Mazin Al-Salihi MBBS PhD Assistant Professor - Faculty of Medicine Acting director of clinical research - Cell Therapy Center The University of Jordan

Nationality: Jordanian

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As a researcher with a clinical background I am passionate in teaching and researching problems immediately relevant to patient health. My postgraduate career started with the prestigious Fulbright pre-doctoral award to work at an NCI-designated cancer center. I had the honor of working with and learning from researchers originally involved in discovery of the *APC* gene in colon cancer.

I am currently interested in colitis associated colon cancer. With only 1 week of prophylactic treatment in an inflammatory bowel disease model colon tumor load is reduced by 50%. I have been successful in obtaining significant funding and publishing my work.

Qualifications:

PhD Oncological Sciences, University of Utah, Salt Lake City, May 2008 (achieved in less than 5 years) **MBBS medicine and surgery** with honors, University of Jordan, Amman, June 2001

Awards, Grants & Honors:

Scientific Research Fund Grant-MOHE 2016-2018 (JD 110,000 preliminary approval) Medical Research Council Career Development Fellowship 2010-2013 (\$200,000) Joan's Legacy Foundation Grant 2007-2008 (\$100,000) Fulbright Pre-doctoral Award - US Department of State, 2003-2005 (\$100,000) Dean of Medicine Award for Academic achievement, 2001 University of Jordan Academic Achievement Scholarship, 1995-2001

Research Experience:

2014-presentAssistant professor, Principal investigatorUniversity of Jordan, Faculty of Medicine & Cell Therapy Center

- Investigating the effect of mesenchymal stem cell endoscopic injections on inflammatory bowel disease pathogenesis & neutrophil myeloperoxidase.
- Characterizing the contribution of COX-2 signaling in "Never-Smokers" lung cancer in order to overcome resistance to tyrosine kinase therapy.

2010-2013 Career Development Fellow Medical Research Council - Protein Phosphorylation Unit University of Dundee

- Discovered the deubiquitylating enzymes USP11 and USP15 as novel regulators of TGF β and BMP pathway signaling respectively.
- Documented the effect USP11 has on TGFβ induced epithelial to mesechymal transition, a process thought to be essential for cancer invasion and metastasis.

- USP11 is now being studied by the pharmaceutical companies involved in the division of signal transduction therapy as a druggable target.
- Authored papers on USP11 and USP15 as well as a comprehensive review on deubiquitylation in regulating TGFβ signaling.
- Co-authored the role of the deubiquitylating enzyme OTUB1 in $\mathsf{TGF}\beta$ pathway signaling.
- Taught an MRes class on $\mathsf{TGF}\beta$ in cancer, as part of the cancer biology lecture series.
- Supervised one undergraduate student and co-supervised three graduate students

2008-2010 Postdoctoral Research Fellow

Huntsman Cancer Institute, University of Utah

- Identified and authored the mechanism of action of 5-azacytidine in solid tumors in relation to their p53 and replicative status. This was a preliminary study to Phase II clinical trials of the azacytidine compounds in late stage colon adenocarcinoma patients.
- Studied and authored a paper on neutrophil myeloperoxidase (MPOx) activity as a high impact new therapeutic target in colitis associated colon cancer. <u>1 week of prophylactic treatment reduced tumor load by 50% in an inflammatory colon cancer model. Confirmed genetically.</u>
- Supervised 2 undergraduate students and one lab technician. Co-supervised one graduate student.

2003-2008

Graduate Research Assistant (Predoctoral Fulbright Award)

Huntsman Cancer Institute, University of Utah

- Characterised and authored the mechanism of epidermal growth factor transactivation by prostaglandin E2. Excess PGE2 as a consequence of cyclooxgenase-2 (COX-2) over-activity or over-expression. Both common events in multiple cancers including colon cancer.
- Studied and authored the role of COX-2 in colon cancer initiation and progression in a non-inflammatory colon cancer mouse model based study.
- Devised a new technique for assaying COX-2 activity in colon crypts ex vivo.
- Assisted in exploring transcriptional regulation of stop codon truncating mutations in muscular dystrophy, a cell based study of bypassing stop codons during transcription.
- Assisted in creating a new Drosophila model for amyloidgenic diseases.
- Supervised one lab technician.

Technical Proficiency:

In vitro Biochemical

- Protein biochemistry/analysis
 IP/Western/ELISA/Spec./
 Filtration Chromatography
- DNA/RNA analysis PCR/rtPCR/HPLC/Reporter assays

In vivo Cell

- Cloning/mutagenesis
- 3D cell culture
- Live cell microscopy/IF
- In vivo Model Organisms
- IP/IC/TV/SC injections
- Mouse dissection & surgery
- Tissue processing/sectioning - H&E/IHC/IF
- Light & confocal microscopy
- Drosophila Brain dissection
- Embryo dissection/ISH

Teaching Experience:

2014-Date	Introduction & gastrointestinal pathology for 2 nd year medical students Blood, lymphoreticular, gastrointestinal, & genitourinary systems Virology for 2 nd & 3 rd year medical students Supervisor, Master students in Medical Analysis
2013	Supervisor, Final year intern from University of Han (1 student) Students in their final year apply and are placed at research institutes to gain practical lab experience.
2012	MRes TGF β in cancer lecture Part of the Cancer biology lecture series designed for newly matriculated masters students. College of life sciences, University of Dundee
2010-2013	Co-supervsisor, 3 PhD candidates in Molecular Biology Medical Research Council, University of Dundee
2008-2010	Supervisor, ACCESS Program for Women in Science and Mathematics (2 students) This program is designed to provide early research experience to young women undergraduates. College of Science, University of Utah
2008-2010	Co-Supervisor, successful PhD candidate in Oncological Sciences Huntsman Cancer Institute, University of Utah
2005	Teaching assistant, Human Physiology (pre-med) Department of Biology, University of Utah
2003	International teaching assistant certification Graduate school, University of Utah

Presentations:

Wrestling with USPs: The Oompa Loompas of deubiquitylation. Medical Research Council Christmas symposium, **invited speaker** (2012)

USP11 augments TGFβ signalling by deubiquitylating ALK5. ZOMES VII. International conference and annual meeting of the DFG-SPP1365, poster presentation (2012)

USP11 and 15: New regulators of the TGFβ pathway? Medical Research Council Christmas symposium, **invited speaker** (2010)

Effect of myeloperoxidase inhibitor resorcinol on colon tumorigenesis in APC^{min/+} mice. Proceedings of the American Association for Cancer Research Annual Meeting, poster presentation (2010)

EGFR and COX-2 in Colon Carcinogenesis, Breaking the Vicious Cycle. Oncological Sciences Departmental Retreat, University of Utah, **invited student speaker** (2006)

Cyclooxygenase-2 Transactivates the Epidermal Growth Factor Receptor through Specific Eprostanoid Receptors and Tumor Necrosis Factor- α Converting Enzyme. Molecular Biology, Biological Chemistry Department Retreat, University of Utah, poster presentation (2005)

Further Professional Development:

Writing proposals for funding. Medical Research Council. Nov 2011

The role of minorities in the political process. Fulbright Enrichment Seminar. Mar 2004

Bibliography:

 Al-Salihi MA, Reichert E, Fitzpatrick FA. (2015) Influence of myeloperoxidase on colon tumor occurrence in inflamed versus non-inflamed colons of APCMin/+ mice. Redox biology. 6:218-225.

1a. **Al-Salihi MA**, Reichert E, Fitzpatrick FA. (2010) Effect of myeloperoxidase inhibitor resorcinol on colon tumorigenesis in APC(min/+) mice. Proceedings of the American Association for Cancer Research Annual Meeting. Volume: 51 Page: 552

- 2. Herhaus L, **Al-Salihi MA**, Dingwell K, Cummins T, Wasmus L, Vogt J, Ewan R, Bruce D, Macartney T, Weidlich S, Smith JC, Sapkota GP. (2014) USP15 targets ALK3/BMPR1A for deubiquitylation to enhance bone morphogenetic protein signalling. Open Biol. 4: 140065.
- 3. Herhaus L, **Al-Salihi MA**, Macartney T, Weidlich S, Sapkota GP. (2013) OTUB1 augments TGFβ signalling by inhibiting SMAD3 ubiquitylation. Nature Commun. 4: 2519.
- 4. **Al-Salihi MA**, Herhaus L, Macartney T, Sapkota GP. (2012) USP11 augments TGFβ signalling by deubiquitylating ALK5. Open Biol. 2: 120063.

4a. *AI-Salihi MA*, Herhaus L, Macartney T, Sapkota GP. (2012) USP11 augments TGF8 signalling by deubiquitylating ALK5. ZOMES VII. International conference and annual meeting of the DFG-SPP1365. Volume: 7 Page:1

- 5. **Al-Salihi MA**, Herhaus L, Sapkota GP. (2012) Regulation of the TGFβ Pathway by Reversible Ubiquitylation. Open Biol. 2: 120082.
- 6. **Al-Salihi MA**, Yu M, Burnett DM, Alexander A, Fitzpatrick FA. (2011) The depletion of DNA methyltransferase-1 and the epigenetic effects of 5-aza-2'deoxycytidine (decitabine) are differentially regulated by cell cycle progression. Epigenetics. 6: 1021-28.
- Al-Salihi MA, Pearman T, Reichert E, Rosenberg D, Prescott SM, Stafforini DM, Topham MK. (2009) Transgenic expression of Cyclooxygenase-2 in mouse intestine epithelium is insufficient to initiate tumorigenesis but promotes tumor progression. Cancer Lett. 273: 225-32.
- Al-Salihi MA, Ulmer SC, Doan T, Nelson CD, Crotty T, Prescott SM, Stafforini DM, Topham MK. (2007) Cyclooxygenase-2 transactivates the epidermal growth factor receptor through specific E-prostanoid receptors and tumor necrosis factor-alpha converting enzyme. Cell Signal. 19: 1956-63.

Professional Memberships:

2009-present	Affiliate member, American Association for Cancer Research		
2001-present	Registe	ered Physician, Jordan Medical Association	
2006-2007	Member, Retention, Promotion, and Tenure student committee, Department of Oncological Sciences, University of Utah		
2004-2007	Sponse	pred member, American Association for the Advancement of Science	
Clinical Experience:			
Feb. 2003-May 2003		John Radcliffe Hospital, Oxford Radcliffe Hospital NHS Trust Clinical attachment with junior house officer duties in GI surgery.	
Jul. 2001-Jul.2002		Jordan University Hospital, University of Jordan Clinical Internship; 12-month rotation in the following departments:	
		Radiology & Interventional Radiology Accident & Emergency (Regional Trauma and Burn Center) Obstetrics and Gynaecology Paediatrics Medicine Surgery	
Jul. 2000-Sept. 2000		Aberdeen Royal Infirmary, Grampian NHS Clinical Attachment in Surgery and Paediatrics as part of final year training.	

Medical Licensing Examinations:

Professional and Linguistic Assessments Board examination part 2 pass (pass/fail only), 2003

Professional and Linguistic Assessments Board examination part 1 pass 163/200 (Mark needed to pass 120, Average 125), 2002