Investigation of stress induced Tbx20 function in autophagy in heart with comparative analyses of extracellular matrix remodeling with multiple stress inductions in cultured cardiac cell line.

Thesis submitted for the partial fulfilment of the requirements

for the degree of

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By

Riffat Khanam

Under the supervision of
Dr. Santanu Chakraborty
Department of Life Sciences
Faculty of Natural and
Mathematical Sciences
Presidency University,
Kolkata, India
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Name of the candidate: Riffat Khanam

Registration Number: R-17RS18210101

Date of registration: 14.08.2018

Department: Department of Life Sciences

Signature of candidate with date

Riffet Theran

28.07.2023

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Declaration

I hereby declare that this thesis contains original research work carried out by

me under the guidance of **Dr. Santanu Chakraborty**, Assistant professor,

Department of Life Sciences, Presidency University, Kolkata, India as part

of the PhD programme.

All information in this document have been obtained and presented in

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I also declare that, this work has not been submitted for any degree either in

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Certificate

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List of published research papers



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Identification of Adamts4 as a novel adult cardiac injury biomarker with therapeutic implications in patients with cardiac injuries

Riffat Khanam, ¹ Arunima Sengupta, ² Dipankar Mukhopadhyay, ³ and Santanu Chakraborty. ¹⁰

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Associated Data

- Supplementary Materials
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Abstract Go to: >

Pathological cardiac remodeling as an aftermath of a severe cardiac injury can lead to ventricular dysfunction and subsequent heart failure. Adamts4, a metalloproteinase, and disintegrin with

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Asporin Reduces Adult Aortic Valve Interstitial Cell Mineralization Induced by Osteogenic Media and Wnt Signaling Manipulation *In Vitro*

Anisha Polley ¹, Riffat Khanam ¹, Arunima Sengupta ², Santanu Chakraborty ¹

Affiliations + expand

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Abstract

Worldwide, calcific aortic valve disease is one of the leading causes of morbidity and mortality among patients with cardiac abnormalities. Aortic valve mineralization and calcification are the key events of adult calcific aortic valve disease manifestation and functional insufficiency. Due to heavy mineralization and calcification, adult aortic valvular cusps show disorganized and dispersed stratification concomitant with deposition of calcific nodules with severely compromised adult valve function. Interestingly, shared gene regulatory pathways are identified between bone-forming cells and heart valve cells during development. *Asporin*, a small leucine-rich proteoglycan (43 kDa), acts to inhibit mineralization in periodontal ligament cells and is also detected in normal murine adult

Table of Abbreviations

CVD: Cardiovascular disease

CHD: Congenital heart disease

AV: Atrio-ventricular

SA: Sino-atrial

AVC: Atrioventricular canal

LV: Left ventricle

RV: Right ventricle

OFT: Outflow tract

E: Embryonic day

MI: Myocardial infarction

AMI: Acute myocardial infarction

IHD: Ischemic heart disease

IWMI: inferior wall MI

AWMI: Anterior wall MI

T2D: Type 2 diabetes

ECM: Extracellular matrix

MMP: Matrix metalloproteinase

TIMP: Tissue inhibitor of matrix metalloproteinase

I/R: Ischemia- Reperfusion

DCM: Dilated cardiomyopathy

DCM & D: DCM with T2D.

EMT: epithelial to mesenchymal transformation

Con: Control

Strv: Starvation

Rap: Rapamycin

CS: Carnegie stage

MESP: Mesoderm Posterior Protein 1

BMP: Bone morphogenetic protein

ANF: Atrial natriuretic factor

Cx40: Connexin40

Yap: Yes associated protein

MST1: mammalian STE20-like protein kinase 1

LATS1: Large tumor suppressor kinase 1

SAV1: Salvador homologue1

SHF: Secondary heart field

RA: Retinoic acid

SRF: Serum Response Factor

Mef2c: Myocyte enhancer factor 2c

MLC2v: Myosin light chain 2v

bHLH: Basic helix loop helix

ROS: Reactive oxygen species

Hyp: Hypoxia

LOF: Loss of function

Col: Collagen

HA: Hyaluronic acid

LOF: Loss of function

LAMP-2A: Lysosomal associated membrane protein 2A

DMEM: Dulbecco's modified eagle medium

FBS: Fetal bovine serum

PBS: Phosphate buffered saline

EDTA: Ethylenediaminetetraacetic acid

TAE: Tris-acetate-EDTA

GSK-3β: Glycogen synthase kinase-3β

mTOR: Mammalian target of Rapamycin

TGF- β : Transforming growth factor β

α-SMA: α Smooth muscle actin

Col-III: Collagen III

Sirt1: Sirtuin1

Glut1: Glucose transporter

WGA: Wheat germ agglutinin

SRF: Serum response factor

PBMC: Peripheral blood mononuclear cells

RT-PCR: Reverse transcriptase PCR

qPCR: Quantitative Real time PCR

WB: Western blotting

IF: immunofluorescence

IHC: Immunohistochemistry

CV450: Cytocalcein violet 450

AG: Apoxin green

7-AAD: 7-amino actinomycin D

Tbx20si: Tbx20 siRNA

ATSsi: Adamts4 siRNA

ALKI: ALK inhibitor

ELISA: Enzyme linked immunosorbent assay

PAGE: Polyacrylamide gel electrophoresis

SDS: Sodium dodecyl sulphate

PI-3K: Phosphoinisitide-3- kinase

AKT: Protein kinase B

DAPI: 4',6-diamidino-2-phenylindole

TBS: Tris-buffered saline

TBST: Tris-buffered saline with Tween 20

RT: Room temperature

Scr: scrambled siRNA

HHD: Hypertensive heart disease

Table of Figures

Chapter 1	
Figure 1: MMPs linked with Cardiovascular diseases (CVDs)	20
Chapter 2	
Figure 1: qPCR study of Autophagy induced reactivation of Tbx20	
in H9c2 cells and aged mice	34
Figure 2: Tbx20 upregulation in H9c2 cells following autophagy	
induction depicted by IF and WB	37
Figure 3: H9c2 proliferation assay by Ki67 staining	39
Figure 4: Detection of apoptosis rate under autophagy influence	41
Figure 5: Tbx20-Sirt1 interaction in-silico assay	44
Figure 6: Assessment of cell size in heart tissue samples of mice	
treated with Strv and Rap	45
Figure 7: In-vivo expression of TBx20 along with GSK-3β, Sirt1,	
Nkx2.5 and Gata4 after Strv and Rap treatment	47
Figure 8: Autophagy regulation through mTORC1 and upregulation	
of Tbx20 and Sirt1 in aged mice	50
Figure 9: High levels of GSK-3β and low levels of p-GSK-3β found	
in aged murine hearts following autophagy induction	53
Figure 10: <i>In-vitro</i> assessment of GSK-3β expression and Tbx20	
LOF assay	56
Figure 11: Tbx20 dependent expression of Nkx2.5 and Gata4	59
Figure 12: Tbx20 mediated regulation of Sirt1 and GSK-3β under	

autophagic conditions	62
Chapter 3	
Figure 1: H9c2 Cell morphology after ROS and hypoxia injury	79
Figure 2: Injury induced overexpression of Adamts4 and fibrosis	
related markers in H9c2 cells	81
Figure 3: Adamts4 and Tgf- β1 expression is inhibited following	
pre-treatment with ALKI	84
Figure 4. Inhibition of Col-III and α -SMA by ALKI pre-treatment	87
Figure 5. Periostin expression is reduced after ALKI	
pre-treatment and successful knockdown of Adamt4 mRNA in	
cultured H9c2 cells	90
Figure 6: Tgf- β1 expression remains unaffected by	
Adamts4 knockdown	93
Figure 7: Inhibited expression of Col-III and α -SMA following	
loss of function of Adamts4	96
Figure 8: Reduced expression of Periostin following Adamts4	
loss of function	98
Figure 9: Enhanced expression of Vimentin in adult primary	
cardiac fibroblasts following ROS and hypoxia injury treatment	100
Figure 10: Upregulated expression of Periostin and α -SMA in	
murine adult primary cardiac fibroblasts following ROS and	
hypoxia injury treatment	101
Figure 11: Overexpression of Adamts4 and Tgf-β1 in murine	

adult primary cardiomyocytes after ROS injury
Figure 12: Upregulation of Adamts4 and α -SMA proteins in
adult cardiac patients
Figure 13: Adamts4 expression following hyperglycaemic shock
in H9c2 cells
Figure 14: Measurement of FBG, in T2D induced rats along with
measurement of ADAMTS4 in DCM&D samples
Chapter 4
Figure 1: Hypothetical proposed model depicting the role of
Tbx20 under autophagic influence
Figure 2: Hypothetical Proposed model depicting hierarchy
and inter-relationship between Tgf- β1 and Adamts4

Table of Tables

Chapter 1.	
Table 1: T-box transcription factors in cardiac development	
Chapter 2.	
Table 1: Characteristics of clinical sample groups	. 74
Table 2: Cholesterol and Triglyceride levels of control (C1, C2 and C3)	
and High fat+ Alloxan treated rats (T1, T2 and T3)	112

Conference and seminar certificates







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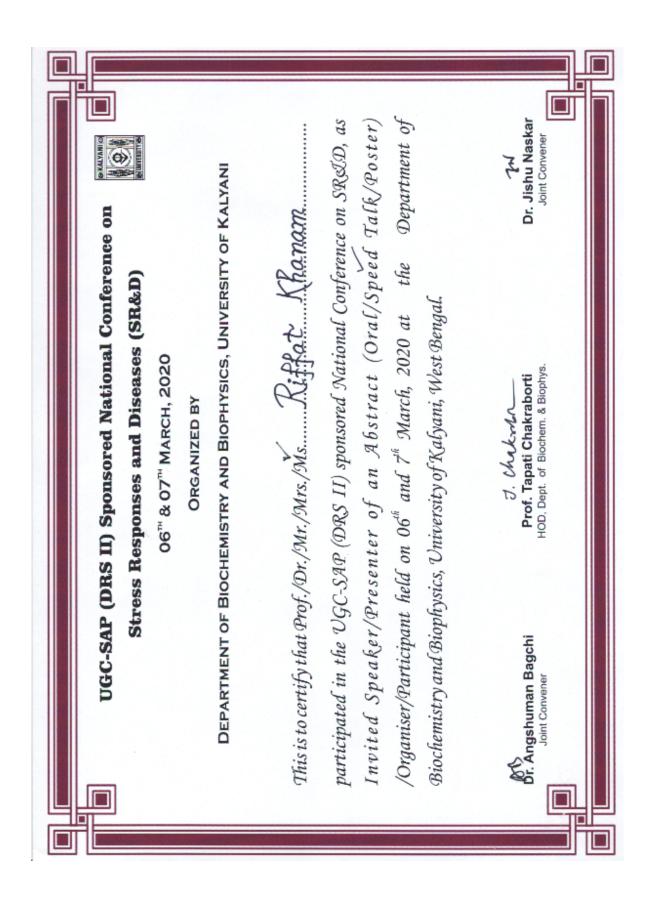
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190



