abortus and confers protection against wild-type challenge.. Infection and immunity. 2006; 74(7); 3874-3879. [PubMed: 16790759]. 3874-3879. [Pubmea: 107907.09].
   Luo D, Ni B, Li P, Shi W, Zhang S, Han Y, Mao L, He Y, Wu Y, Wang X Protective immunity elicited b
   Luo D, Ni B, Li P, Shi W, Zhang S, Han Y, Mao L, He Y, Wu Y, Wang X Protective immunity elicited b
- livalent DNA vaccine encoding both the L7/L12 and Omp16 genes of Bruce ce., Infection and immunity, 2006; 74(5); 2734-2741. [PubMed: 16622210]. Vemulapalli R, Contreras A, Sanakkayala N, Sriranganathan N, Boyle SM, Schurig GG Enh
- of recombinant *Brucella abortus* RB51 vaccines against *B. melitensis* i ary *microbiology*, 2004 Sep 8; 102(3-4); 237-45. [PubMed: 15327798].
- Rupprecht CE, Hanlon CA, Slate D Oral vaccination of wildlife against rabies: opportunities and challenges in prevention and control.. Developments in biologicals. 2004; 119; 173-184. [PubMed
- Pasquali P, Rosanna A, Pistola C, Petrucci P, Ciuchini F Brucella abortus RB51 induces protection mice orally infected with the virulent strain B, abortus 2308, Infection and immunity, 2003; 71(5); 2326-2330, [PubMed: 12704101]. He Y, Vemulapalli R, Schurig GG Recombinant Ochrobactrum anthropi expressing Br
- iperoxide dismutase protects mice against *B. abortus* infection only after switching of imr es to Th1 type.. *Infection and immunity*. 2002 May; 70(5); 2535-43. [PubMed: 11953393].
- Schurig GG, Sriranganathan N, Corbel MJ Brucellosis vaccines: past, present and future... Veterinary microbiology. 2002 Dec 20; 90(1-4); 479-96. [PubMed: 12414166].
- Ko J, Gendron-Fitzpatrick A, Ficht TA, Splitter GA Virulence criteria for Brucella abortus strains as determined by interferon regulatory factor 1-deficient mice.. Infection and immunity. 2002; 70(12) 7004-7012. [PubMed: 12438380].
- Novino La Forma Da, Michael A, Azevedo V, Campos E, Cravero SL, Rossetti C, Spiller G, Oliveira SC. Mentication and characterization of a *Bruestia about* ATP-binding cassette transpor homolog to *Rhizobium mellioti* Essa and its role in virulence and protection in mice... *Mection and Immunity*. 2002; 70(9): 5036-504. [PubMed: 12163550]. He Y, Vemulapalli R, Zeytun A, Schurig GG Induction of specific cytotoxic lymphocytes in mice vaccinated with Brucella abortus RB51. Infection and immunity. 2001 Sep; 69(9); 5502-8. [PubMed
- Vemulapalli R, He Y, Cravero S, Sriranganathan N, Boyle SM, Schurig GG Overexpression
   Vemulapalli R, He Y, Cravero S, Sriranganathan N, Boyle SM, Schurig GG Overexpression
- protective antigen as a novel approach to enhance vaccine efficacy of Infection and immunity. 2000 Jun; 68(6); 3286-9. [PubMed: 10816475].
- Vermulapalli R, He Y, Buccolo LS, Boyle SM, Sriranganathan N, Schurg GG Complementation of Brucella abortus RES with a functional wood gene results in O-antigen synthesis and enhanced vaccine efficacy but no change in rough phenotype and attenuation... Infection and immunity. 2000 Jul; 68(7): 3927-32. [PubMed: 10655205]. 18 Cheville NF Development, testing and commercialization of a new brucellosis vaccine fo Annals of the New York Academy of Sciences. 2000; 916; 147-53. [PubMed: 11193615].
- 19. Mackowiak M, Maki J, Motes-Kreineyer L, Harbin T, Van Kampen K Vaccination of wildlife agai rables: successful use of a vectored vaccine obtained by recombinant technology. Advances in veterinary medicine. 1999; 41; 571-583. [PubMed: 980044].
- Winter AJ, Schurig GG, Boyle SM, Sriranganathan N, Bevins JS, Enright FM, Elzer PH, Kopec JD Protection of BALB/c mice against homologous and heterologous species of Brucella by much st
- vaccines derived from Brucella methodologuus and heterologous spacies of Brucella Muper J research. 1996 May; 57(5); 677-83. [PubMed: 8723861]. Schurg GG, Rono PM 9-1 er -----Schurig GG, Roop RM 2nd, Bagchi T, Boyle S, Buhrman D, Sriranganathan N Biological propertie RBST: a stable rough strain of *Brucella abortus.*, Veterinary microbiology, 1991 Jul; 28(2); 171-88 [PubMed: 1905158]. 21
- 22.
- Brochier B, Kleny MP, Costy F, Coppens P, Bauduin B, Lecocq JP, Languet B, Chappuis G, Desmettre P, Aflademanyo K Large-scale eradication of rables using recombinant vaccinia-vaccine...*Nature*. 1991: 354(6354): 520-522. (PubMed: 1758494).

NOTE: Go to Litesearch, Vaxpresso and Vaxmesh for advanced literature search.

### REFERENCES

- Xiang, Zuoshuang. "VIOLIN: Vaccine Investigation and Online Information Network." VIOLIN: Vaccine Investigation and Online Information Network. University of Michigan Medicine School, n.d. Web. 29 Sept. 2013.
   <a href="http://www.violinet.org/protegen/">http://www.violinet.org/protegen/</a>>.
- Yang, Sayers, Xiang, et al. "Protegen: a web-based protective antigen database and analysis system." 16 August 2010.