



# Rabies surveillance and control programs in Cambodia

Philippe Dussart, PharmD, PhD

Head of Virology Unit, Institut du Cambodge

2018 Launching Meeting of OIE Twinning Project for Rabies between Anses-Nancy and AHRI – Taiwan – 17-18 October 2018



# Institut Pasteur du Cambodge - 2018





- The IPC is a **non-profit, international research organization** of public utility under the patronage of the Cambodian Ministry of Health
  - The missions of IPC are:
    - 1. Support to public health
    - 2. Research in public health in Cambodia and in Southeast Asia
    - 3. Training in research and public health
- For the past 65 years in Cambodia, the main activities of IPC have been focused on the fight against **infectious diseases and emerging pathogens** in Cambodia and in Southeast Asia.
- Member of the Institut Pasteur International Network
- 230 staff 4 reference lab (Influenza, Arboviruses, Rabies, Malaria RDT) – 6 research units & platforms – BSL3 & BLS2 facilities, clinical diagnostic laboratory, vaccine center, water and food safety laboratory



# **RABIES RISKS IN CAMBODIA**

# Dog bites injuries are frequent in Cambodia

- Around 600,000 dog bites per year
  - 3-4 injuries for 100 persons per year
  - 80% are severe injuries (WHO category 3)
  - 60% among children under 17 years old
  - ➔ An extremely high probability of being bitten in a rabies-endemic country
  - Only around 5% consulted a rabies vaccination center
  - → 95% individual did not receive proper rabies prophylaxis after dog bites







# **Evidence of virological burden is missing in Cambodia**

- Around 5 millions dogs -> ~1 dog for 3 humans
  - Most not vaccinated against rabies
  - Most are owned dogs, but range freely in the community
- No veterinary surveillance or quarantine
- Biting dogs:
  - Are put down
  - Escape
  - But also consumed for food
- No (or low) virological confirmation despite competent lab







# Rabies caused human death in Cambodia

- Mathematical estimation in 2007
  - Based on data collected from IPC vaccination center (PEP)
- 810 human rabies deaths in 2007
- Incidence = 5.8 / 100,000 inhabitants
- That year: 54 cases notified

Ly S et al., 2009

# **Rabies Post-Exposure Prophylaxis at IPC**

- Since 1995 in Phnom Penh
- > 20,000 new patients per year since 2008
- 2017: 22,421 patients
- 60% among children under 17 years old
- 50% from Phnom Penh and Kandal (Southern Cambodia)

Rabies Vaccination Center, IPC



New Patients who Received PEP at IPC





# **Origins of patients who received Rabies PEP**



#### Data from IPC, 2013-2016, N= 64,158 patients



OIE Twinning Project for Rabies between Anses-Nancy and AHRI, Taiwan – 17-18 October 2018

Randomized dots by province

- Genome detection: RT-PCR

### **Participation to Inter-laboratory test for Rabies diagnostic**

Organised by the Nancy Laboratory for Rabies and Wildlife from ANSES, France (European Union Reference Laboratory (EURL) for Rabies)

### **Perspective: set-up of seroneutralization assays**

- FAVN assay with support from ANSES Nancy (dog samples; in progress)
- RFFIT assay with support from Institut Pasteur, Paris (human samples)

OIE Twinning Project for Rabies between Anses-Nancy and AHRI, Taiwan - 17-18 October 2018

# **Rabies diagnosis at IPC**

- Lab testing at IPC
  - Virology Unit: reference laboratory for rabies diagnosis
  - Free of charge for PEP patients
  - Animal brains (98% from dogs)
    - Cat, swine, cattle, monkey
  - Antigen detection: DFAT







## Results in dogs, 2002-2017

- 226 samples per year
- 50% positive for rabies virus
- Around 2 positive dogs per week

#### OIE Twinning Project for Rabies between Anses-Nancy and AHRI, Taiwan – 17-18 October 2018

COont, Dogs tested

Testing in Dogs for Rabies, Institut Pasteur in Cambodia (Direct Immunoflurescence Assay)

#### 400 80% 350 70% 60% 300 50% 250 200 40% 150 30% 100 20% 50 10% 0 2012 2013 2015 2016 2017 2002 2003 2005 2006 2009 2010 2011 2014 2008 2018 (Jan-Sep) 2004 2007 ---% of positive Tested

# **Dogs tested positive for Rabies**



of positive dogs

\*

# **Origins of lab confirmed Rabid dogs**



#### Data from IPC, 1998-2016, N=1,842 samples



OIE Twinning Project for Rabies between Anses-Nancy and AHRI, Taiwan – 17-18 October 2018

Randomized dots by province



# IMPROVEMENT OF RABIES PREVENTION IN CAMBODIA

#### Reduce **distance**

so people don't have to travel and make hard choices



#### Reduce duration

so people don't have to travel pay taxis several times



#### Reduce cost

so people don't have to contract debts to get PEP



### Reduce **distance**

so people don't have to travel and make hard choices



## Objective

- Facilitate access to rabies PEP
- By bringing rabies PEP center closer to the Cambodian population
- ➔ Reduce number of human rabies death by half over 5 years
- How?
  - Opening a second Rabies Prevention Center
- Where?
  - In Northwestern Cambodia: Battambang province



#### Reduce **distance**

so people don't have to travel and make hard choices



### **Battambang province**



# **Opening of the center**

Open to public on 27 July 2018 



Inauguration 27-28 September 2018 



OIE Twinning Project for Rabies between Anses-Nancy and AHRI, Taiwan – 17-18 October 2018





មជ្ឈមណ្ឌល



# **Communication, Education and Information on rabies and its prevention**



The most deadly disease known
No cure, only palliative care
100% lethal disease

Human Rabies is 100% preventable by vaccination after infections caused by animals

An acute neurological syndrome due to a virus

Infects domestic and wild animals

Spreads to people through close contact with animal infected saliva via bites or scratches

In Cambodia around 800 deaths a year

Implementation of Cambodia Rabies National Control Program in progress (MOH, MAFF)

As an ASEAN Member State, Cambodia has committed to eliminate rabies by 2025

#### RABIES AND DOGS

• In Cambodia rabies mostly transmitted by dogs

More than 4 million dogs in Cambodia
 Most of them not vaccinated

 Most dogs in Cambodia have owners, only few are stray dogs

#### WHEN BITTEN WHAT TO DO ?

• You have to seek for vaccination against rabies immediately for example at Pasteur Phnom Penh

The intradermal vaccination consists of 4
 sessions (Day 0, 3, 7 and 28) for a total cost
 of 12 USD
 Immunoglobulins are required in addition to the
 vaccine in case of severe wound

 More than 20 000 patients per year are vaccinated at Pasteur Phnom Penh Children under 17 years old : 60%



# Remedicence and and and a set of a set

### As its rabies plan lags,

The Phnom Penh Post

Cambodia feels the bite

Despite killing some isoo people in the Kingdom each year, rabies has been largely off the public health radiar. Plans are in the works for a nationwide response, but can funding each up?



we been received, one of four rounds of some exposure rables vectored a re Penh this month. Hone Menee

When her 15-year-old desylter Thai Sophesk was bitten by a dog last mosth, Kat Solitin took no shances. Die had know is man from a nearby vilage who died following a dog bite the year before, and vie warrit going to let her desylter softer the same fem.

September of the lady over the the part month, the basises leading ter village in Kampong Charris Rey Charr district every leadapant of the manning to beact in Phone Perch. The destination is the Partner further to comparison output, when a the carriest approximation is before the tradition of the



- Post-exposure prophylaxis
- 4 sessions
  - Day 0, D3, D7, D21/28
- With or without immunoglobulins
- Reduce the number of injection

→ Prospective documentation of antibody response after 3 and 4 sessions in people bitten by tested dog

#### Reduce duration

so people don't have to travel pay taxis several times



#### Reduce cost

so people don't have to contract debts to get PEP





# **New PEP protocol recommended by WHO**



Table 1 Post-exposure prophylaxis (PEP) by category of exposure

Tableau 1 Prophylaxie post-vaccinale par catégorie d'exposition

	Category I exposure – Exposition de catégorie I	Category II exposure – Exposition de catégorie II	Category III exposure – Exposition de catégorie III
Immunologically naive individuals of all age groups – Individus jamais encore immunisés quel que soit le groupe d'âge	Washing of exposed skin surfaces. – Lavage des surfaces cutanées exposées. No PEP required. – Aucune PPE n'es requise.	Wound washing and immediate vaccination: – Lavage de la plaie et vaccination immédiate: • 2-sites ID on days 0, 3 and 7 <sup>73</sup> – Injections ID en 2 sites aux jours 0, 3 et 7 <sup>73</sup>	<ul> <li>Wound washing and immediate vaccination – Lavage de la plaie et vaccination immédiate</li> <li>2-sites ID on days 0, 3 and 7<sup>73</sup> – Injections ID en 2 sites aux jours 0, 3 et 7<sup>73</sup></li> <li>Or – Ou</li> <li>1-site IM on days 0, 3, 7 and between day 14–28<sup>74</sup> – Injections IM en 1 site aux jours</li> </ul>
		<ul> <li>1-site IM on days 0, 3, 7 and</li> <li>between day 14, 2974</li> </ul>	0, 3, 7 et entre 14 et 28 jours <sup>74</sup>
Dne-week, 2-site ID regimen Institut Pasteur du Cambodge regimen: 2-2-2-0-0 Duration of entire PEP course: 7 days		<ul> <li>Detween day 14–28<sup>77</sup> –</li> <li>Injections IM en 1 site aux jours</li> <li>0, 3, 7 et entre 14 et 28 jours<sup>74</sup></li> <li>or – ou</li> <li>2-sites IM on days 0 and 1-site</li> <li>IM on days 7, 21<sup>75</sup> – Injections IM en</li> </ul>	<ul> <li>2-sites IM on days 0 and 1-site IM on days 7, 21<sup>75</sup> – Injections IM en 2 sites au jour 0 et en 1 site aux jours 7, 21<sup>75</sup></li> <li>RIG administration is recommended. – L'administration d'immunoglobuline</li> </ul>
		<ul> <li>2 sites au jour 0 et en 1 site aux jours 7, 21<sup>75</sup></li> <li>RIG is not indicated. – L'immuno- globuline antirabigue n'est pas indiguée</li> </ul>	antirabique est recommandée.

international network

World Health Organization. Rabies vaccines: WHO position paper. Wkly Epidemiol Rec 2018; 2018: 201–20.



# RECENT AND CURRENT RESEARCH ACTIVITIES IN ANIMAL

# Study of dog population, vaccination coverage and bite risk factors

- 2 Cambodian provinces (10 villages): 2017-2019
  - Kandal and Battambang
- **2 year follow-up**, with door to door census
- Dog identification, vaccination and sampling

## • Questionnaires

- Dog characteristics
- Dog use
- Dog slaughter value chain
- Assessment Knowledge, Attitude and Practice of dog owners, animal health and public health professionals towards rabies
- Bite incidence
- Bite risk factors









# **Preliminary result in Kandal**

- 5,400 identified dogs
- Ratio dog/human : 1:4
- High turn-over rate
- Dog trade and consumption
- Bite incidence rate : 2-4%/year
- Risk of being bitten increases
  - With number of dog per family
  - Where dogs are sold for consumption
  - With number of children < 15 years old
- Good acceptability for dog vaccination
- But poor knowledge of rabies







Strata 🛨 Male 🕂 Female

#### **Survival rate**

1 y/o : 0.62 (IC 0.59-0.65) 2 y/o : 0.49 (IC 0.46-0.52) 5 y/o : 0.19 (IC 0.17-0.23)



## Way forward

- Neutralization assay: FAVN tests 

   vaccination coverage dynamic
- Modelling → adapted vaccination strategies
- Phylogeographic models 

   better understanding of spread mechanisms of rabies virus
- Many dogs are aggressive and can not be handled, even by owners → evaluation of oral vaccination may be useful completing parenteral vaccination
- Involvement of all stakeholders and inter-sectoral collaboration









# **National Plan to Fight against Rabies**



National Guideline for Rabies Surveillance, Prevention and Control for Cambodia

December 2017

Ministry of Health Ministry of Agriculture, Forestry and Fisheries

```
ព្រះរាជាឈាចគ្រងអគីជា
                  ខាឌិ សាសនា ព្រះមហាភ្យគ្រ
       សៀទនៅមគ្គខ្លេសក៏ខាគិស្តីពី
       ສາເສເອສສາຍສາຂ ສາເຍອາເ
តិខភាររួមយុន្តតី១៩ំខឺឆ្លែឆ្លួងនៅអូខរួមនេស
                  ខែជ្ ឆ្នាំ២០១៧
```

# In conclusion



- ✓ Institut Pasteur du Cambodge
- Ministry of Health
- $\checkmark$  Ministry of Agriculture, Forestry and Fisheries
- Ministry of Education, Youth and Sports
  - Aim to decrease the number of death from 800 to 400 deaths per year
  - Save 2,000 lives in 5 years
  - Help to reach national coverage of rabies prevention



- Stating the obvious
  - Rabies, as neglected disease, is still a major public health concern in Cambodia
  - → Fund raising remains challenging
  - → Funds and vaccines to study dog population dynamic and define the best vaccination strategy for Cambodia









# Acknowledgements



- Didier Fontenille
- Sowath Ly
- Véronique Chevalier
- Laurence Borand
- Tineke Cantaert
- Arnaud Tarantola



Hervé Bourhy



**cirad** 

international network

Institut Pasteur

• Florence Cliquet









World Organisation

for Animal Health



