



Code No. KAL-KH001-04-EX

For research use only

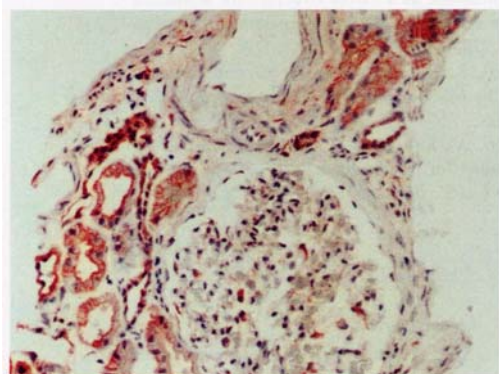
**Advanced Glycation End Products (AGE)  
Anti AGE Monoclonal Antibody,  
FITC conjugated**

(Clone No. 6D12)

Reaction of protein amino groups with glucose leads, through the early products such as a Schiff base and Amadori rearrangement products, to the formation of advanced glycation end products (AGE). Recent immunological studies using anti-AGE antibody (6D12) have demonstrated the presence of AGE-modified proteins in several human tissues: ( i ) human lens (nondiabetic and noncataractous), ( ii ) renal proximal tubules in patients with diabetic nephropathy and chronic renal failure, ( iii ) diabetic retina, ( iv ) peripheral nerves of diabetic neuropathy, ( v ) atherosclerotic lesions of arterial walls, ( vi )  $\beta_2$ -microglobulin forming amyloid fibrils in patients with hemodialysis-related amyloidosis, ( vii ) senile plaques of patients with Alzheimer's disease, ( viii ) the peritoneum of CAPD patients, ( ix ) skin elastin in actinic elastosis, and ( x ) ceroid/lipofuscin deposits. These results suggest that AGE-modification has a potential role of AGE-modification in normal aging as well as age-enhanced disease processes. This antibody named as 6D12 has been used to demonstrate AGE-modified proteins in these human tissues, indicating potential usefulness of this antibody for histochemical identification and biochemical quantification of AGE-modified proteins.

Package Size	10 $\mu$ g (40 $\mu$ L/vial)
Format	Mouse monoclonal antibody 0.25 mg/mL
Buffer	Block Ace as a stabilizer, containing 0.1%Proclin as bacteriostat
Storage	Store below $-20^{\circ}\text{C}$ . Once thawed, store at $4^{\circ}\text{C}$ . Repeated freeze-thaw cycles should be avoided.
Clone No.	6D12
Subclass	IgG1
Purification method	The splenic lymphocytes from BALB/c mouse, immunized with AGE-BSA were fused to myeloma P3U1 cells. The hybrid cells were screened, and the cell line (6D12) with positive reaction to AGE-human serum albumin but negative to BSA was selected through successive subclonings and grown in ascitic fluid of BALB/c mouse, from which the anti-AGE antibody was purified by Protein G affinity chromatography. (Reference No.1)

Working dilution for immunohistochemistry: 2  $\mu$  g/mL; for ELISA: 0.1-0.5  $\mu$  g/mL



Immunohistochemical staining of renal proximal tubules and glomeruli in patients with diabetic nephropathy, using anti-AGE antibody 6D12.

Yamada, K. et al,  
*Clinical nephrology*, Vol.42, 354-361, 1994



Immunohistochemical staining of the early stage of human atherosclerotic lesions of the aorta with anti-AGE antibody 6D12.

Kume, S. et al,  
*American Journal of Pathology*, Vol.147, 654-667, 1995



Code No. KAL-KH001-04-EX

## Anti AGE Monoclonal Antibody

(Clone No. 6D12)

### 【Specificity】

The initial study (Ref. 1) revealed that 6D12 does not recognize early products (Schiff base and Amadori products), but shows a positive reaction to AGE-samples obtained either from proteins, lysine derivatives or monoamino-carboxylic acids, indicating the immunospecificity to a common structure among AGE-structures. The subsequent study (Ref. 10) revealed of 6D12 is an N<sup>ε</sup>-carboxymethyllysine(CML)-protein adduct.

### 【References】

1. Horiuchi, S. et al.: Immunochemical approach to characterize advanced glycation end products of the Maillard reaction; Evidence for the presence of a common structure. *J. Biol. Chem.* 266: 7329, 1991. 2. Araki, N. et al.: Immunochemical evidence for the presence of advanced glycation end products in human lens proteins and its positive correlation with aging. *J. Biol. Chem.* 267: 10211, 1992. 3. Miyata, T. et al.:  $\beta_2$ -Microglobulin modified with advanced glycation end products is a major component of hemodialysis-associated amyloidosis. *J. Clin. Invest.* 92: 1243, 1993. 4. Yamada, K. et al.: Immunohistochemical study of human advanced glycosylation end-products (AGE) in chronic renal failure. *Clin. Nephrol.* 42: 354, 1994. 5. Kume, S. et al.: Immunohistochemical and ultrastructural detection of advanced glycation end products in atherosclerotic lesions of human aorta using a novel specific monoclonal antibody. *Am. J. Pathol.* 147 : 654, 1995. 6. Makino, H. et al.: Ultrastructure of nonenzymatically glycated mesangial matrix in diabetic nephropathy. *Kidney International* 48: 517, 1995. 7. Mori, T. et al.: Localization of advanced glycation end products of Maillard reaction in bovine tissues and their endocytosis by macrophage scavenger receptors. *Exp. Molec. Pathol.* 63:135, 1995. 8. Miyata, T. et al.: Identification of pentosidine as a native structure for advanced glycation end products in  $\beta_2$ -Microglobulin forming amyloid fibrils in patients with dialysis-related amyloidosis. *Proc. Natl. Acad. Sci. USA.* 93: 2353, 1996. 9. Kimura, T. et al.: Accumulation of advanced glycation end products of the Maillard reaction with age in human hippocampal neurons. *Neurosci. Lett.* 208: 53, 1996. 10. Ikeda, K. et al.: N<sup>ε</sup>-(carboxymethyl) lysine protein adduct is a major immunological epitope in proteins modified with advanced glycation end products of the Maillard reaction. *Biochemistry* 35: 8075, 1996. 11. Horiuchi, S. et al.: AGE modified proteins and their potential relevance to atherosclerosis. *Trends Cardiovasc. Med.* 6: 163, 1996. 12. Hammes, H-P et al.: Modification of vitronectin by advanced glycation alters functional properties in vitro and in the diabetic retina. *Lab. Invest.* 75: 325, 1996. 13. Kimura, T. et al.: Identification of advanced glycation end products of the Maillard reaction in Pick's disease. *Neurosci. Lett.* 219: 95, 1996. 14. Nakayama, M. et al.: immunohistochemical detection of advanced glycosylation end-products in the peritoneum and its possible pathophysiological role in CAPD. *Kidney International* 51: 182, 1997. 15. Mizutani, K. et al.: Photo-enhanced modification of human skin elastin in actinic elastosis by N<sup>ε</sup>-(carboxymethyl)lysine, one of the glycoxidation products of the Maillard reaction. *J. Invest. Dermatol.* 108: 797, 1997. 16. Murata, T. et al.: The relationship between expression of advanced glycation end products and vascular endothelial growth factor in human diabetic retinas. *Diabetologia* 40: 764, 1997. 17. Sugimoto, K. et al.: Localization in human diabetic peripheral nerve of N<sup>ε</sup>-carboxymethyllysine-protein adducts, one of advanced glycation endproducts. *Diabetologia* 40: 1380, 1997. 18. Shimokawa, I. et al.: Advanced glycosylation end-products in adrenal lipofuscin. *J. Gerontol.* 51A: B49, 1998. 19. Yoshida, S. et al.: Immunohistochemical study of human advanced glycation end-products and growth factors in cardiac tissues of patients on maintenance dialysis and with kidney transplantation. *Clin. Nephrol.* 49:273, 1998. 20. Matsuse, S. et al.: immunohistochemical localisation of advanced glycation end products in pulmonary fibrosis. *J. Clin. Pathol.* 51: 515, 1998

### Distributor



COSMO BIO Co., LTD.  
Inspiration for Life Science

TOYO EKIMAE BLDG. 2-20, TOYO 2CHOME  
KOTO-KU, TOKYO 135-0016, JAPAN  
TEL : +81-3-5632-9617  
FAX : +81-3-5632-9618  
URL : <http://www.cosmobio.co.jp/>  
e-mail : [export@cosmobio.co.jp](mailto:export@cosmobio.co.jp)

### Manufacturer

Trans Genic Inc.

7-1-6 Minatojimaminami-machi,  
Chuo-ku, Kobe, 650-0047 JAPAN  
TEL : +81-78-306-0590  
FAX : +81-78-306-0589  
URL : <http://www.transgenic.co.jp/>  
e-mail : [techstaff@transgenic.co.jp](mailto:techstaff@transgenic.co.jp)