



University of Miyazaki

## The identification and characterization of Interleukin-17 family ligands and receptors in fish

Hiroki Korenaga\*, Tomoya Kono, Haruka Kuse,  
Hiroaki Takayama and Masahiro Sakai

University of Miyazaki, JAPAN

6th International Symposium on Aquatic Animal Health



### - Contents -

- Search and analysis of teleost IL-17L/R genes
  1. Analysis of IL-17 ligand genes (medaka)  
genome synteny, homology analysis,  
expression study (in un-stimulated tissues,  
LPS stimulated intestine)
  2. Analysis of IL-17 receptor genes (*Fugu*)  
genome synteny, homology analysis

## IL-17 ligands isolation

Human	IL-17 family ligands	secreting cells	main functions
	IL-17 (IL-17A)	T cells (T <sub>H17</sub> )	Neutrophil recruitment and immunity to extracellular pathogen
	IL-17F	T cells (T <sub>H17</sub> )	
	IL-17B	Numerous tissues	Pro-inflammatory ?
	IL-17C	Numerous tissues	
	IL-17D	Numerous tissues	
	IL-17E (IL-25)	Th2 cells	Induces Th2 immune responses

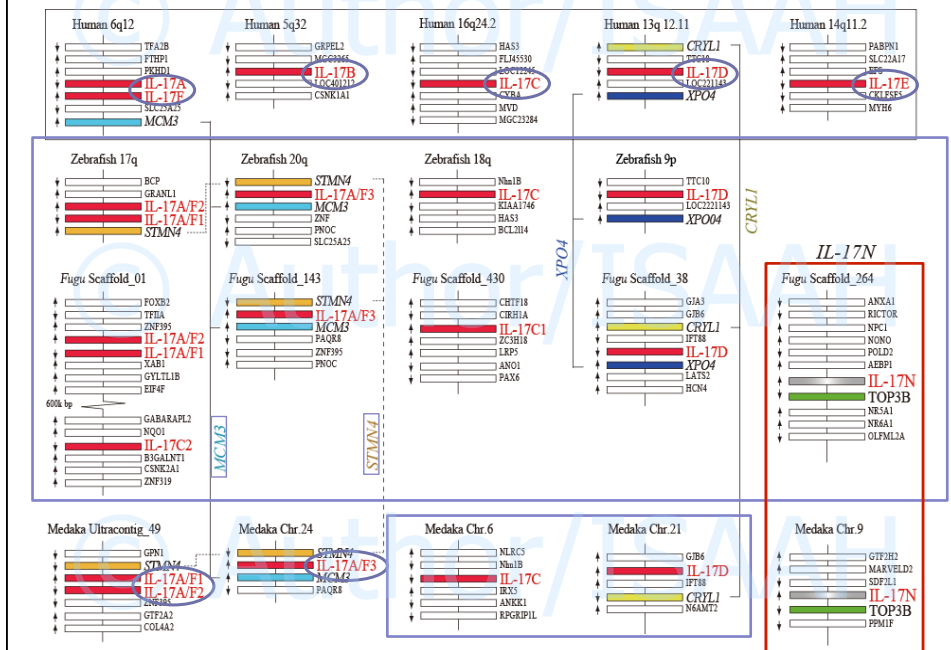
### Teleost

IL-17 family ligands	Fish species
IL-17A/F	zebrafish, trout, salmon, <i>Fugu</i>
C	zebrafish, trout, <i>Fugu</i>
D	zebrafish, trout, salmon, <i>Fugu</i>
N	<i>Fugu</i>

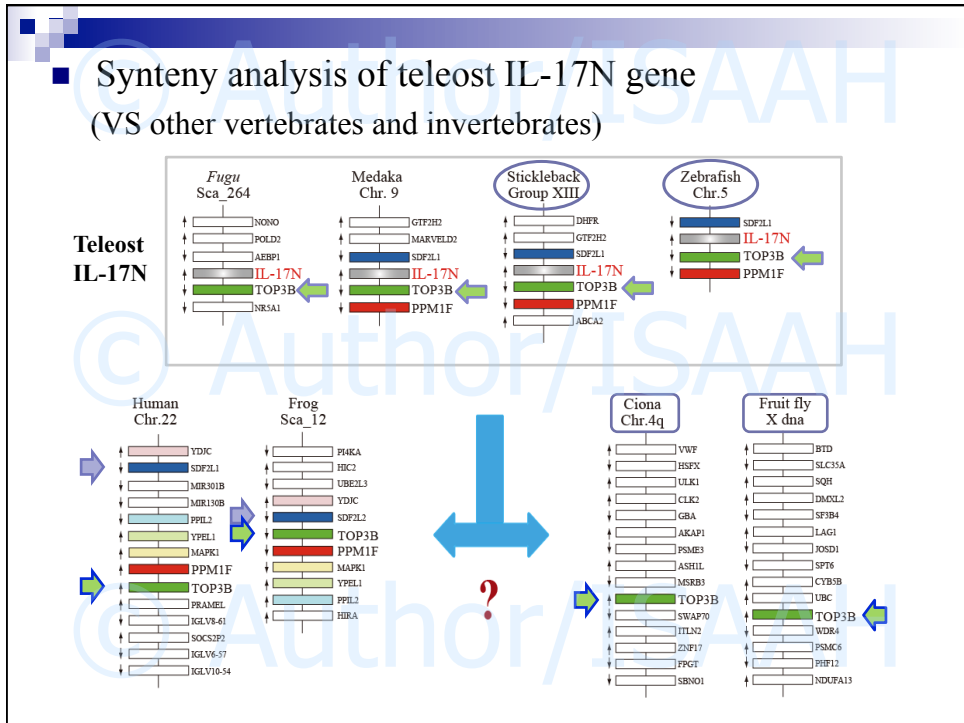
### Invertebrate

IL-17 family ligands	species
IL-17	<i>Ciona</i> , Oyster, Sea urchin

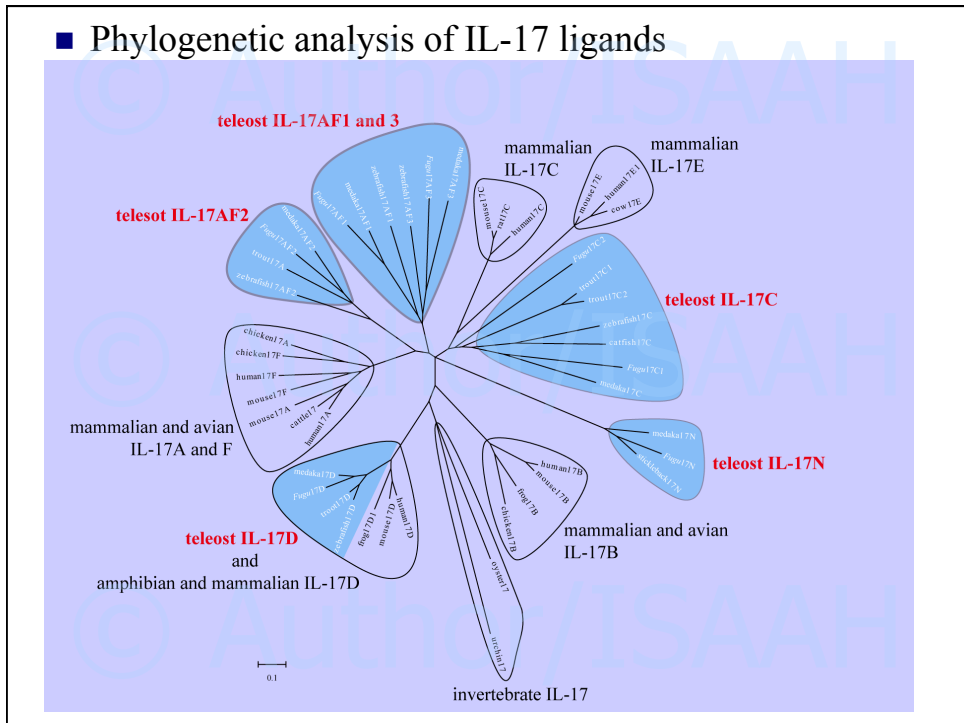
## Synteny analysis of IL-17 genes (Human and Teleost)



■ Synteny analysis of teleost IL-17N gene (VS other vertebrates and invertebrates)



■ Phylogenetic analysis of IL-17 ligands



## ■ Expression analysis of IL-17 ligands in medaka



tissues:

Intestine, Kidney, Spleen, Liver, Heart,  
Brain, Skin, Muscle, Gill

Intestine stimulated with  
LPS(20  $\mu$ g/ml) for 1, 4, 8,  
12 and 24 h

Total RNA extraction



cDNA synthesis

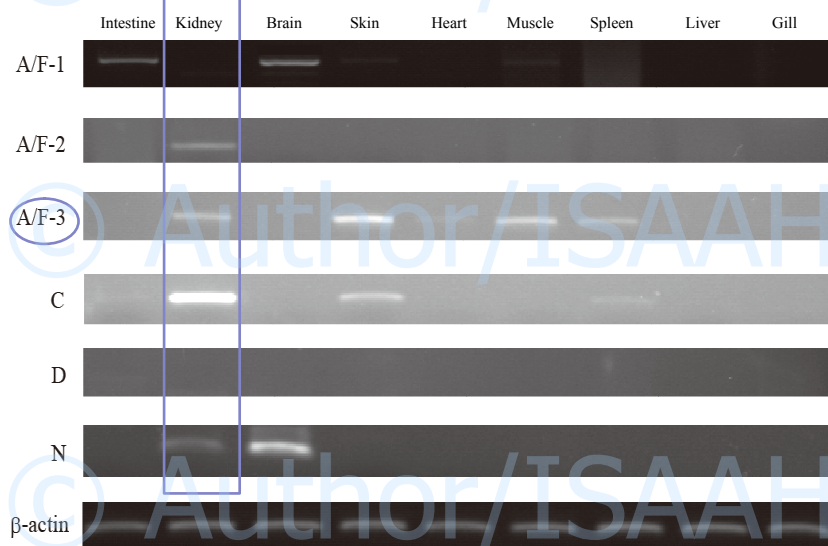


PCR using specific primers for IL-17 /  $\beta$ -actin genes

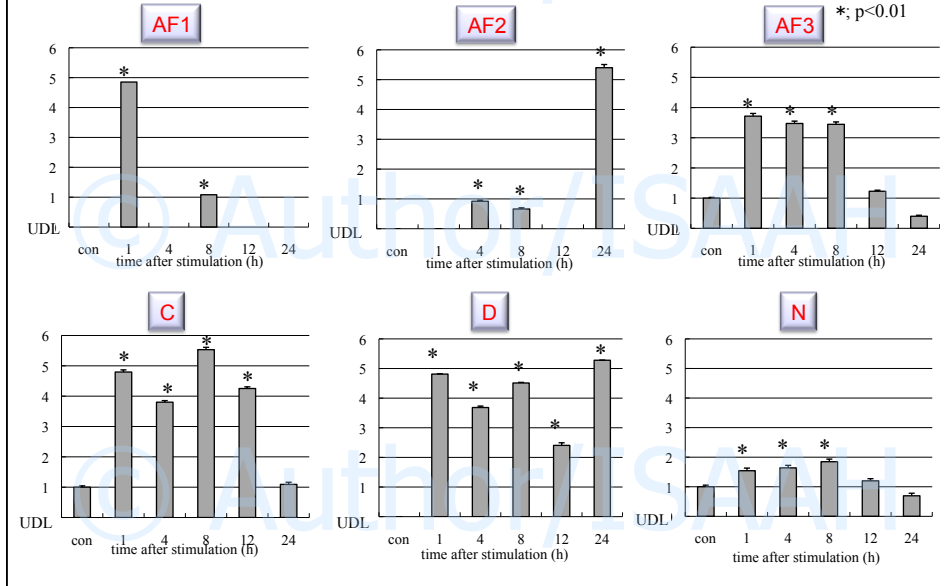


Semi-quantitative analysis

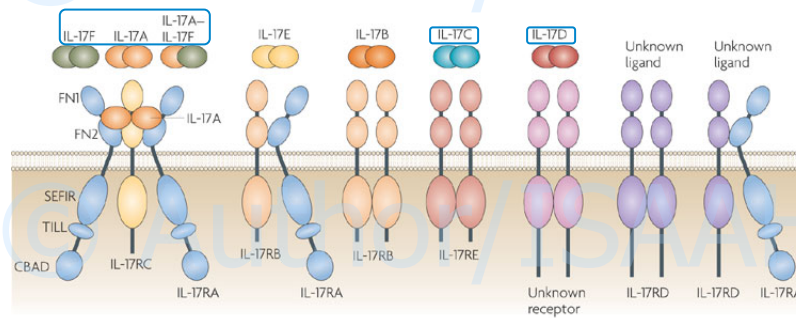
## ■ Expression level of IL-17 family genes in organs



## ■ Expression level of IL-17 family genes in intestine stimulated with LPS



## ■ IL-17 receptors isolated from human and teleost



Legend:  : Ligands isolated in teleost

Nature Reviews | immunology

**Teleost**

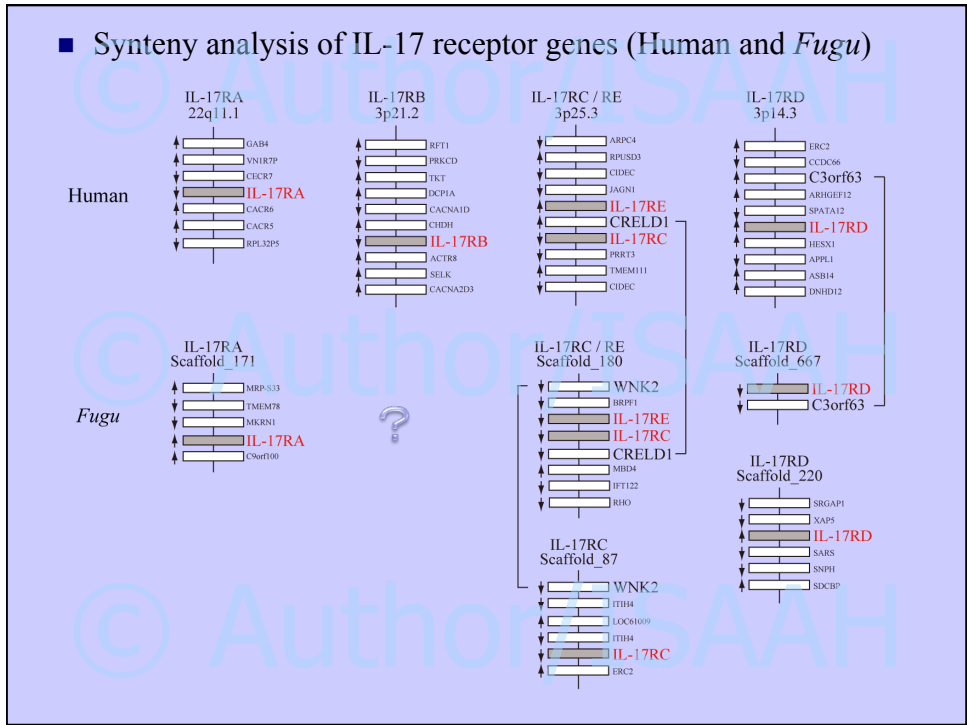
*IL-17 receptors*

IL-17RA  
IL-17RD

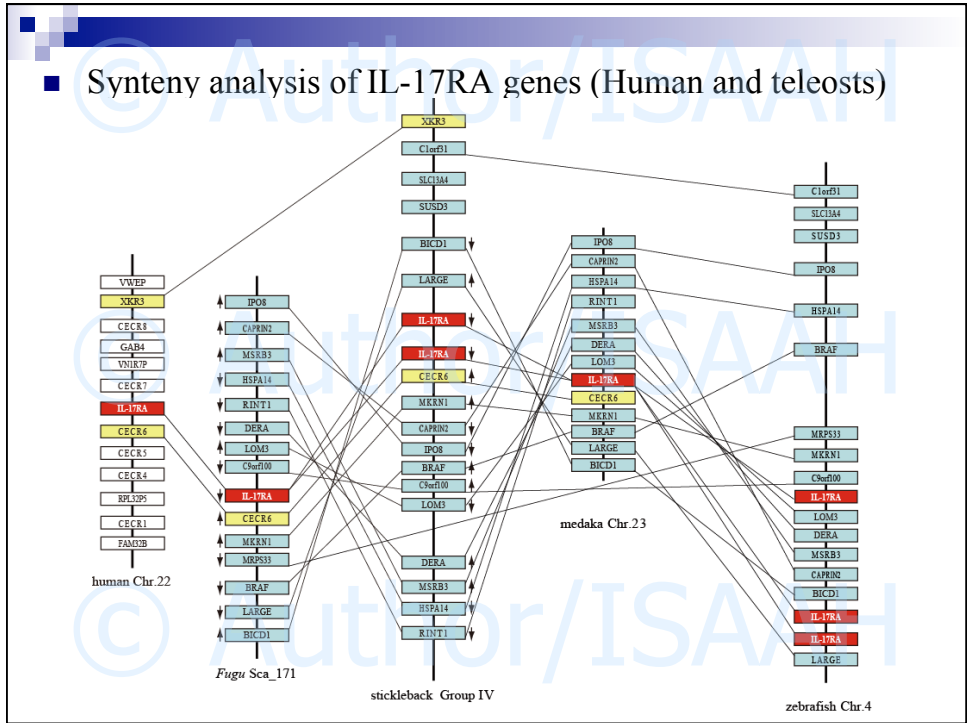
*Fish species*

salmon, trout  
salmon, trout, zebrafish

■ Synteny analysis of IL-17 receptor genes (Human and *Fugu*)



■ Synteny analysis of IL-17RA genes (Human and teleosts)



- Conclusions -

- IL-17 family genes (A/F-1, 2, 3, C, D, N) were isolated from medaka
- IL-17N might be unique ligand in teleost
- The expression profile of medaka IL-17 family genes in tissues was quite different in each subtype
- Medaka IL-17 family may be involved in immune response because the expression of the genes was increased by the stimulation with LPS
- IL-17RA, RB, RC, RD, RE homologue genes were confirmed on fish genome.

