Development of Deep Learning-integrated Futuristic Biomedical Platforms for Translational Digital Healthcare

unist

Doctoral Candidate

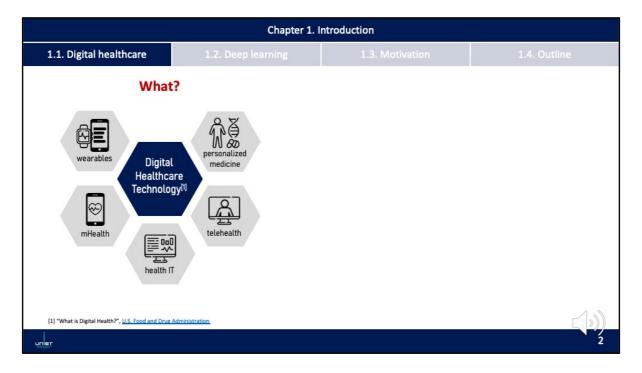
Sanzhar Askaruly

(February 24, 2022)

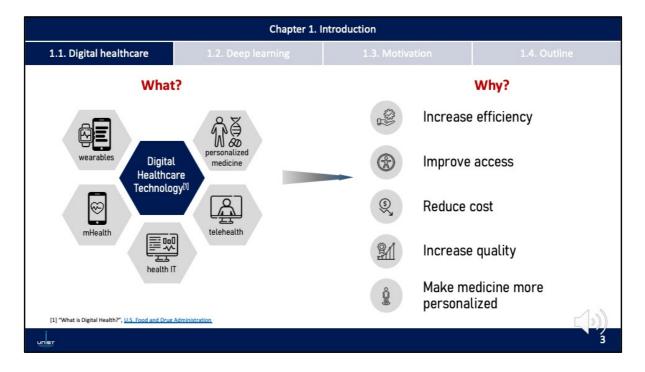
Supervised by Professor Woonggyu Jung Department of Biomedical Engineering College of Information-Bio Convergence Engineering Ulsan National Institute of Science and Technology (UNIST)

Dear committee, good day, this is a post-recorded PhD defense presentation of mine. It is done with the purpose to inform the committee members who could not participate, as well as our lab members.

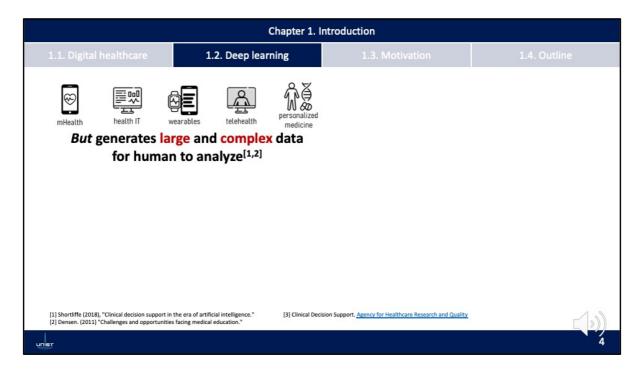
Dear all, I am Sanzhar Askaruly, a PhD candidate in Biomedical Engineering at UNIST. The topic of my Ph.D thesis is called Development of Deep Learningintegrated Futuristic Biomedical Platforms for Translational Digital Healthcare. My research has been done under supervision by Professor Woonggyu Jung.



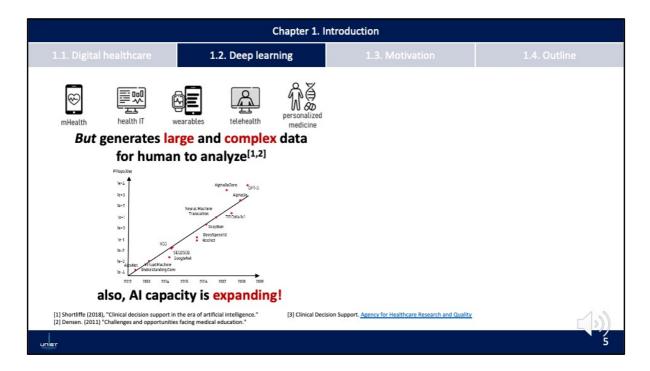
To begin with, I would like to introduce what is digital health. The broad scope of digital health includes categories such as wearable devices, mobile health, health IT, telehealth and personalized medicine. These digital tools give a more holistic view of human health through access to data and control.



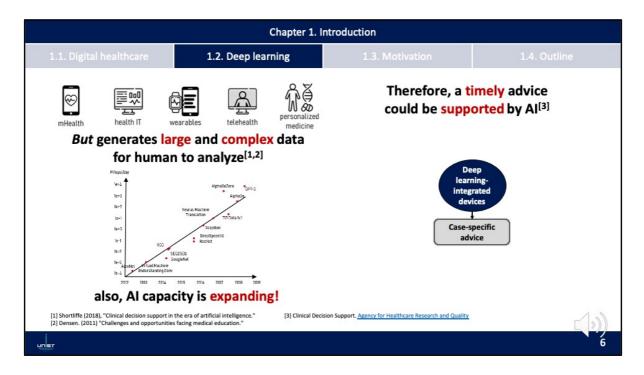
It offers real opportunities to improve medical outcomes and enhance efficiency. Providers and users of this tech have several benefits, including cost reduction, quality improvement, personalization and others.



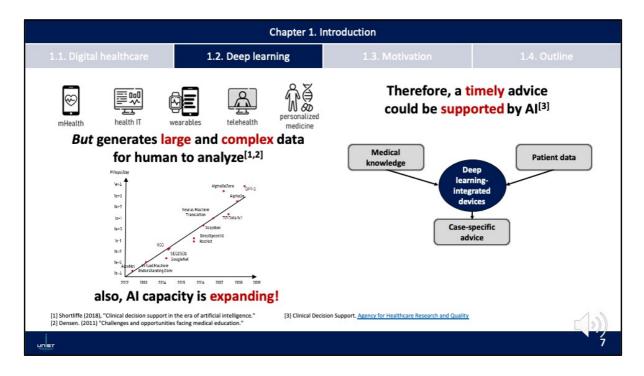
However, the extensive use of technologies within digital health creates a bottleneck of analysing large and complex amounts of medical data, where human is incapable.



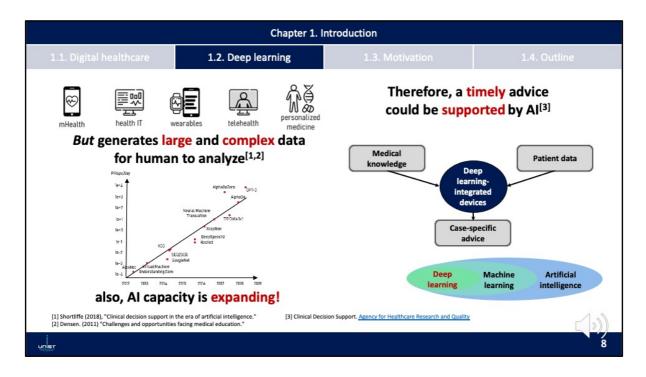
At the same time, the capacity of artificial intelligence keeps expanding.



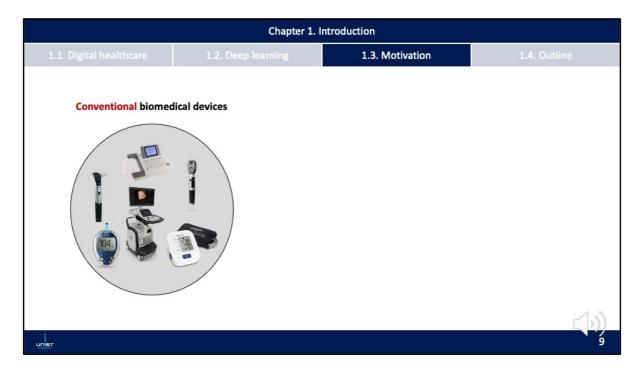
One of the opportunities for this challenge could be the suggestion of assistive intelligent devices. Their goal is to support the decision by giving a timely advice at the point of care.



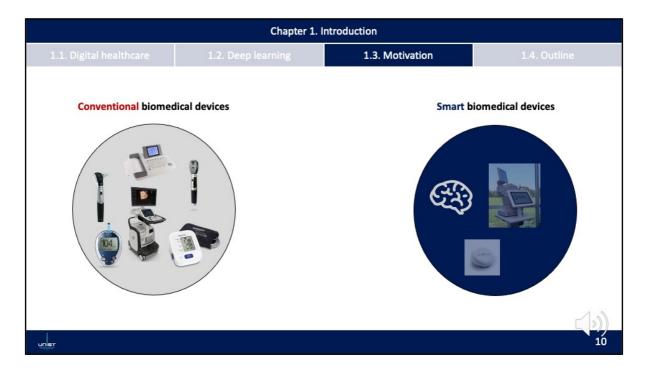
For this reason, they may need to utilize medical knowledge and patient data.



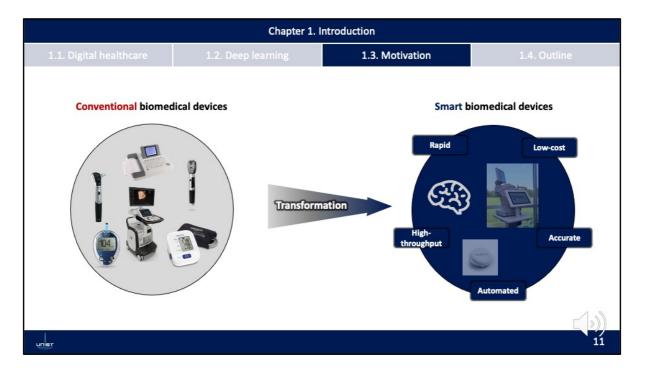
Deep learning is a computational technique, which is subset of a broader term of artificial intelligence. It was repeatedly reported to achieve success for various biomedical problems.



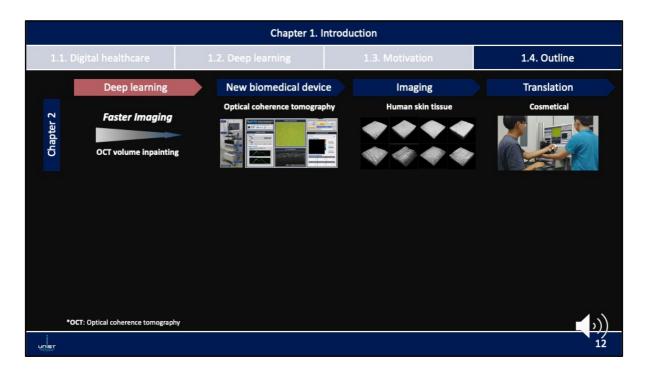
Currently, conventional biomedical devices are developed with fixed requirements. They are often primitive, static, and are even analogue sometimes in their nature.



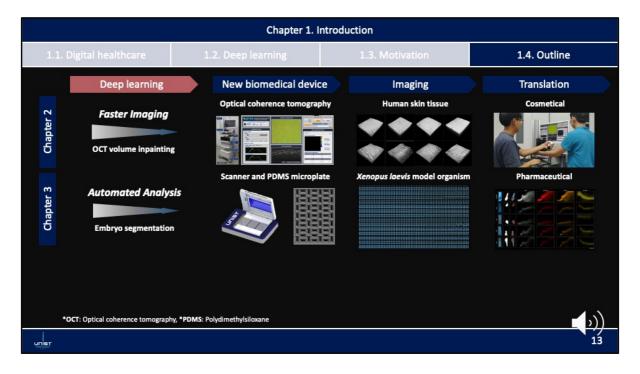
In contrast, smart biomedical devices possess built-in intelligence to support human decision. They exist, but rare, partly because they require multiple considerations. Also, their objective is not always obvious.



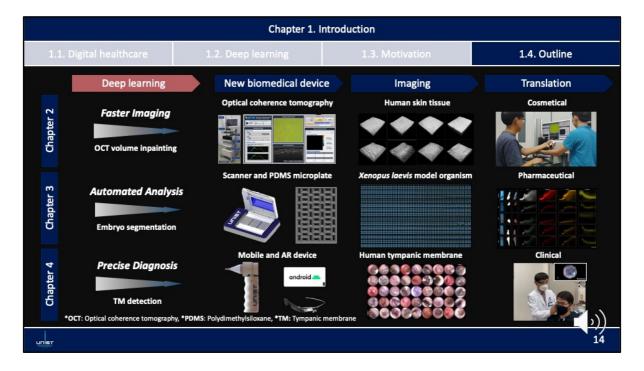
The transformation from one category to another is an open question. However, the exploration of opportunities within was my primary interest during the PhD program.



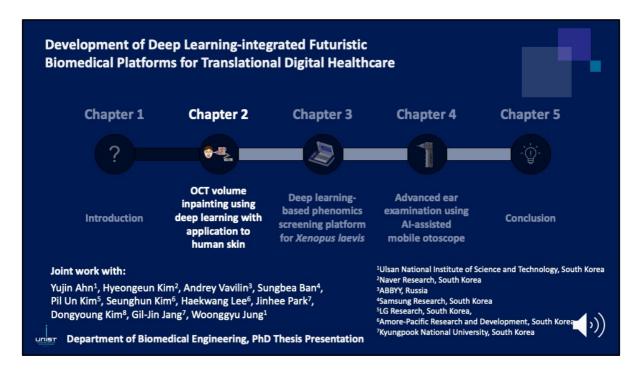
In this slide, I present with the outline of research works. Initially, I will describe how deep learning could be integrated to accelerate OCT imaging of human skin for cosmetical fields.



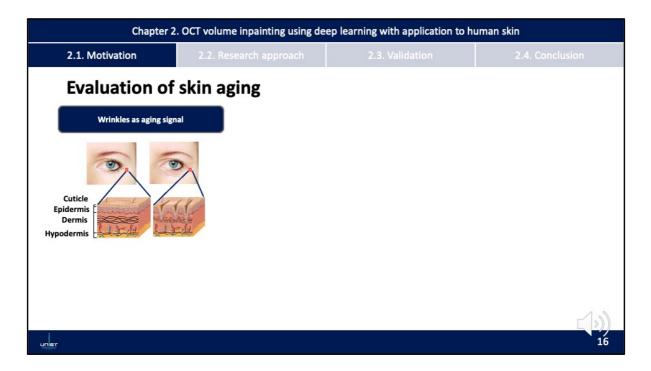
Next, I demonstrate the results of leveraging deep learning for automated analysis of an aquatic animal model, Xenopus, having implications for pharmaceutical research.



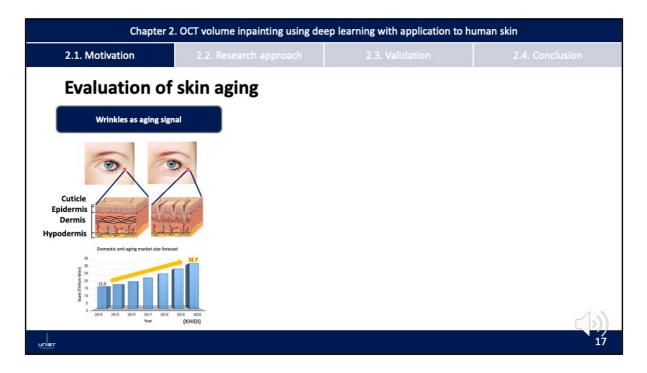
Finally, in chapter 4, I will present the outcome of incorporating AI model to enhance ear examination procedure to support clinical diagnosis.



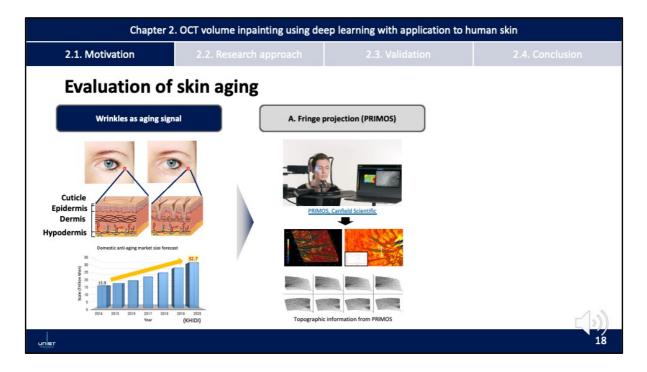
Let me introduce the first project OCT volume inpainting using deep learning with application to human skin. This is a joint work with Professor Jang's laboratory from Kyungpook National University and several researchers.



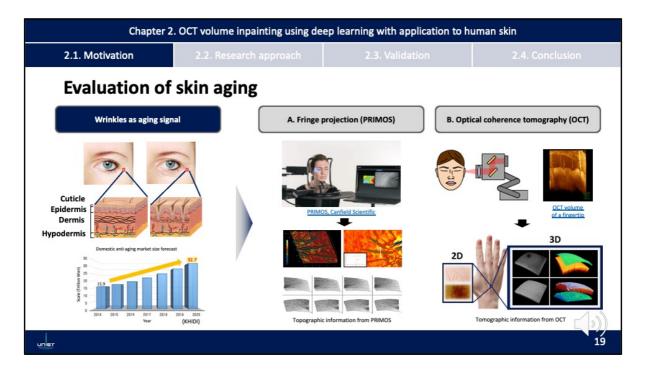
Wrinkle formation serves a representative signal of ageing in human skin.



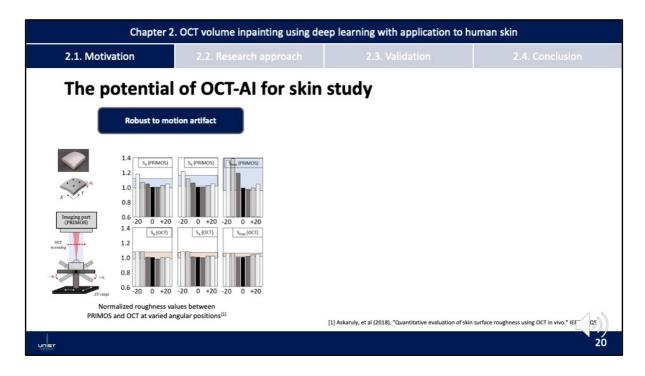
The anti-aging market is expanding, and the companies are driven to develop not only high-quality products, but also utilize high standard product assessment instruments. In particular, the *in vivo* and quantitative observation of skin is required for an accurate evaluation of the effectiveness of the product.



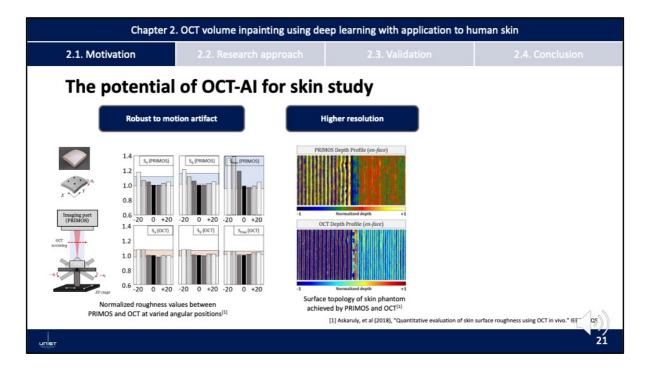
One of the most commonly used imaging systems for skin measurement, PRIMOS, was proposed as an objective tool in the cosmetics industry for studying skin topography and the volume of wrinkles. PRIMOS is a non-invasive, fast and direct measurement of the skin surface with high precision.



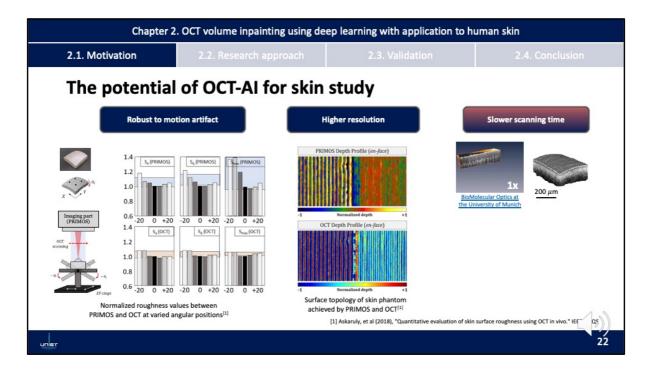
Another potential, but not so popular tool is OCT, which is a non-invasive, volumetric, real-time imaging modality providing a powerful tissue inspection. Detailed structural information is provided by OCT with high-resolution capability.



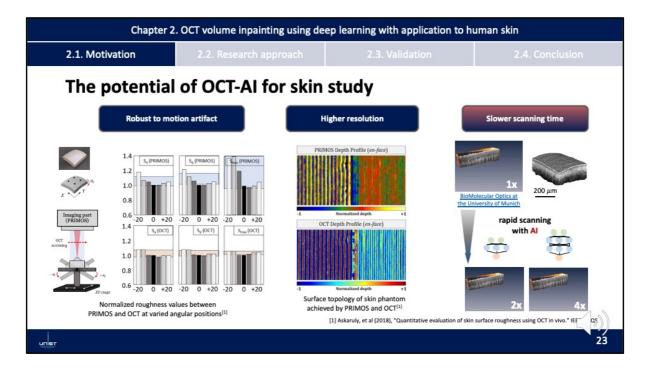
From our previously reported comparison study, it was identified PRIMOS has difficulty in providing accurate and reliable skin analysis because its results can vary according to orientation, motion artifacts, as well as back scattering of the subject.



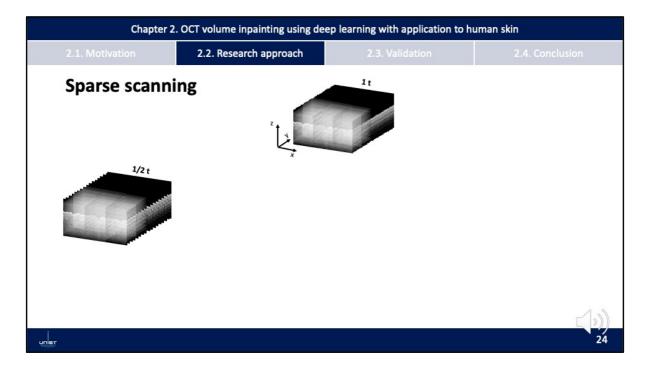
Moreover, in order to observe the compact and periodic structure of wrinkle, imaging device requires high resolution. Topologic image from OCT has well defined periodic structures and clear distinction of patterns compared to PRIMOS.



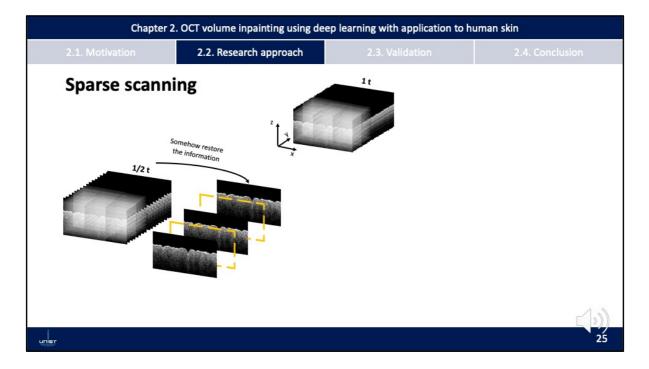
OCT inherently maintains essential advantages of in vivo skin imaging. However, OCT setups are often available in limited research environments. One obstacle is lengthy scanning procedure of the subject.



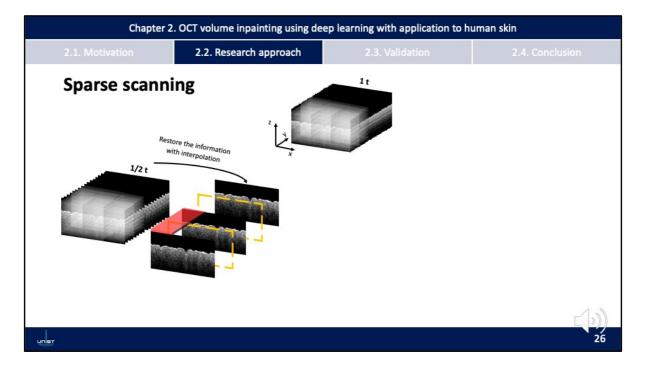
Essentially, we can overcome this issue with the help of deep learning technique to reduce the time of imaging.



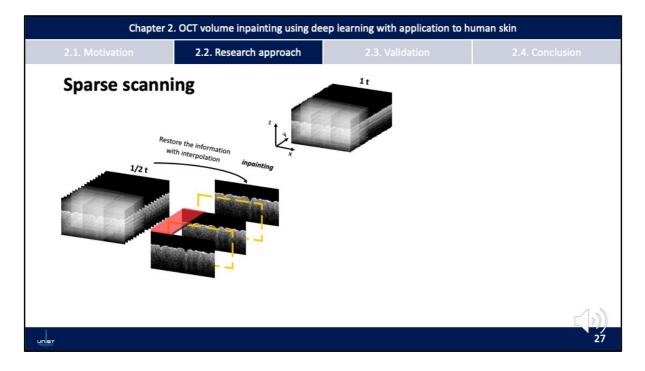
In the principle of the suggested technique, we start by low-sample scanning to reduce time, in this case half.



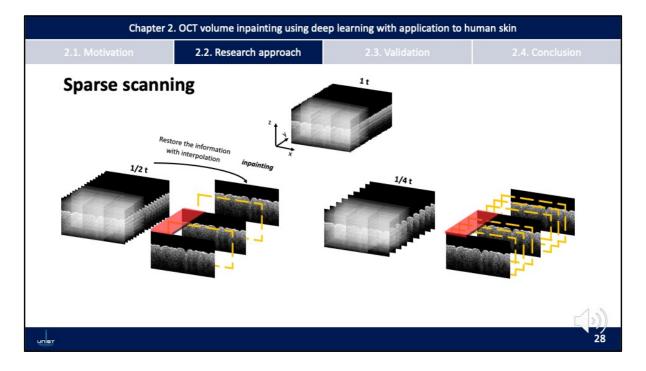
Further we somehow should restore the slices in between.



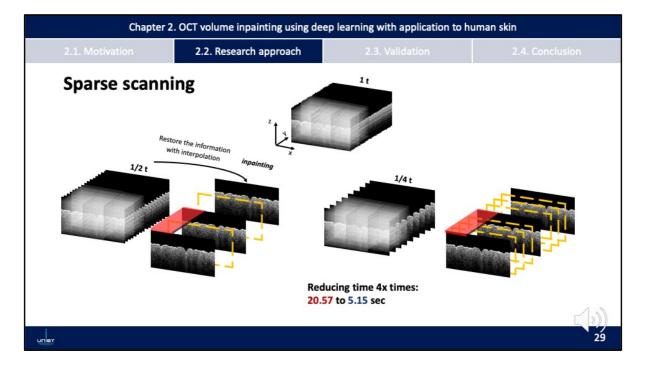
To do this, one straightforward way could be interpolation of so-called patches between two successive slices at each height and width. Basically, we can save halftime of full volume scanning at the cost of post-processing.



The common terminology for the information restoration represented here is called inpainting, and we will further refer to it this way.



Similarly, we can go further and scan only every fourth slice to save even more.



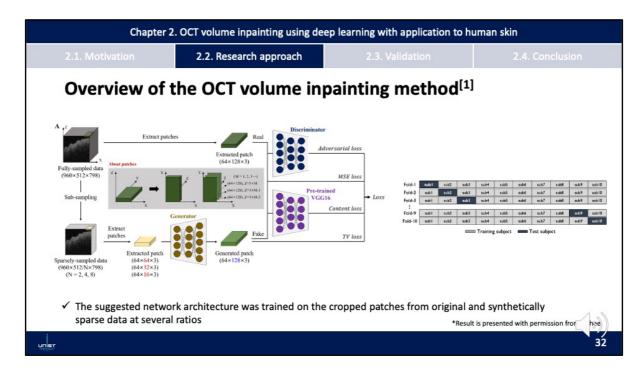
In this case, the average scan time of typical 1cm2 FOV would reduce from 20 seconds to 5 seconds.

Chapter 2. OCT volume inpainting using deep learning with application to human skin						
2.1. Motivation	2.2. Research approach	2.3. Validation	2.4. Conclusion			
Volume inpainting using GAN						
Ground truth 1//	2 Sparse 1/4 Sparse 1/8 Sparse					
Magnited						
CONTRACTOR OF CONT		Criginal image Generated image by interpolation Generated image by GAN (general	tive adversarial network) model			
		*Result	is presented with permission from Ahi			
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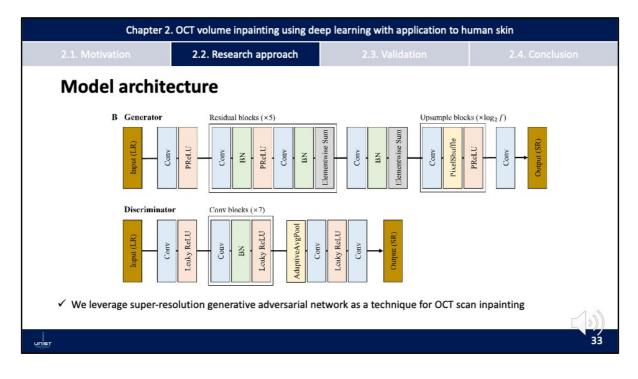
As we can expect, the described interpolation method could have adverse effect onto the accurate representation of structure, causing interrupted details, discontinuous surfaces, and other phenomena, reducing research and diagnostic significance. In this study, we present another, more precise approach based on deep learning technique.

Chapter 2. OCT volume inpainting using deep learning with application to human skin					
2.1. Motivation	2.2. Research approach				
Dataset					
Summary of skin dataset details 정응규 (2018), 근적외선 3차원 광학영상기법을 이용한 피부 정당화 기술개별					
			Participant		
Young (20s) Sample	OCT volumes Old (50s)	Number of subjects	10		
		Patients' age (years)	35.3 ± 8.5 40		
	(Stars-	Gender (% of female) S _a	40 2.23±0.57		
		Sa Sa	2.84±0.71		
		S _{max}	14.36±2.66		
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The collection of data was approved by the IRB protocol. A total of 10 people participated, average of 35 years and skin measurement using OCT conducted. Images on the bottom left visualize sample volumes of human skin.



Briefly speaking, we tried to restore the original dimension of the cropped patches of the intentionally dropped slices along C-mode scan axis. This patch is further utilized as input to the deep neural network. As for the model, we utilize generative adversarial network architecture. During training, we carried out 10-fold crossvalidation to evaluate the performance. In each fold, a model is trained using 9subjects data, and then tested using the other subject data not used for training.



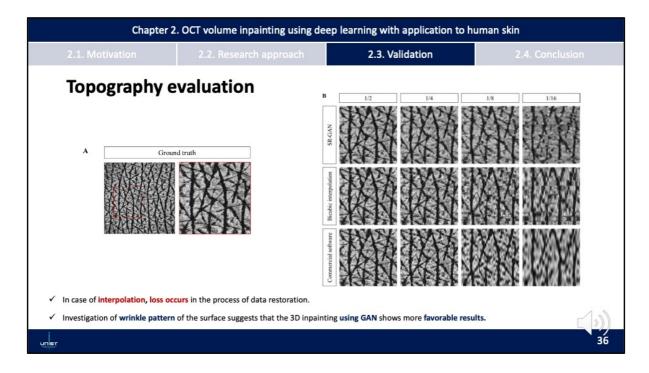
For deep neural network, we utilize SRGAN-like architecture. Its original purpose is the estimation of high-resolution images from its low-resolution images. Likewise, we can utilize the SRGAN for inpainting. Using the technique, we can obtain fully sampled OCT volume from sparsely sampled OCT scans.

Chapter 2. OCT volume inpainting using deep learning with application to human skin							
2.1. Motivation	2.2. Research approach	2.3. Validation	2.4. Conclusion				
Flattening algorithm							
Input	Input image Curvature estimation Flat						
2004	Flattening Surface detection Gaussian filter → Differentia Gaussian filter → Differentia	filter → Differential filter → Extract curvature Final result	300/m				
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One of the useful algorithms to know before the evaluation of surface is flattening. Its goal is to minimize the natural curvature of skin in order to perform correct measurements. Home-built combination of image processing steps produces the output as shown on the far right.

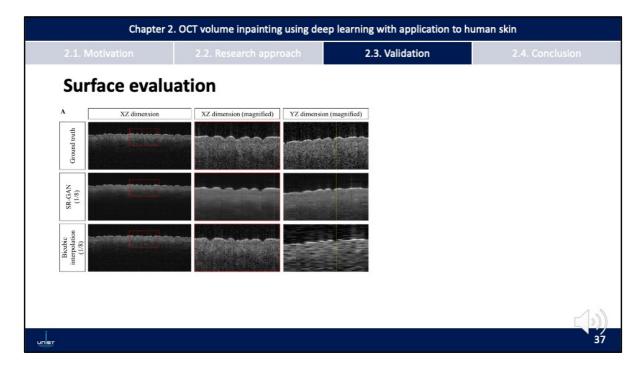
Chapter 2. OCT volume inpainting using deep learning with application to human skin						
2.1. Motivation	2.2. Research approach	2.3. Validation				
Evaluation metrics						
A. Su	rface roughness	B. Image quality				
Average surface roughness:		The mean squared error:	The mean squared error:			
$S_a = \frac{1}{nx \times n}$	$\frac{1}{ny}\sum_{i=1}^{nx}\sum_{j=1}^{ny} z(x_i, y_j) $	$MSE(x, y) = \frac{1}{N} \sum_{i=1}^{N} (x_i - y_i)^2$				
Root mean square roughness:		Peak signal-to-noise ratio:				
$S_q = \sqrt{\frac{1}{nx \times ny} \sum_{i=1}^{nx} \sum_{j=1}^{ny} z(x_i, y_j)}$		$PSNR = 20 \log_{10} \frac{(MAX_I)^2}{\sqrt{MSE}}$				
Maximal roughness: $S_{max} = \max(z) - \min(z)$		Multiscale structure similarity				
		$SSIM_{MS} = [l_M(x,y)]^{\alpha_M} \cdot \prod_{j=1}^{M} [c_j(x_j)]^{\alpha_M}$	$[x,y)]^{\beta_j} [s_j(x,y)]^{\gamma_j}$			
[1] Wang, et al (2003) "Multiscale structural similarity for image quality assessment" ACSSC						
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We evaluated performance based on surface roughness and image quality. Roughness characterizes irregularities on surfaces, providing information on the geometry of structure. Additionally, image quality was assessed using using MSE, multi-scale structural similarity index, and peak signal-to-noise ratio.

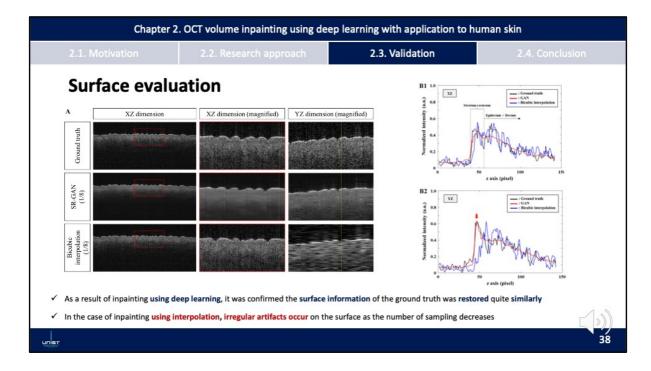


One of the common applications of OCT imaging in skin is the observation of skin topography. It is useful for estimation of wrinkle width, wrinkle depth and overall geometric orientations.

Here, we compared bicubic interpolation, commercial software and the suggested method at various sampling rates. Although better at half sampling rate, during interpolation, loss occurs in the process of data restoration for higher sampling. As a result of qualitative investigation of wrinkle pattern of the surface, we confirm that the 3D inpainting using GAN shows more favorable results.



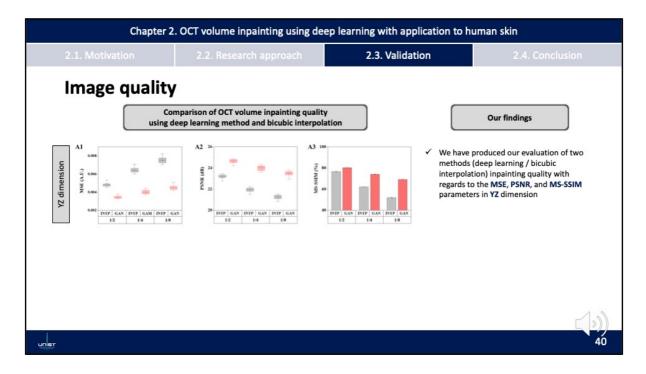
We investigated inpainting quality in cross-sectional dimensions. We can observe that interpolation could amplify artifact in the dimension, where it has less information. Although pixel information is blurred, the surface information is still restored with GAN.



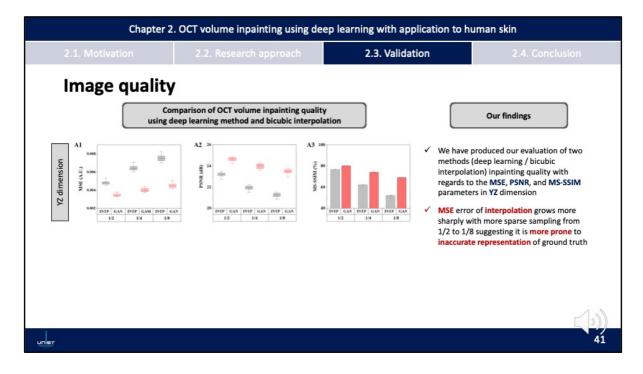
If we plot the depth profile along the axis, we can observe that bicubic interpolation tends to have more deviation from the ground truth compared to the deep learning method.

	2.2. Resear	ch approach	2.3. Validation	
Survey				
		Mean opinion score (MOS) test by 14 subjects		
	Sampling	SR-GAN	Bicubic interpolation	
	1/2	4.25	3.96	
	1/4	3.68	3.50	
	1/8	3.18	2.36	
		Preference test by 14 subjects		
	Sampling	SR-GAN	Bicubic interpolation	
	1/2	8 (57.1%)	6 (42.9%)	
	1/4	12 (85.7%)	2 (14.3%)	
	1/8	13 (92.9%)	1 (7.1%)	
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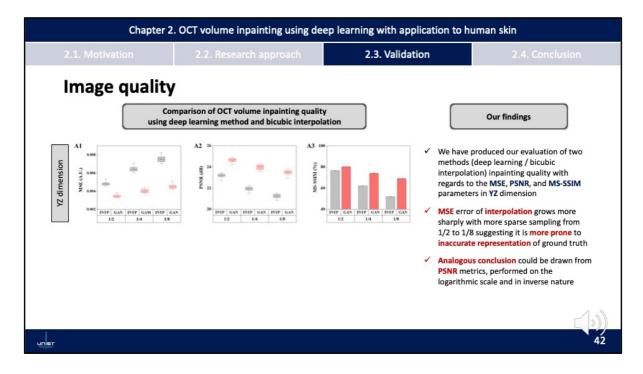
We evaluated the inpainting methods in terms of human opinion. To conduct survey, we provided the respondents the resultant images from interpolation and the deep learning approach. Table above reports mean opinion scores and table below describes the results of preference test. In both surveys, the deep learningbased method suggests prevalence over interpolation.



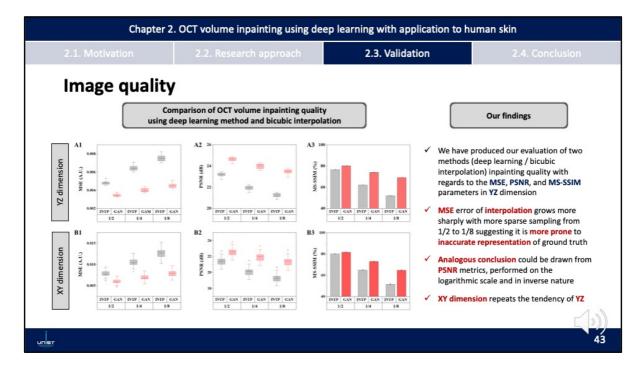
We investigated inpainting quality in cross-sectional dimension. In accordance with definitions, we compared MSE, PSNR and MS-SSIM parameters for bicubic interpolation and generative adversarial network.



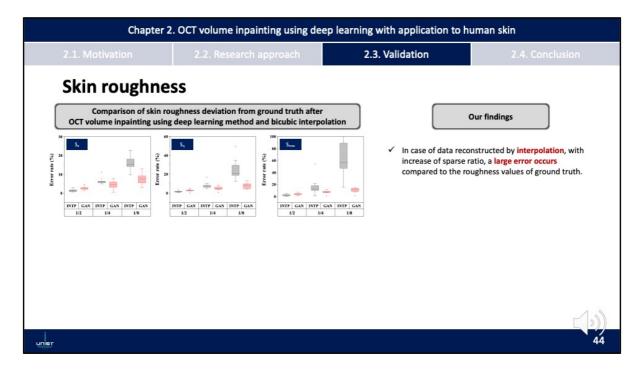
Here, MSE error of interpolation grows more sharply compared to the GAN with more sparse sampling from half to one-eighth suggesting it is more prone to inaccurate representation of ground truth



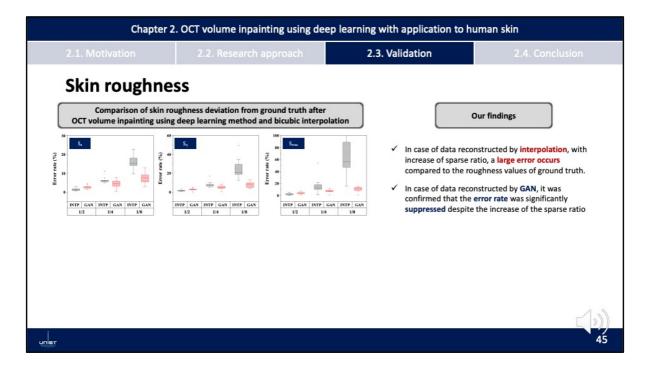
Analogous conclusion could be drawn from PSNR metrics, performed on the logarithmic scale and in inverse nature



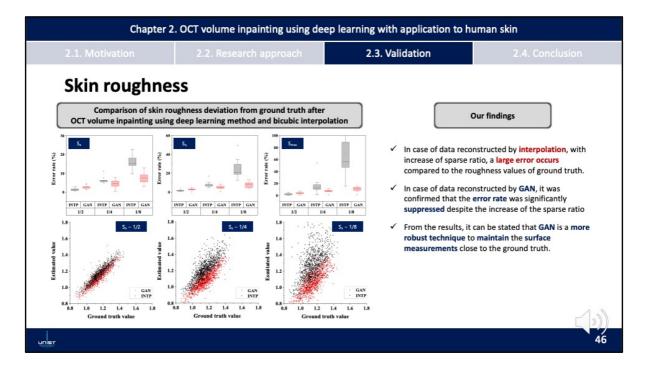
As for the top-view XY dimension, MSE and PSNR have analogous tendency and similar deviation ranges. SSIM measurements drop less abruptly for GAN.



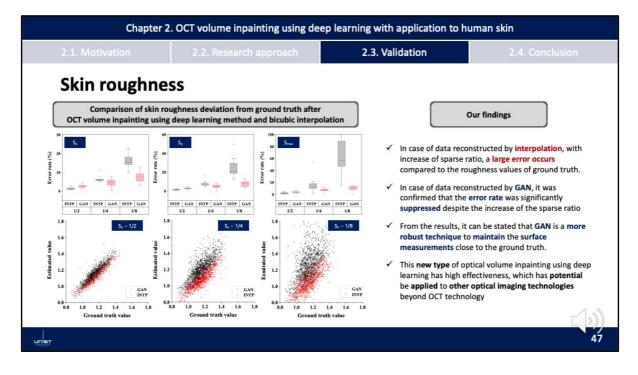
Another valuable perspective to inspect inpainting quality is the examination of roughness data with the increase of sampling ratio. In particular, here we present error rates for the measured roughness parameters. A large error occurs compared to the roughness values for interpolation cases.



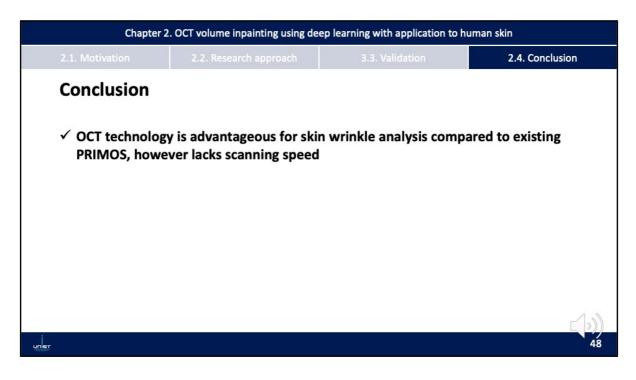
On contrast, in case of data reconstructed by GAN, it was confirmed that the error rate was significantly suppressed despite the increase of the sparse ratio.



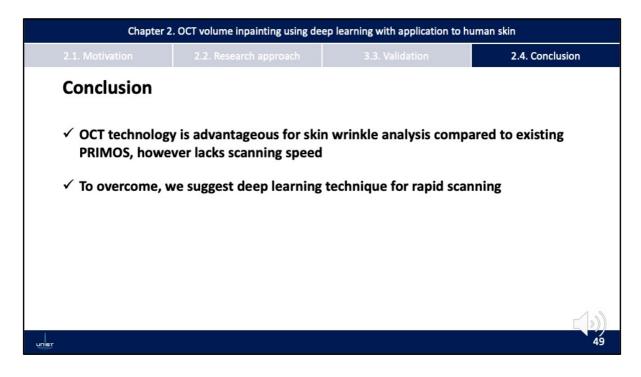
From the results, it can be stated that GAN is a more robust technique to maintain the surface measurements close to the ground truth.



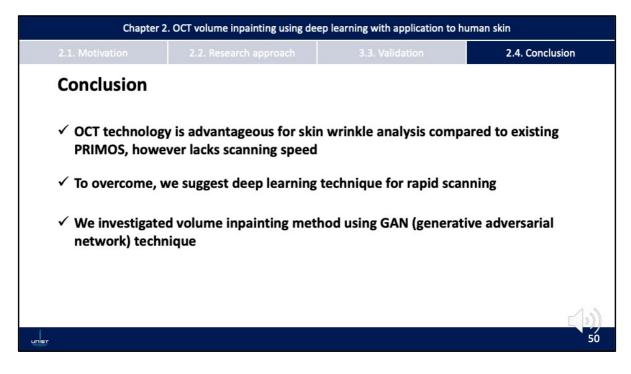
This new type of optical volume inpainting using deep learning has high effectiveness, which has potential be applied to other optical imaging technologies beyond OCT technology.



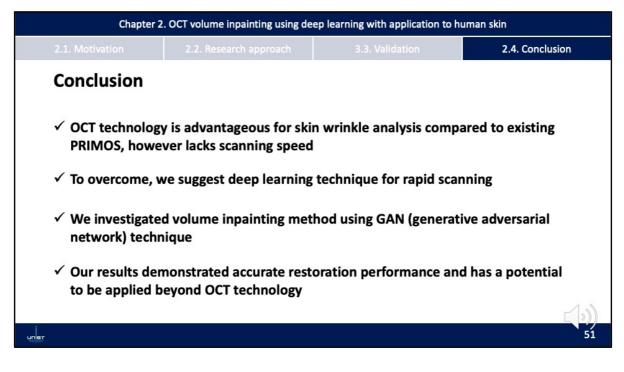
OCT technology is advantageous for skin wrinkle analysis compared to existing PRIMOS, however lacks scanning speed



To overcome this issue, we suggest deep learning technique for rapid scanning.



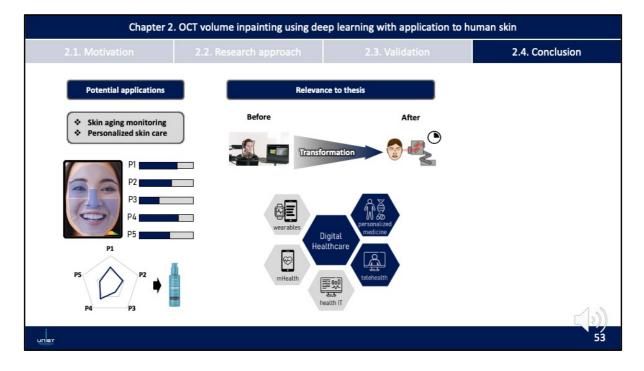
We investigated volume inpainting method using GAN (generative adversarial network) technique.



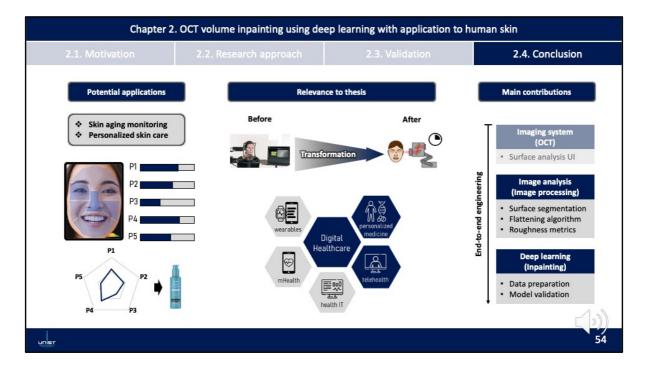
Our results demonstrated accurate restoration performance and has a potential to be applied beyond OCT technology.

Chapter 2	Chapter 2. OCT volume inpainting using deep learning with application to human skin					
2.1. Motivation		2.3. Validation	2.4. Conclusion			
Potential applications						
 Skin aging monitoring Personalized skin care 						
P1 P2 P3 P4 P5						
P1 P5 P4 P3 P2						
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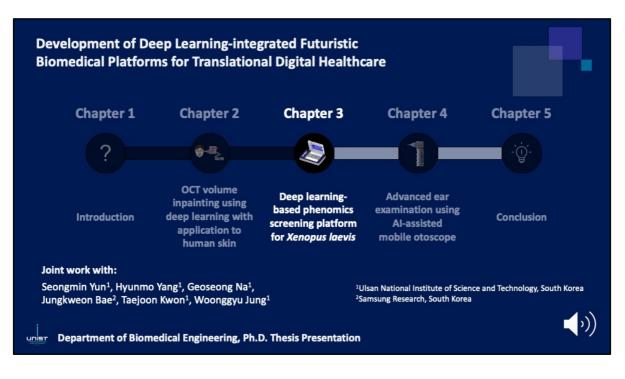
This project could potentially be useful for skin aging monitoring or personalized skin care.



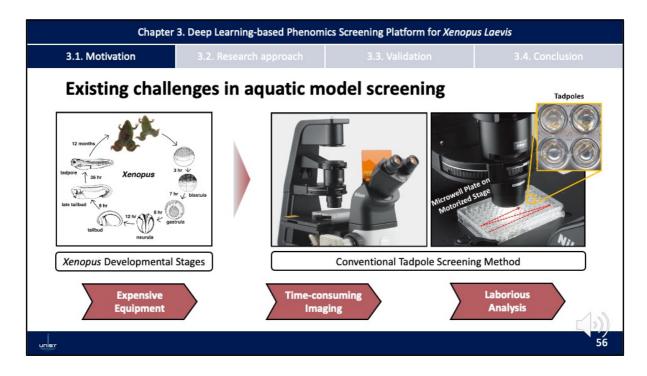
Within bounds of my thesis, I have explored the transformation of traditional skin assessment towards novel deep learning-integrated smart device. It could in perspective contribute towards the telehealth, or personalized medicine directions of digital healthcare technology.



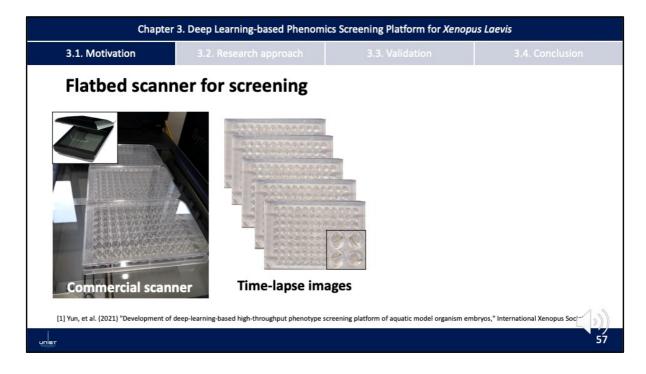
Although my experience with hardware was limited in this project and supported by my senior, here I outlined my personal contributions inside the research work towards end-to-end engineering.



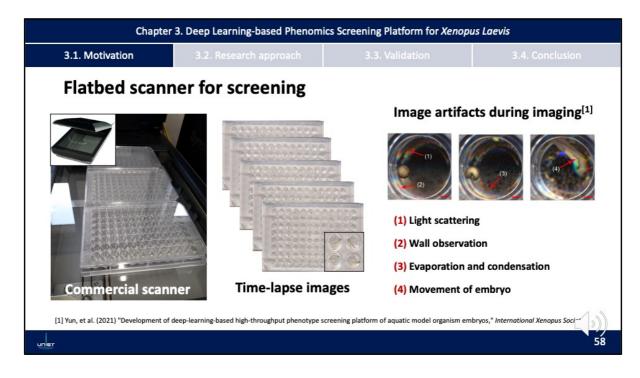
In the third chapter, I will present about the development of deep learning-based phenomics screening platform for xenopus laevis. This is a joint work with Professor Kwon's laboratory from UNIST and several researchers.



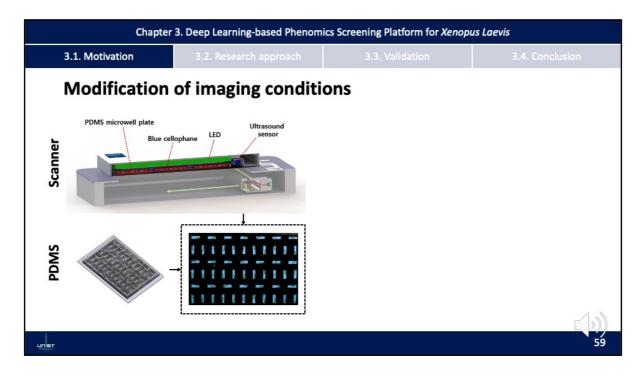
Xenopus laevis is emerging model to study human disease and to investigate pharmaceutical effects *in vivo* due to smaller size and faster developmental rates. It is also an effective organism to observe drug effects on phenotypic characteristics because it can provide many biological systems in a short time and remain optically accessible at the early stages of development. However, traditional screening of massive *Xenopus* data requires expensive equipment and labor-intensive inspection under an optical microscope.



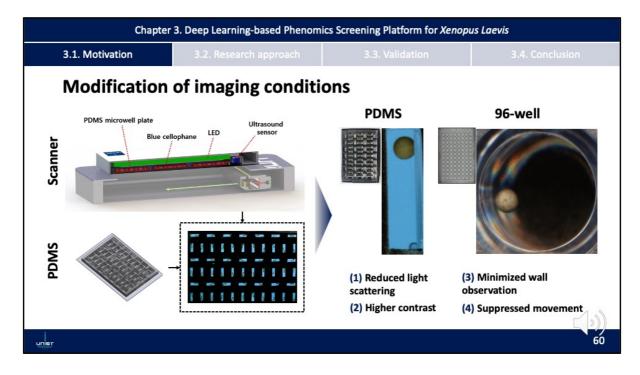
Alternatively, the flatbed scanner was considered to obtain quantitative images for large-scale phenotype assay. Its large field-of-view and low cost are advantageous. We could observe the phenotypic characteristics with a reasonable resolution as time-lapse images.



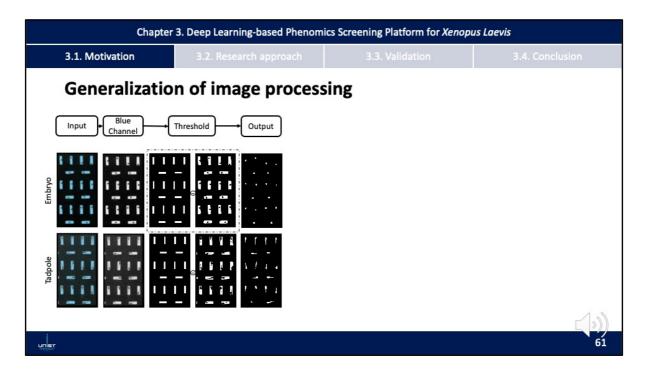
However, it was found that there were issues with the downstream image analysis, such as light scattering, wall observation and other artifacts.



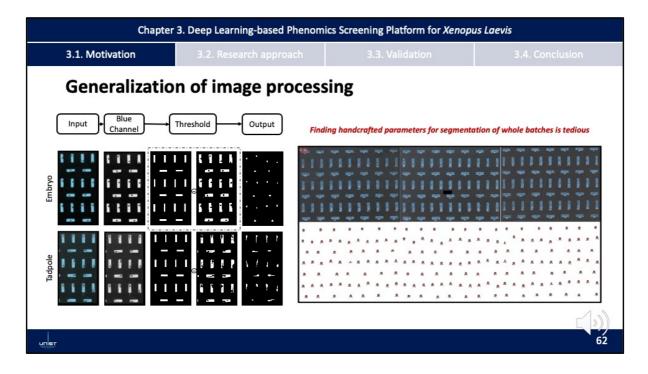
To overcome these issues, we have suggested modifications to the commercial scanner, converting it into a custom imaging device. We also fabricated the customized PDMS plate for efficient and stress-free imaging of living Xenopus samples in normal and drug environments.



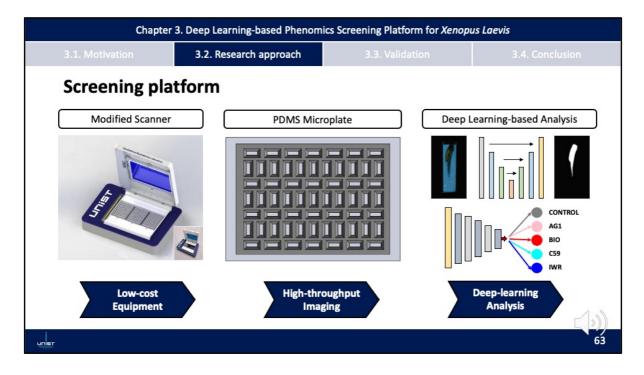
Our goal is to achieve reduced light scattering, higher contrast, and minimize wall observation.



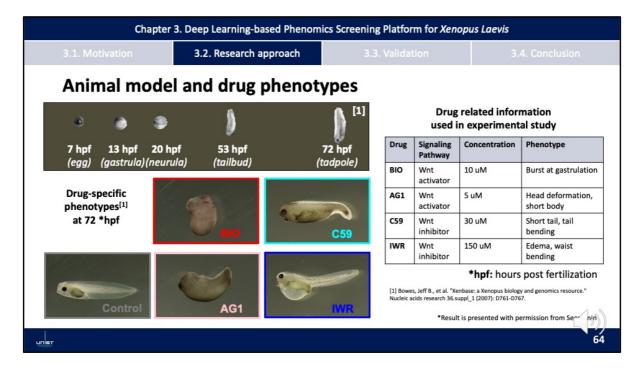
Our preliminary approach of analyzing embryos consisted of image processing pipeline for identification of the regions of interest with a few simple steps.



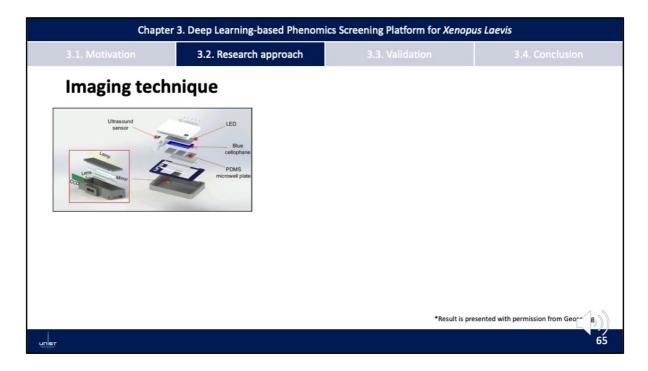
But as we expanded the technique to large-scale, we had to look for handcrafted parameters, which was necessary for better segmentation. We realized this computational approach quickly became manual task.



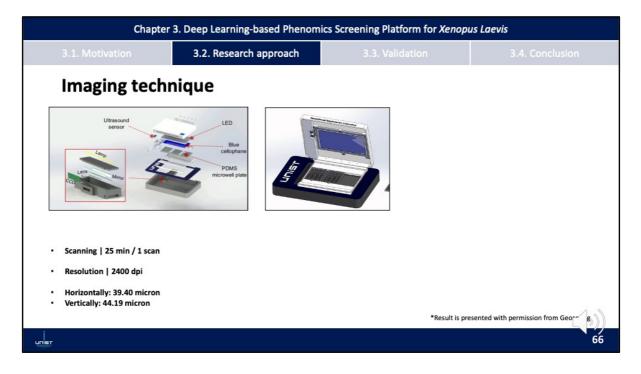
Therefore, in this work we suggest a screening platform consisting of modified flatbed office scanner, PDMS plate and utilize deep learning technique for automated phenotypic analysis. Our suggested system is low-cost and high-throughput, with automated analysis.



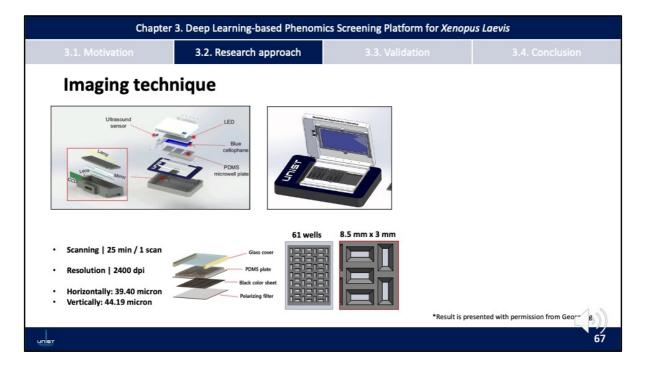
For drug screening, four different types were used under specific concentrations to cause noticable phenotypes under 72 hours post fertilization, affecting physical changes in body and tails of embryos. Figure shows the deformations visually and table summarizes the concentrations of each drug.



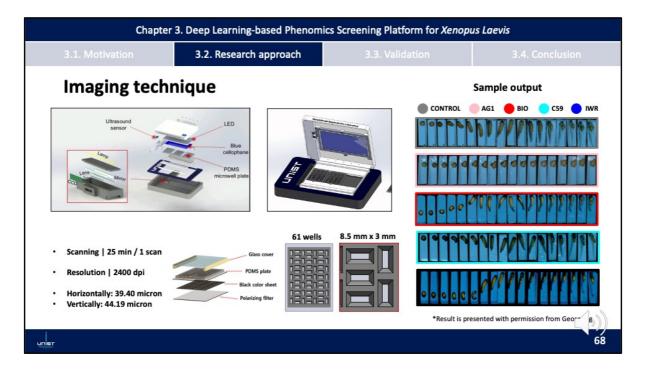
The figure demonstrates modified elements of the scanner. LED and blue cellophane was required to enhance contrast, while ultrasound was put to achieve automation.



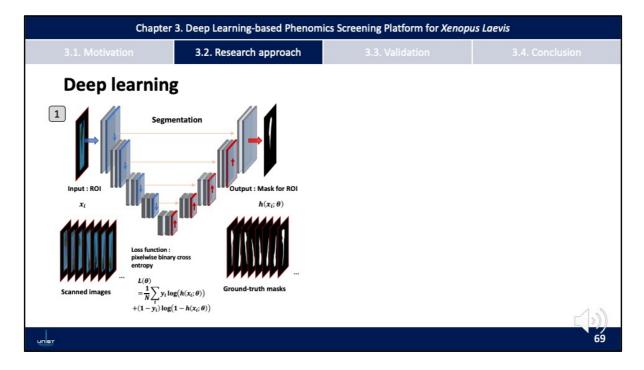
In this way, we performed imaging within 4 days by modified scanner taking 25 minutes per scan. The acquired image resolution is enough to resolve phenotypes. The temperature was below 25°C.



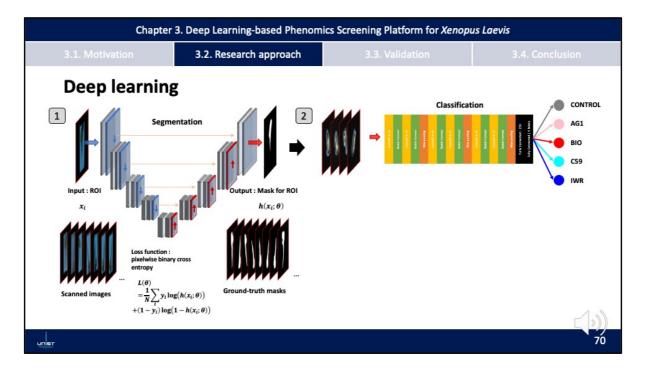
The suggested PDMS plate contains 61 wells and its design primarily targets for emrbyo studies.



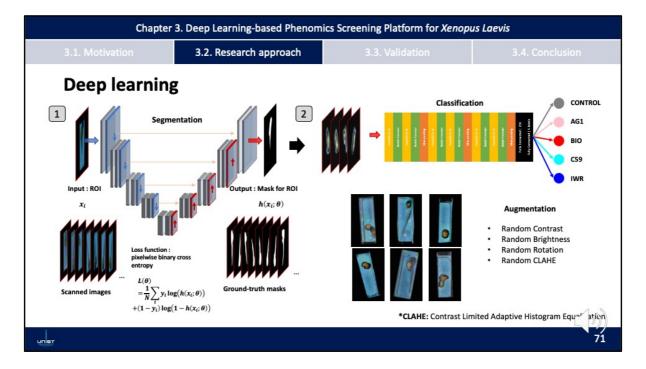
Sample output demonstrates the reduced effect of imaging artifacts in the figure.



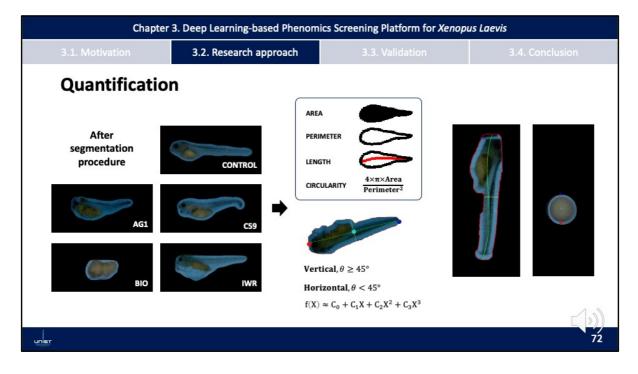
For analysis using deep learning technique, we acquired large number of corresponding manual masks. We utilized U-Net architecture for segmentation to locate embryo ROIs. At the input of our architecture, each individual well was cropped for the size of 900 by 600 pixels.



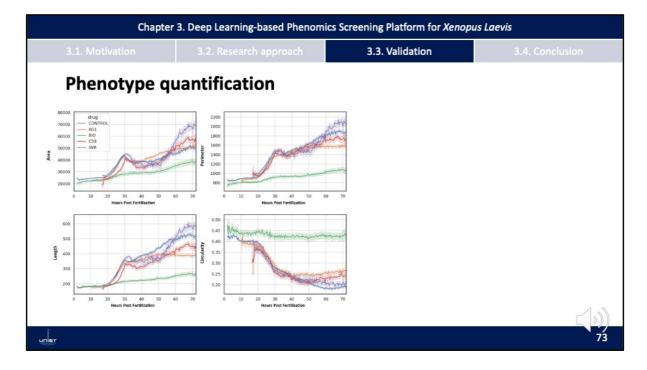
The segmented output was then further processed to the input of another CNN for the definition of drug related phenotypes.



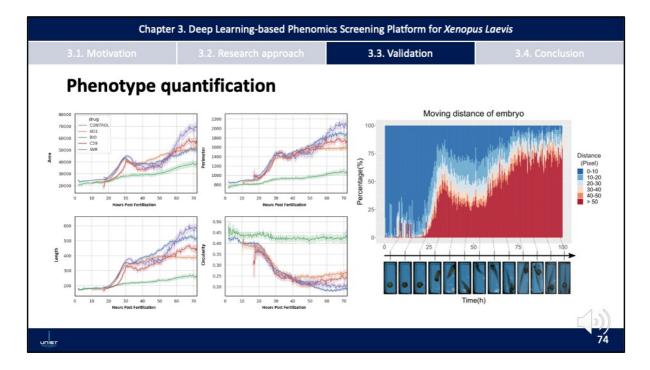
We applied augmentation strategy focusing on the generalization of the model towards color information.



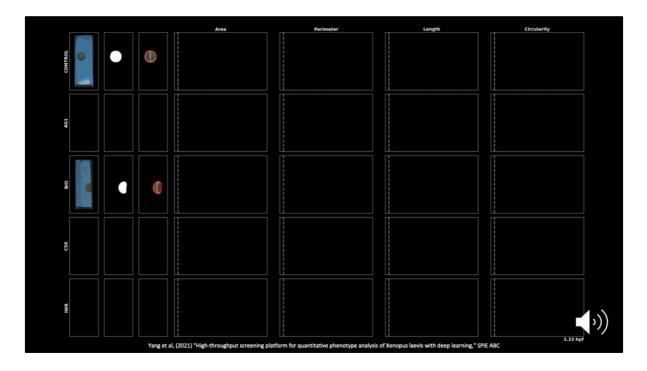
Acquired segmentation is basis for further quantitative analysis. In this outcome, we investigated four geometrical parameters: area, perimeter, length, and circularity. Our suggested evaluation could contribute to the observation of variant development dynamics as well as assist to early differentiation of signals.



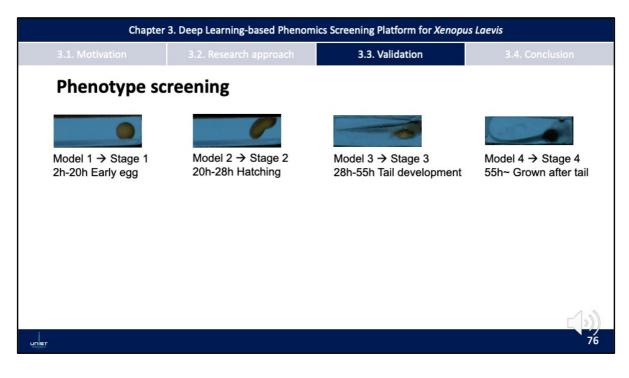
In this result, I will describe the quantitative evaluation of embryo's phenotypic changes from morphological perspective. We could observe various development dynamics under different drugs. Specifically, tracking the beginning of hatching stage or alterations in growth of tail at ~30 hpf and corresponding >60 hours post fertilization could provide meaningful insights for discovery of drugs, designed to target certain pathways.



Another characterization could possibly be the tracking of embryo activities with time by determination of the moving distance. For later stages of control cases, as expected, it is increased.



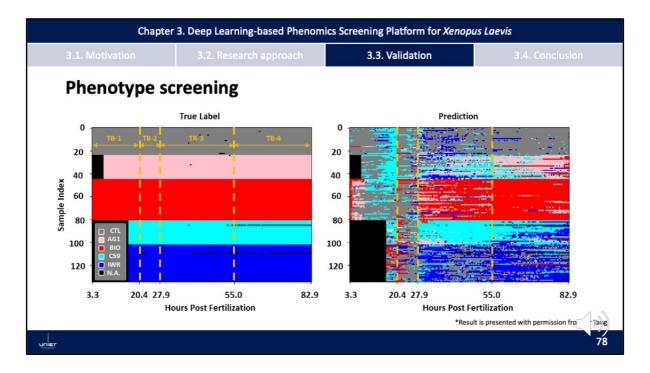
In this video, we present the results of quantification of segmented regions with deep learning. Control group and four drug cases are presented. Although, the work has been carried in post-processing mode, in the perspective, this results could contribute to building real-time, large-scale monitoring platforms for small aquatic animal models.



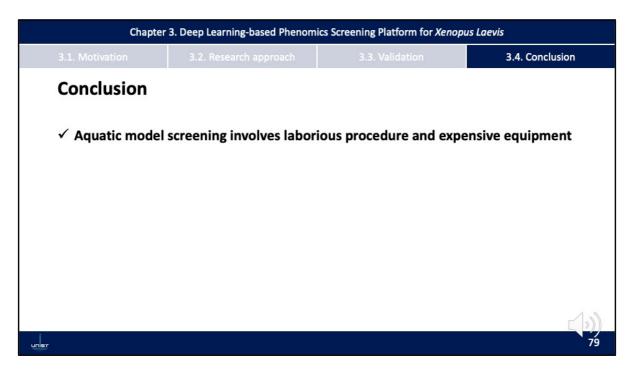
Although quantitative variations are provided, they are not sufficient to differentiate clearly which type of drug were treated before dramatic morphological changes. Therefore, we further developed CNN model for embryo images. We trained separate CNN classifiers for 4 developmental stages of Xenopus laevis: early egg, hatching, tail development, and grown tail.

	3.2. Research approach	3.3. Validation	3.4. Conclusion				
Phenotype screening							
	-		(in)				
Model 1 → Stage 1 2h-20h Early egg	Model 2 → Stage 2 20h-28h Hatching	Model 3 → Stage 3 28h-55h Tail development	Model 4 → Stage 4 55h~ Grown after tail				
Confusion matrix	CTL 0.474 0.035 0.125 0.331 0.035 0.9	Confusion matrix CTL 0.447 0.057 0.000 0.065 0.431 0.9	Confusion matrix CTL 0.891 0.001 0.000 0.037 0.072 0.9				
AG1 0.622 0.000 0.003 0.324 0.051 -0.7 BIO 0.785 0.009 0.101 0.085 0.020 -0.5 -0.6	AG1 - 0.391 0.005 0.095 0.378 0.130 - 0.8 - 0.7 - 0.6 BIO - 0.194 0.009 0.623 0.143 0.031 - 0.5	AG1 - 0.107 0.212 0.059 0.431 0.191 - 0.8 - 0.7 - 0.6 - 0.9 - 0.9 - 0.6 - 0.5 - 0.5	AG1 0.021 0.713 0.058 0.200 0.009 -0.7 BD0 0.004 0.023 0.942 0.020 0.010 -0.5 -0.4				
C59 0.249 0.085 0.116 0.545 0.005 - 0.4 WR 0.205 0.000 0.721 0.074 0.000 - 0.2	C59 0.531 0.108 0.281 0.080 0.000 - 0.3 - 0.2	P -0.4 C59 0.031 0.077 0.021 0.862 0.010 -0.4 IWR 0.133 0.002 0.007 0.61 0.797 -0.1	₽ 0.007 0.071 0.074 0.794 0.055 0.2 IWR 0.111 0.044 0.163 0.149 0.534 0.2				
Cr predicted label	Cr for for cr fo	Cr ↓ Predicted label	C' \$P & d Mr				

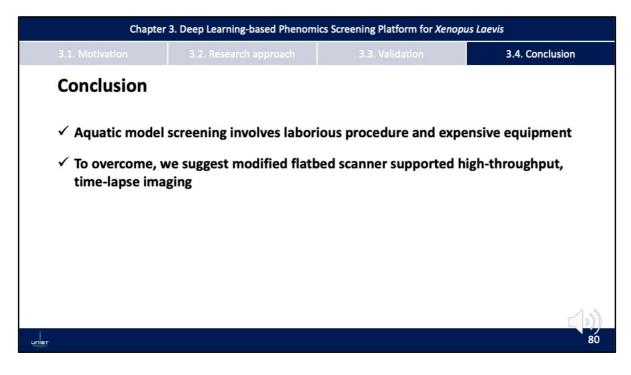
The acquired confusion matrix suggests the screening functionality can be confirmed at each developmental period, showing higher capability at later stages.



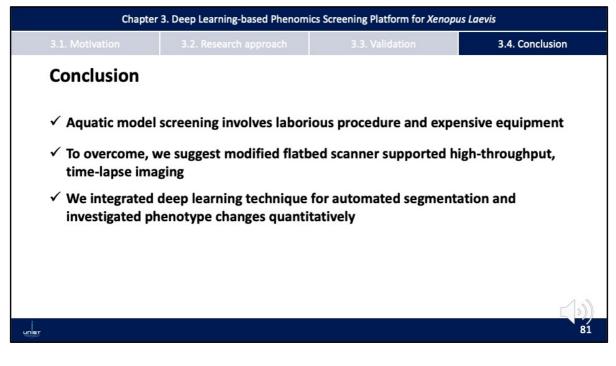
Below figure on the right shows classification results as heatmap of predicted labels compare to true labels from the validation dataset. In the egg and hatching stages classification accuracy is not sufficient for all drug types, however, along the tail development stage, drug treated samples resulted in higher classification accuracy levels.



To conclude, Xenopus screening involves laborious procedure and expensive equipment.



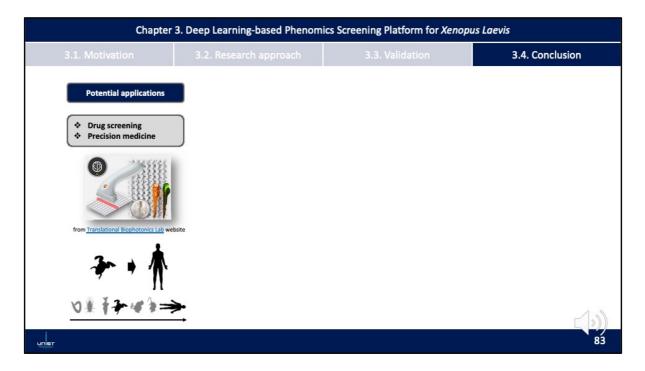
Therefore, in this study, we developed the high-throughput screening platform with modifying flatbed office scanner.



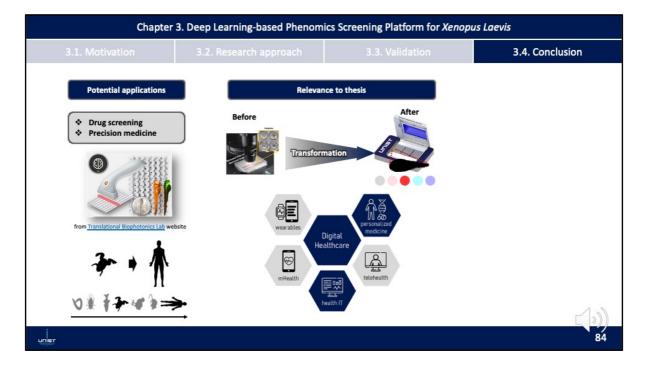
Based on the power of massive data acquisition and the deep learningbased technique, quantitative phenotype analysis, and automated screening capability.

Chapter 3. Deep Learning-based Phenomics Screening Platform for Xenopus Laevis							
	3.2. Research approach	3.3. Validation	3.4. Conclusion				
Conclusion							
\checkmark Aquatic model screening involves laborious procedure and expensive equipment							
✓ To overcome, we suggest modified flatbed scanner supported high-throughput, time-lapse imaging							
 We integrated deep learning technique for automated segmentation and investigated phenotype changes quantitatively 							
 Our results suggest proposed platform has potential to become a promising tool for massive and dynamic observation, and could be applied to developmental studies, drug testing, and phenotype-genotype assays 							
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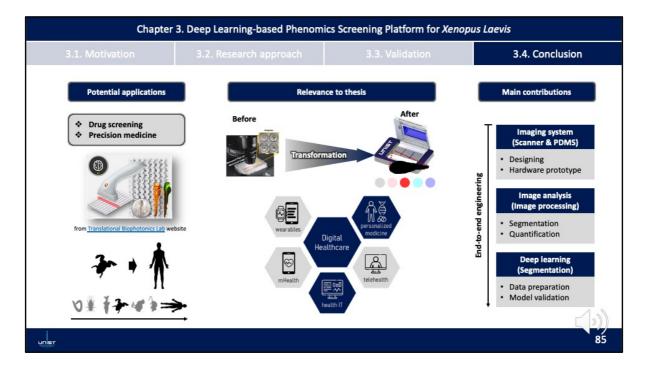
The proposed platform could become a promising tool in massive and dynamic observation based biological studies, such as developmental studies, drug testing, and phenotype-genotype assays.



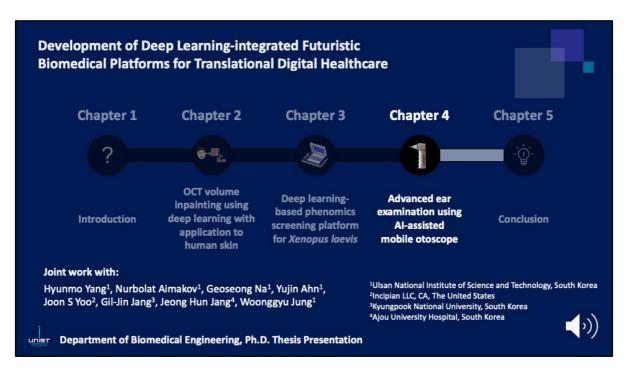
This project could potentially be useful for drug screening or precision medicine applications.



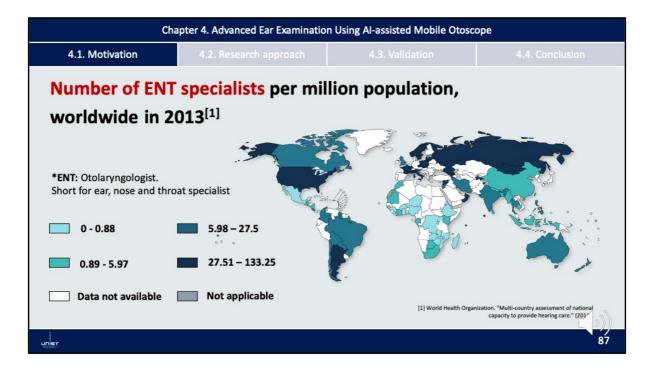
Within bounds of my thesis, I have explored the transformation of aquatic model screening towards novel deep learning-integrated smart device. It could in perspective contribute towards the personalized medicine, or health IT directions of digital healthcare technology.



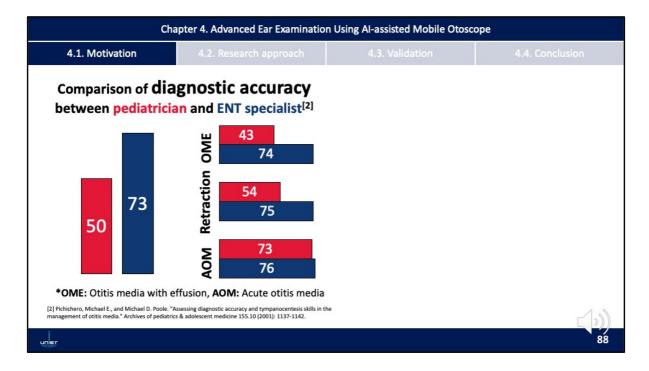
Finally, here I outlined my personal contributions inside the research work towards end-to-end engineering.



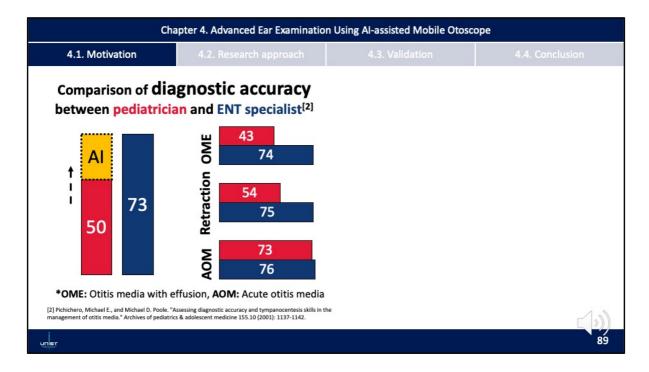
In the last chapter, I will present about the development of "Advanced ear examination using AI-assisted mobile otoscope". This is a joint work with Professor Jang from Ajou university hospital and several researchers.



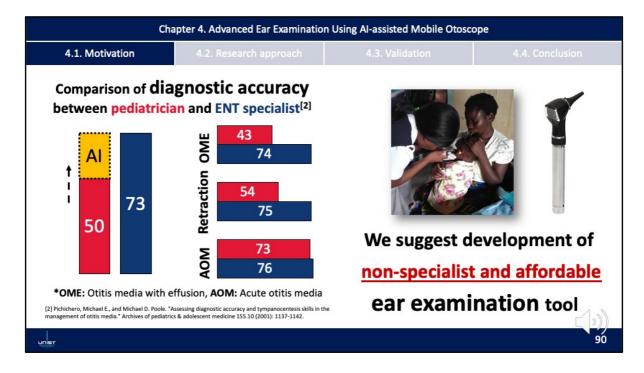
In 2013 multi-country assessment of hearing care capacity held by World Healthcare Organization, there was a statistics regarding the number of ENT specialists' availability across various countries. Just to clarify, ENT specialist is an otolaryngology doctor equivalent and stands short for ear, nose and throat. A critical finding to pay attention in this report is that there is a considerable number of countries which lacks ENT doctors in low and lower-middle income segment.



Essentially, such an absence of expertise knowledge in this domain could negatively affect in proper medical treatment decisions. For instance, studies have indicated a 23% distinction in technical competence between pediatricians and specialists to correctly diagnose possible a condition.



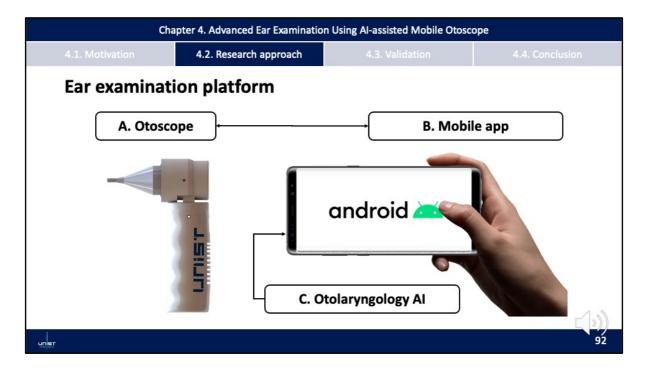
However, the recent advancement of the deep learning opens a possibility to compensate the current limitation of physician knowledge to reach ENT level diagnosis.



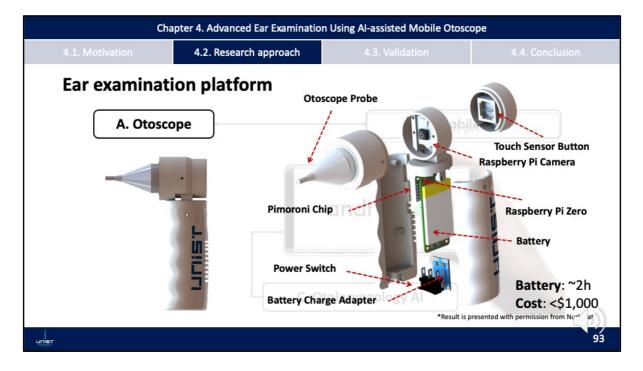
Therefore, we believe equipping non-specialists with an assistive technology in an affordable manner could support a more accurate diagnosis, thereby improving the ear examination situation within low resource settings.

Study		Application	Accuracy]
Basaran (202	20) ^[3]	Diagnosis of middle ear inflammation	90.48%	
Cha (2019) ^{[4}]	Detection of ear and mastoid disease	93.67%	
Livingstone	(2019) ^[5]	Otologic disease screening	84.4%	1

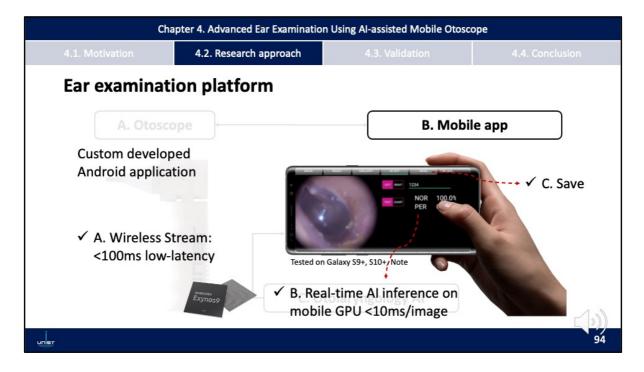
A survey of previously reported research efforts has demonstrated on the potential of AI in otolaryngology. These consider different applications, including diagnosis of middle ear inflammation and otologic screening purposes. More specifically, they unite convolutional neural networks as dominant machine learning technique with high performance.



However, to tackle the real-world problem a holistic approach in the form of targeting platform is required. Therefore here, we suggest a custom otoscope and mobile phone application with on-device AI an accessible platform to enhance traditional ear examination with minimal training. A physicial component was determined from the perspective of the end user, who is potentially a general physician residing without superior infrastructure.



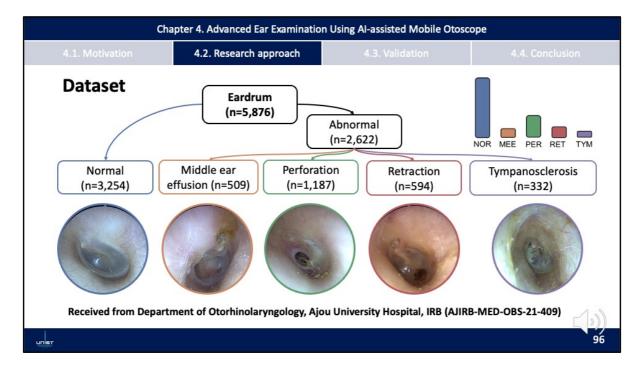
Our suggested otoscope is minimalist and effectively adapted for the ear inspection purpose. Empirically, it withstands 2 hours of continuous operation and lies under \$1000 price range.



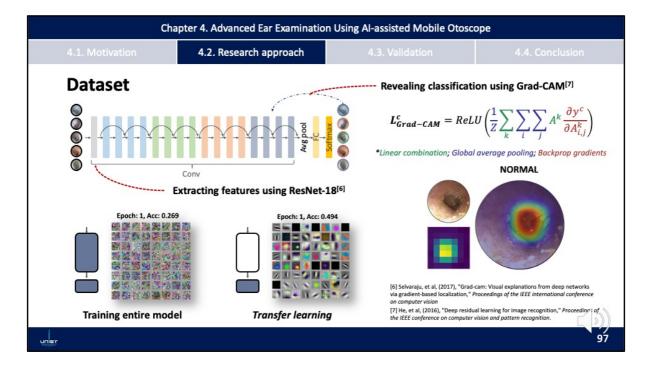
As of the mobile application side, it is based on the customly developed Android software merging several functionality, such as low-latency video streaming, saving image for later review, and real-time on-device AI inference

Chapter 4. Advanced Ear Examination Using Al-assisted Mobile Otoscope						
4.1. Motivation	4.2. Research approach		4.4. Conclusion			
Ear examinat	ion platform					
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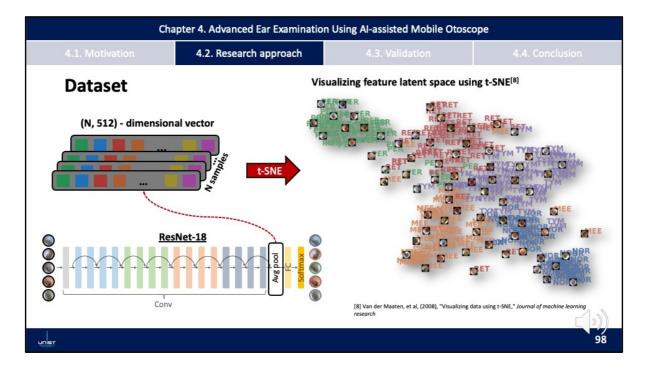
Further, let me focus your attention on our deep learning approach used for this study.



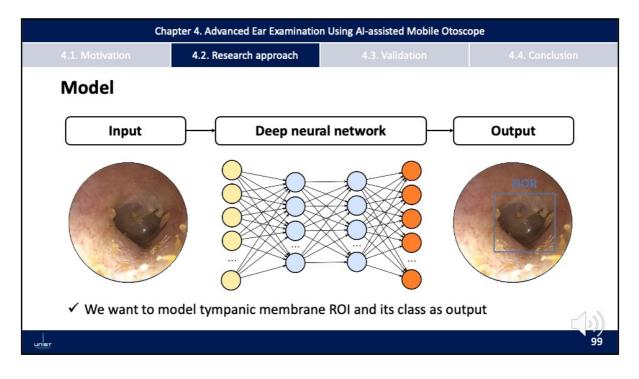
The integral part of the deep learning technique is data acquisition. Following the IRB protocol, our collaborators from the Department of Otolaryngology of Ajou University Hospital in South Korea provided us with 5 commonly represented ear conditions accumulated within a clinical setup, resulting in over 5,800 unique patient cases.



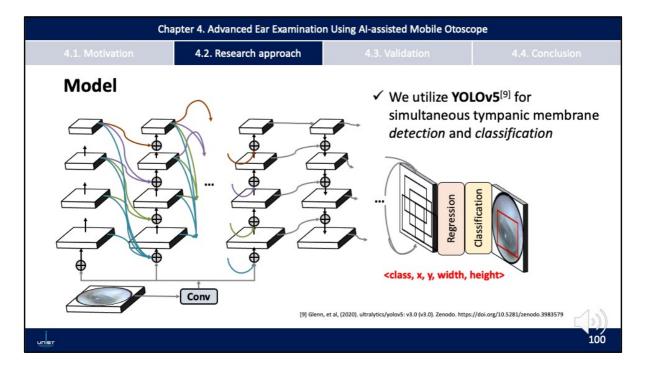
Based on acquired data, with the help of existing explainable AI techniques, we were curious to identify which patterns and meaningful regions contribute to the discrimination of ear diseases. Our observation led to constitute that region inside the actual tympanic membrane is more important to successfully classify diseases.



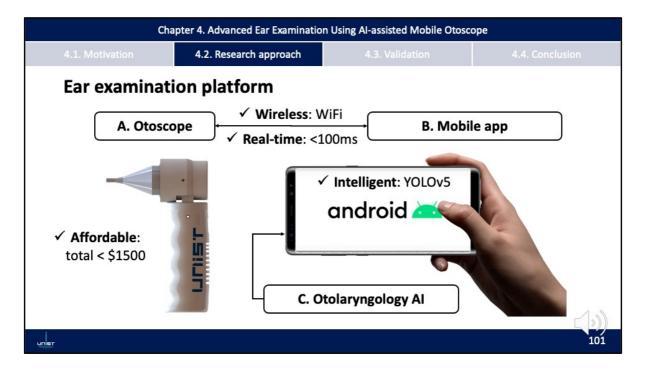
A further interesting investigation would be knowling how machine could independently group various diseases. For this purpose, we utilized latent space of feature vector, extracted from the last convolutional layer of basic ResNet architecture. This vector was consequently transformed into 2D space using t-SNE dimensionality reduction technique. This small experiment, apparently, demonstrates the formation of clusters.



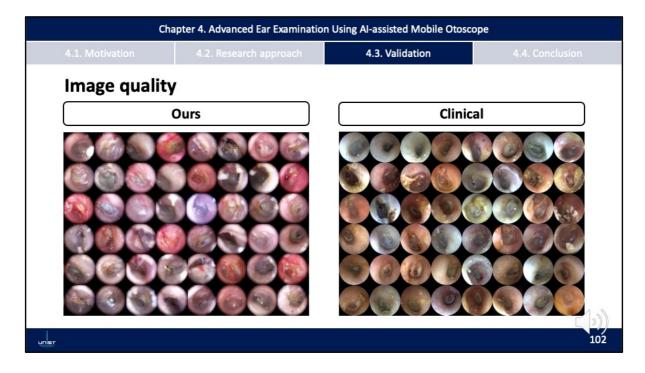
Our previous observations suggest that AI is in favor to focus on the tympanic membrane region and could cluster clearly. Therefore, it led us to impose a requirement in the development of deep neural network component with object detection-oriented task.



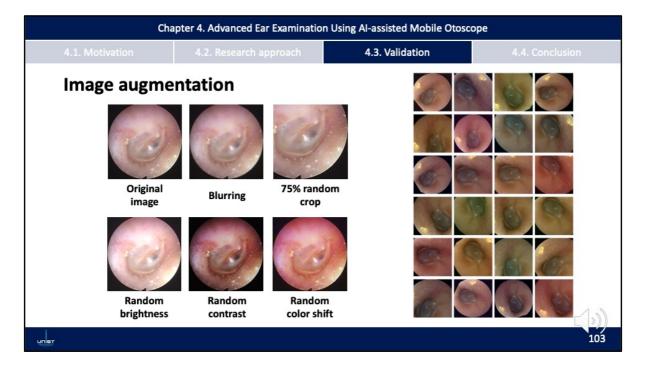
When we addressed our attention to the computer vision field, YOLOv5 has been a prominent method combining robust performance and quick response. Although at architectural complexity, the efforts of open-source community made this model significant and well-engineered, which influenced our decision to utilize it for tympanic membrane data.



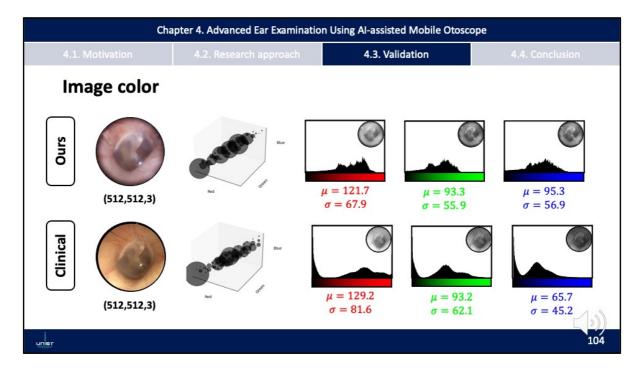
Let me reiterate on our suggested ear examination platform one more time. It consists of the custom otoscope, mobile application, and on-device AI model to support the real-time diagnosis.



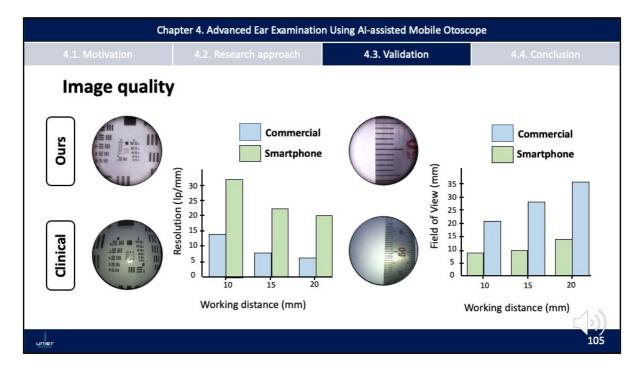
We were interested to visually compare images acquired from our developed otoscope and the clinical setup. Therefore, in this slide, we summarize the results of 48 randomly picked samples after the conducted field test examinations in Ajou University Hospital.



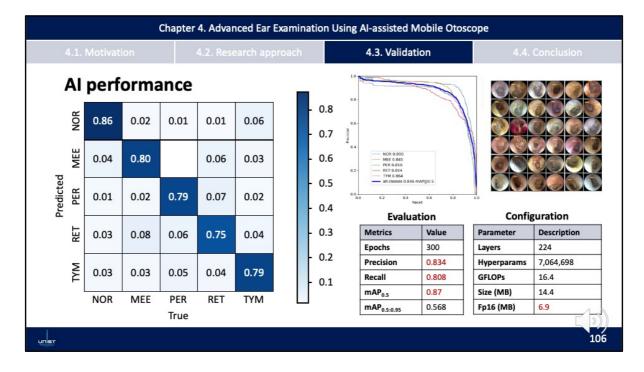
To minimize the effect of inherent distinctions present between clinical setup and our platform in image color and image quality, we additionally introduced blurring and random color shift operations during dataset augmentation to increase the overall accuracy of the deep learning model.



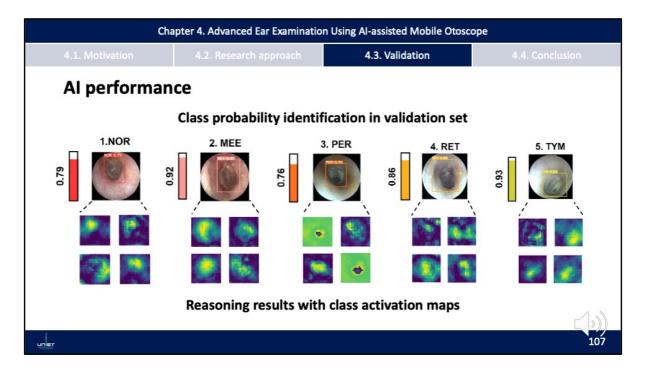
For instance, in this slide, you can see a summary of color variance investigation between our device and clinical setup.



Let's also take a closer look at both images in terms of the quality, characterized by resolution and field of view parameters. At the smaller field of view, our suggested setup provides a superior resolution at working distances of 10, 15, and 20 mm, within which tympanic membrane distance is typically represented.



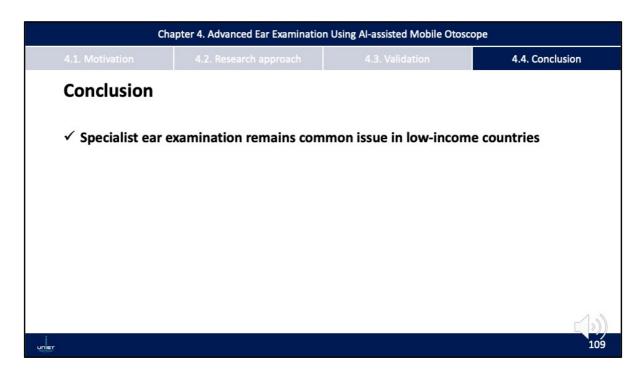
Our inspection of the AI model performance suggested on the accurate results for tympanic membrane detection and classification, with overall of over 83% precision and 80% recall. Interestingly, the compressed trained model produced only 7MB of size, which states on the practicability to process inference on the mobile devices in real-time.



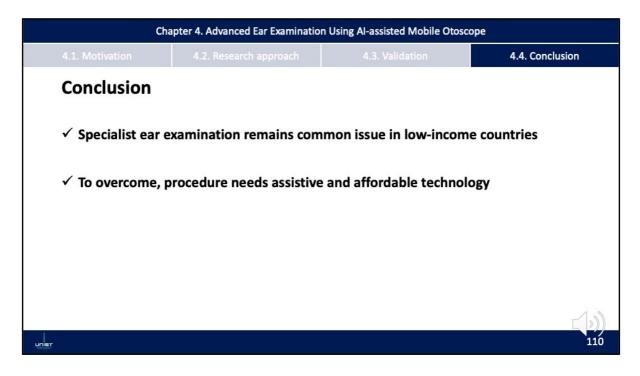
One can further analyze the reasoning behind the deep learning model performance and find important regions contributing inference through the investigation of activation maps at certain levels. This research direction is another central topic of the deep learning community and is under active exploration.



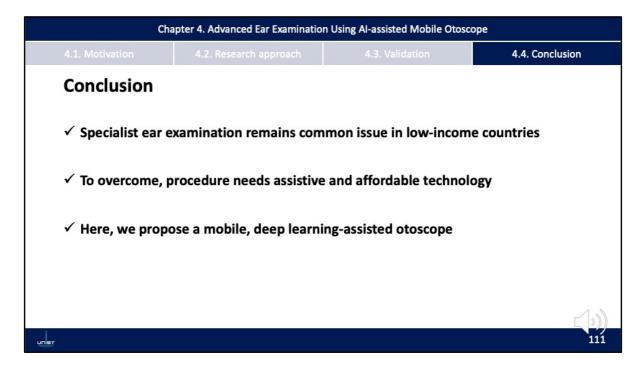
We finally demonstrate the actual protocol of user application in this video. First, the doctor connects the device, enters relevant patient information and does the necessary setup. After that he can start the ear inspection procedure. He is supported with real-time AI diagnosis and can capture images for later review.



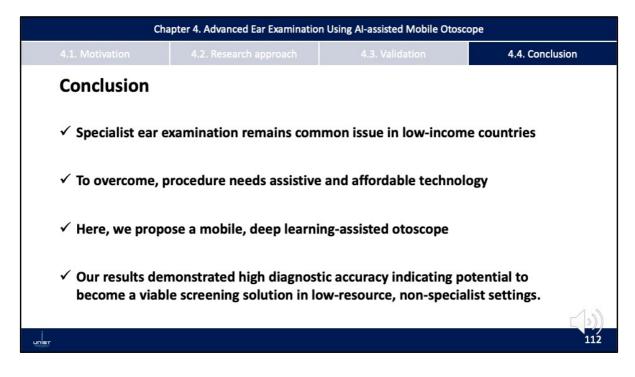
Specialist ear examination remains common issue in low-income countries.



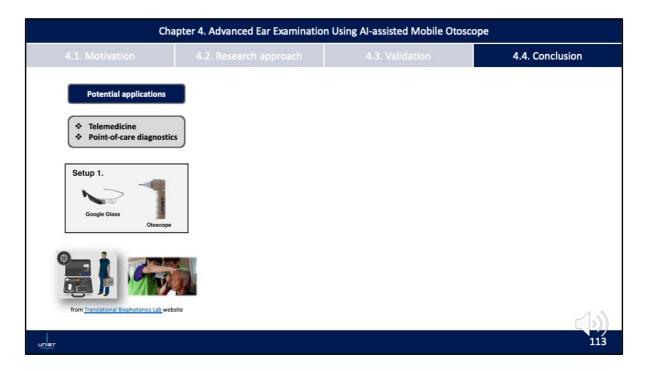
To overcome this problem, we believe procedure needs a supportive and affordable technology.



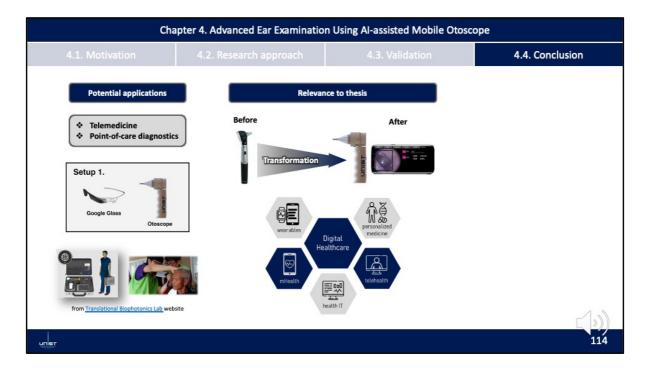
In this study, we propose a mobile, deep learning-assisted otoscope for this role.



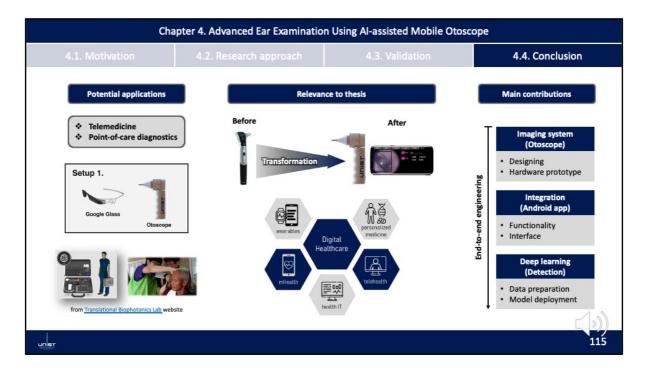
Our results demonstrated high diagnostic accuracy indicating potential to become a viable screening solution in low-resource, non-specialist settings.



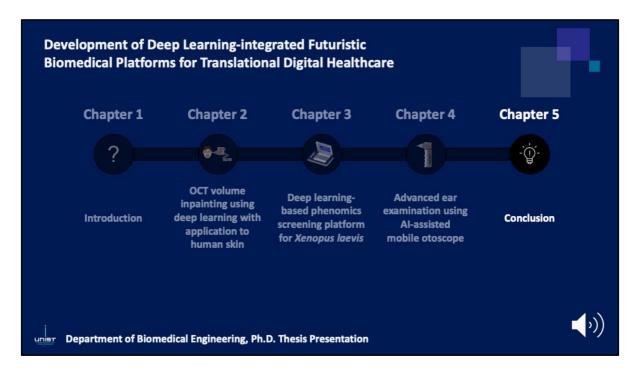
This project could potentially be useful for telemedicine or point-of-care diagnostics.



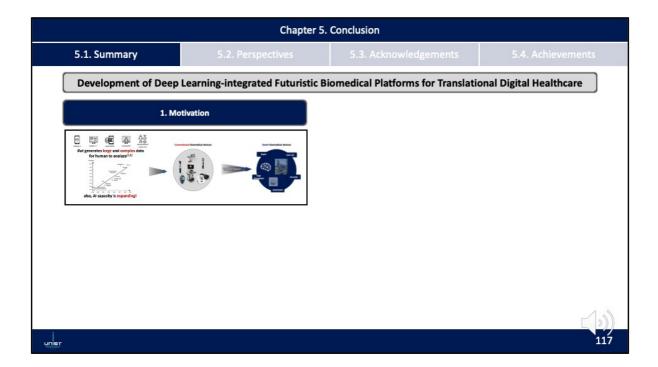
Within bounds of my thesis, I have explored the transformation of traditional ear examination tool towards novel deep learning-integrated smart device. It could in perspective contribute towards the mobile health, or telehealth directions of digital healthcare technology.



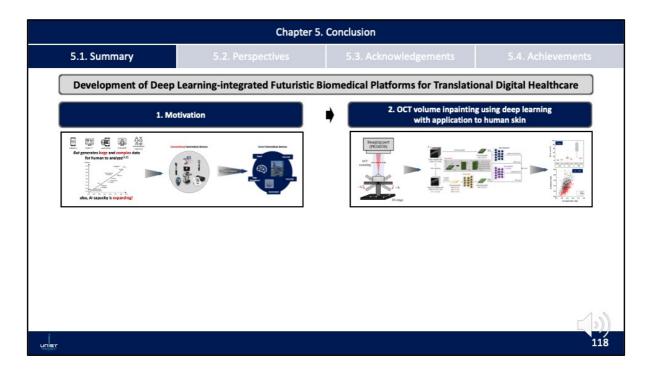
Finally, here I outlined my personal contributions inside the research work towards end-to-end engineering.



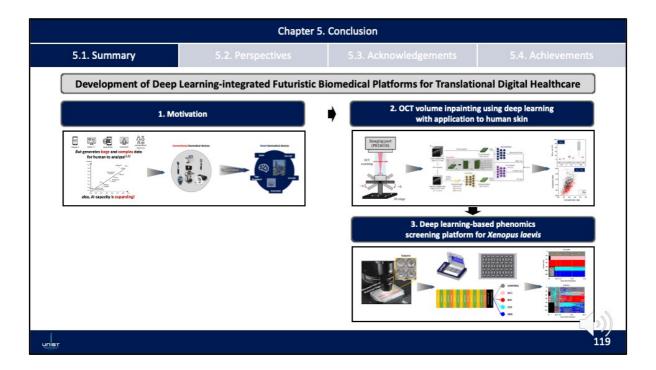
Let me finally conclude today's presentation.



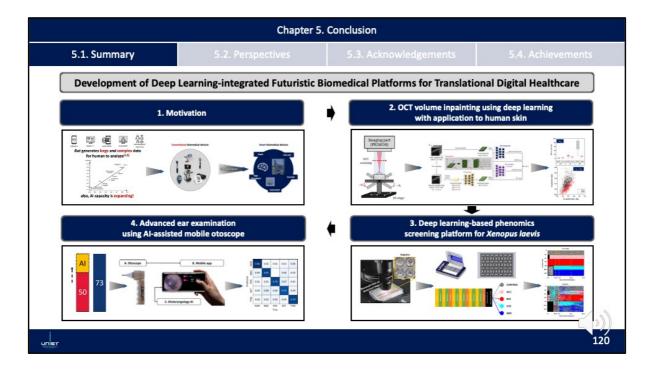
The extensive use of digital health technologies creates challenge of analysing large and complex data, where human is incapable. At the same time, the capacity of artificial intelligence keeps expanding. One of the opportunities could be the suggestion of assistive smart devices. Currently, conventional biomedical devices are developed with fixed requirements. They are often primitive and static. In contrast, smart biomedical devices possess built-in intelligence to support human decision. The transformation from one category to another is an open question. And the exploration of such opportunities was my primary interest during the PhD program.



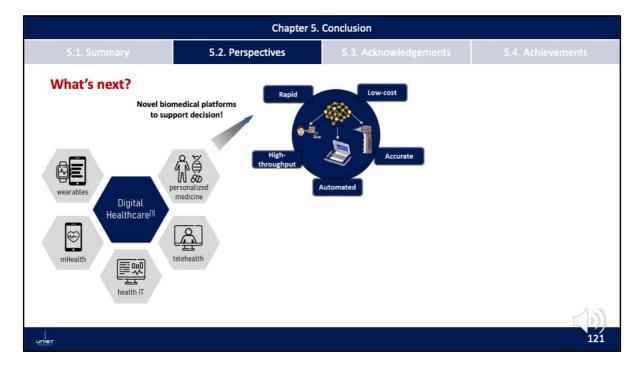
In chapter 2, I presented advantages of OCT for human skin study and how deep learning technique could be integrated for volume inpainting in order to make OCT scanning more efficient. Our results are promising, showing little deviation of image quality restoration parameters as well as roughness compared to the conventional interpolation method.



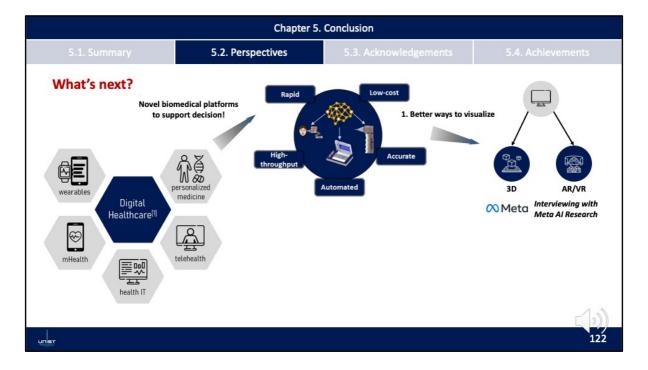
In chapter 3, we addressed challenges of phenotype screening problem by the development of the high-throughput, modified scanning platform. Here, deep learning technique was utilized for further automated phenotypic analysis and screening of massive embryo timelapse images. The proposed platform could become a promising tool in dynamic observation based developmental studies or drug testing applications.



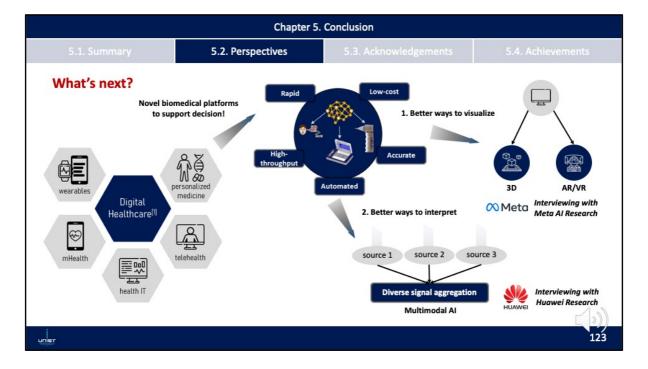
Finally, in chapter 4, we target the shortage of ENT doctors in low-income countries. For that reason, we suggest mobile otoscope platform with on-device AI to equip non-specialists with assistive technology in an affordable manner. Our results suggest it could support a more accurate diagnosis, thereby improving the ear examination situation within low resource settings.



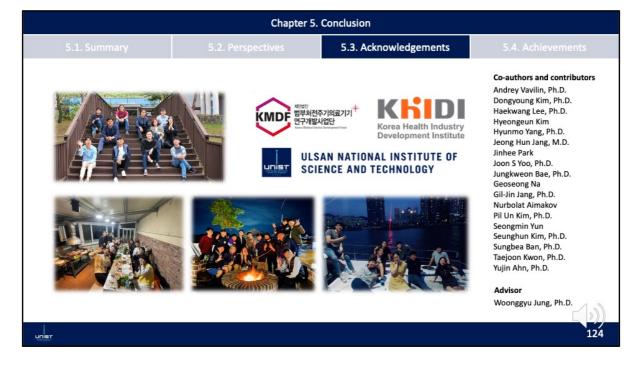
In this work, I focused on the development of novel biomedical platforms for digital healthcare, integrating deep learning techniques. In the perspective, I believe there are two interesting ways for my research and professional trajectory.



One could possibly be exploring better ways of visualization and making more the overall experience of delivery more convenient. These could include extension for three-dimensional technologies, as well as adaptation of AR/VR technology.



Another direction is investigating additional ways to interpret information by understanding the richness and diversity from various source signals. For this reason, I believe multimodal AI techniques could assist this task.



I would like to thank my advisor; all the contributors participating in the projects, acknowledge research fundings supporting projects and you, committee members for feedbacks.

	Chapter 5.	Conclusion	
		5.3. Acknowledgements	5.4. Achievements
 Syun*, H Yang*, S Askaruly*, G Na, J Bae, W Ji Y Ahn*, J Park*, S Askaruly*, D Kim, G Jang, W Ji S Askaruly*, H Yang*, N Aimakov*, G Na, Y Ahn Jik Bae, H Roh, JS You, K Kim, Y Ahn, S Askaruly, Inspection after acetic acid using machine Ban, PU Ki Quantum Electronics 25 (2), 1-8 S Kim, Yujin Ahn, S Askaruly, P Kim, W Jung, H U Chal Presentations G Na, H Yang, N Aimakov, G Na, Y Ahn, JS G Na, H Yang, U Aimakov, G Na, Y Ahn, JS G Na, H Yang, U Aimakov, G Na, T Kwon, W Jung, T W S Yang, S Askaruly, S Yun, G Na, T Kwon, W Jung, T W S Askaruly, Y Ahn, J Bak, A Vavilin, G Jang, P Kin S Askaruly, N Ahn, J Bak, A Vavilin, G Jang, P Kin S Askaruly, N Aimakov, A Islakov, H Cho, Y Ahn S Askaruly, N Aimakov, A Islakov, H Cho, Y Ahn S Askaruly, Y Ahn, H Kim, A Vavilin, PU Kim, H Li 	ung, T Kwon, "XenoScan: Deep learning-based phenomics si tung, "OCT volumetric inpainting using deep learning netwo IS You, G Jang, H Jang, W Jung, "Advanced ear examinati K Park, H Yang, G Jang, K Moon, W Jung (2020). "Quantitating ing techniques," <i>JMIR mHealth and uHealth</i> 8 (3), e16467 m, S Kim, H Lee, W Jung (2018). "Quantitative evaluation o ee (2017). "Evaluation of skin texture and wrinkle using opi You, G Jang, JH Jang, W Jung (2022) "Advanced ear examin. , Y Lee, W Jung (2022) "High-throughput screening with de (2021), "Evaluation of deep-learning-based high-thro von (2021), "Development of deep-learning-based high-thro , H Lee, W Jung (2018), "Quantitative classification of OCT , MH Choi, H Yang, W Jung, "Farabio: Deep learning for bior ee, W Jung (2017), "Evaluation of age-related effects on hu aratus for detecting wrinkle of skin using optical coherence r 2021, <u>https://tbl-unist.github.jo/mobile-ai-21/</u>	creening platform for aquatic model organism development, rk with application to human skin", <i>in preparation to SCI</i> no using deep learning-assisted mobile ootscoep", <i>in prepara</i> f skin surface roughness using optical coherence tomograph sical coherence tomography (Pilot study)," <i>Journal of the Soc</i> ation using deep learning-assisted mobile otoscoepe", <i>SPIE Ph</i> ep learning for quantitative phenotype analysis of zebrafish' the phenotype analysis of <i>Xenopus leavis</i> with deep learning bughput phenotype screening platform of aquatic model org skin images with deep learning." <i>SPIE Photonics West</i>	ation to SCI ation to SCI its study of smartphone-based endoscopic visual y in vivo," IEEE Journal of Selected Topics in lety of Cosmetic Scientists of Korea 43 (3), 247-254 otonics West SPIE Photonics West "SPIE Advanced Biophotonics Conference anism embryos," International Xenopus Society aphy," The optical society of Korea
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Here I have listed research activities during the period of my Ph.D. program. I am currently working on final preparations of manuscripts for submission.

5.1. Summary		5.3. Acknowledgements	5.4. Achievements
in paper is in preparation to OPTIO	. Will be submitted in March.		
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One of them regarding the skin project will be edited more by Professor Jung. He has not seen it yet however we believe it to be submitted within March.

Closing remarks

Dear committee members,

Before we go to the Q&A session, I would like to say a few words of closing remarks.

I would like to thank you for the valuable feedbacks which helped me to grow not only from our last meeting, but also for the continuous support during the whole period of my PhD program.

When I look back, I feel to have grown a lot. I widened my outlook and deepened expertise. Also, I acquired many skills, which hopefully will serve well in my later career. But more importantly, I believe I became more patient and humble. Although this journey was challenging, quite often I was lost in the curiosity of

research and passion for engineering. My acknowledgement here primarily goes to my family, friends, labmates. The role of my supervisor, professor Woonggyu Jung is doubtlessly huge. He always kept advising, supporting and cheering up for whom I am grateful, especially I admire his endless enthusiasm :)

Further, I hope to wrap up the works within the soonest period and concentrate on the next stage of life, building professional career. Given the time and resources, I am now confident to explore opportunities in the development end-to-end solutions for biomedical fields and not only, thanks to obtained research training. I am also grateful to UNIST, and hope I can contribute to its international reputation later in my career as its alumni.

That said, I am ready for your comments and expert opinions, to evaluate the qualification of the presented thesis.



Finally, thank you for your time and attention. If you have any questions regarding the defense presentation, I would be glad to answer them now.