## **Original Article**

# A critical appraisal of usage of rabies immunoglobulin in an anti-rabies clinic in India\*

M.K.Sudarshan<sup>1</sup>, D.H.Ashwath Narayana<sup>2</sup> and Giriyanna Gowda<sup>3</sup>

Rabies is a practically 100% fatal viral encephalomyelitis. It is transmitted to man mostly following bite by a rabid dog or cat. In India annually an estimated 20,000 human rabies deaths occur consequent to about 17 million animal/dog bites<sup>1</sup>. Following bite by a rabid animal, administration of a modern rabies vaccine using a regimen approved by World Health Organization (WHO) stimulates production of neutralizing antirabies antibodies by the patient's immune system. Protective levels of these antibodies are seen 7 to 14 days after the first dose of vaccine. Adequate serotitres can be expected in all the vaccinees by day 14. When the bites are on the head, neck, face & hands, the incubation period will be shorter. Thus, such patients are vulnerable to develop rabies during this window period of 7 to 14 days despite the timely and full course of any antirabies vaccine and wound care. The rabies immunoglobulins (RIGs) are readymade antirabies antibodies which provide passive immunity and thus offer immediate protection. Hence, administration of RIGs, after thorough cleansing of wounds, following severe (category III) exposure to rabies is life saving as their timely and proper administration neutralizes the virus in the wound and aborts the risk of developing rabies.

According to WHO all of the RIG, or as much as anatomically possible (but avoiding possible compartment syndrome), should be administered into or around the wound site or sites. The remaining immunoglobulin, if any, should be injected intramuscularly at a site distant from the site of vaccine administration. RIG may be diluted to a volume sufficient for all wounds to be effectively and safely infiltrated<sup>2</sup>.

These guidelines are followed by Government of India<sup>3</sup> and professional bodies like Association for Prevention and Control of Rabies in India<sup>4</sup>. However, some experts consider systemic administration of RIGs as wasted<sup>5</sup>. At a recent expert consultation of WHO, the usefulness of systemic administration of RIG was debated and recommended that new in-vitro and in-vivo research be encouraged to determine the quantity of RIG (in IU) required on site, with or without distal parental RIG administration<sup>6</sup>.

Hence, this study was conducted with the objective of finding out the ratio of volume of RIG injected into/around the wound/s (local infiltration) to the volume of RIG injected by intramuscular (systemic) route. In this context, the data from the antirabies clinic of Kempegowda Institute of Medical Sciences, Bangalore, a centre specialized for

Table-I: Details of exposure to rabies

SI. No.		Exp	osure to rabies	No.	%
1	Total cases			6664	100
	Categ	ory III (	severe exposure) cases	3109	46.6
	Biting	anima	/rabies exposure	3109	100
	Dog			2918	93.8
	Monk	ey		98	3.2
2.	Cat			45	1.5
	Wilda	animals		23	0.7
	Exposure to rabid cow's milk/secretion			19	0.6
	Exposure to case of Hydrophobia			6	0.2
	Profile of Patients/subjects			3109	100
	Sex	Male		2182	70.2
		Fema	ale	927	29.8
3.	Age Groups		0-4	418	13.5
	(in ye	ars)	5-14	907	29.2
			15-44	1207	38.8
			45-64	434	13.9
			65+	143	4.6
4.	Type of animal bite wounds / injuries (multiple responses)			3345	100
	Abrasions/scratches			1589	47.5
	Lacerations			1284	38.4
	Transdermal bite / puncture			330	9.9
	Cuts			138	4.1
	Avuls	ions/fr	actures	4	0.1
5.	Distribution of wounds (multiple responses			3366	100
	Lowe	r limbs		1763	52.4
	Uppe	r limbs		961	28.5
	Head	/neck		421	12.5
	Trunk			207	6.2
	Genit	alia		14	0.4

<sup>&</sup>lt;sup>1</sup>Dean/Principal and Professor, <sup>2</sup>Professor, <sup>3</sup>Lecturer/Asst. Professor, Department of Community Medicine, Kempegowda Institute of Medical Sciences, Bangalore 560 070, INDIA [Corresponding author: <a href="mailto:mksudarshan@gmail.com">mksudarshan@gmail.com</a>]

<sup>\*</sup>This paper was presented at the third rabies in Asia conference (RIACON) organized by Rabies in Asia foundation from November, 28-30, 2011 at Colombo, Sri Lanka

Table-2
Usage of RIGs in persons exposed to rabies

S1. No.	Usage of RIGs	No.	%			
1	Rabies exposed persons who received RIGs	3109	100			
	ERIG	2904	93.4			
	HRIG	199	6.4			
	Refused / could not afford / referred to Go hospital	ovt. 4	0.6			
2.	Volume (in ML) of RIGs received					
	Children (<15 years)	1323	-			
	Mean	2.85 ML	-			
	Range	0.4 to 10.00 ML	-			
	Adults (>15 years)	1780	-			
	Mean	7.89 ML				
	Range	3.30 to 10.00 ML				
3.	Volume (in proportion) of RIGs used by different routes of administration					
	ERIG	2904	-			
	Local infiltration only	-	73.5			
	Local infiltration plus systemic injection	-	25.8			
	Systemic injection only	-	0.7			
	HRIGs	199	-			
	Local infiltration only	-	74.6			
	Local infiltration plus systemic injection	-	23.0			
	Systemic injection only	-	2.4			

administration of RIGs, was analyzed for 3 years duration from July, 2008 to June, 2011. In this centre the dosage of RIG required for each patient was calculated as per the guidelines of the manufacturer. When the calculated dose of RIG was not adequate sterile normal saline was used as diluent to make up for the volume sufficient to infiltrate all wounds.

A total of 6664 cases of rabies exposures were treated during this period and of these 3109 (46%) were categorized as "severe exposures "or belonging to Category III. This lower percentage/proportion of category III exposures is due to the fact that many trivial and minor exposures come for consultations and counseling and others visit /referred from outside centres come to receive vaccine only. Majority (93%) of these were due to dog

bites. The victims were mostly males (70%), adolescents and adults (57%). The bite wounds were mostly abrasions/ scratches (47%) & lacerations (38%) and found on the upper and lower limbs (81%) (Table-1). In about 93% cases ERIG was used, in 6% HRIGs and the remaining ( $\sim$ 1%) could not afford / referred/refused RIG. In children (<15 years age) the mean volume of RIG used was 2.85 mL per patient and in adolescents/ adults (>15 years) 7.89 mL. In these cases about three fourths ( $\propto$  74%) of the total dose/ volume of RIG was used exclusively for local infiltration of wounds, 25% injected both locally and systemic and 1% injected by only systemic route (Table-2).

In conclusion, the usage of RIGs was found be in accordance with the current recommendations of WHO. Besides it shows that only about one fourth (25%) volume of the total dose of RIG was injected systemically or the ratio of local: systemic administration of RIG was 3: 1.

### **Acknowledgments**

The authors thank the post graduates of community medicine viz. Drs.Ramesh Holla, V.Veena, U.Malatesh, AR Rachana and interns Drs.Kavitha, Jawad, Manoj, Manjusha, Mona, Manjula, Shantanu and Abhinav for compiling and collating the data.

#### References

- Sudarshan MK, Madhusudana SN, Mahendra BJ, Rao NSN, Ashwath Narayana DH, Rahman SA, et al. Assessing the burden of human rabies in India: results of a national multicentric epidemiological survey. Int. J Infect. Dis. 2007; 11:29-35.
- World Health Organization (WHO). Rabies vaccines, WHO position paper. Weekly Epidemiological Record, 6 Aug 2010;32(85):309-20.
- National Institute of Communicable Diseases. National guidelines for rabies prophylaxis and intradermal administration of cell culture rabies vaccines, 2007, New Delhi, India.
- 4. Association for Prevention and Control of Rabies in India. Manual on rabies immunoglobulin administration. Feb. 2009, Bangalore, India.
- David C Anderson. WHO guidelines dealing with immunoglobulin use impede rabies prevention. Asian Bio-medicine. June.2007; 1(1):103-7.
- World Health Organization. Human and dog rabies prevention and control, Report of the WHO/ Bill & Melinda Gates Foundation Consultation, Annecy, France, 2009.

## **Please Visit**

The APCRI web site at www.apcri.org for all information about APCRI