PHARMACHAIN: A BLOCKCHAIN TO ENSURE COUNTERFEIT FREE PHARMACEUTICAL SUPPLY CHAIN

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ABSTRACT

Access to essential medication is a primary right of every individual in all developed, developing and underdeveloped countries. This can be fulfilled by pharmaceutical supply chains (PSC) in place which will eliminate the boundaries between different organizations and will equip them to work collectively to make medicines reach even the remote corners of the globe. Due to multiple entities, which are geographically widespread, being involved and very complex goods and economic flows, PSC is very difficult to audit and resolve any issues involved. This has given rise to many issues, including increased threats of counterfeiting, inaccurate information propagation throughout the network because of data fragmentation, lack of customer confidence and delays in distribution of medication to the place in need. Hence, there is a strong need for robust PSC which is transparent to all parties involved and in which the whole journey of medicine from manufacturer to consumer can be tracked and traced easily. This will not only build safety for the consumers, but will also help manufacturers to build confidence among consumers and increase sales. In this article, a novel Distributed Ledger Technology (DLT) based transparent supply chain architecture is proposed and a proof-of-concept is implemented. Efficiency and scalability of the proposed architecture is evaluated and compared with existing solutions.

Keywords Distributed Ledger Technology (DLT) · Blockchain · Smart Contracts (SC) · Pharmaceutical Supply Chain (PSC) · Transparent Supply Chain · Counterfeit Medication

I. Introduction

Essential medicine access is one of the main objectives of healthcare systems in and pharmaceutical supply chains ensure that the right amount of medicine, with acceptable quality, will reach the customers in need, at the right time [1]. The supply chain provides smooth operation of cross-organizational business processes like procurement of raw materials, manufacturing, and delivery of finished goods to the customers. This not only helps customers but also the organizations to reshape their workable boundaries [2]. The pharmaceutical supply chain on the other hand is very complex in terms of data and payment flow as it involves multiple entities, including manufacturers, mail-order retailers and brick and mortar pharmacies, wholesalers, health insurance corporations, Pharmacy Benefit Managers (PBM) and the consumer. Multiple factors, including storage environment parameters, demand and usage patterns affect the flows and associations between these entities. The complexity of flows and the limited visibility into the supply chain activities has led to multiple risk factors which have not only been wasting the resources available but also are risking the lives of consumers [3]. Among this complex and untraceable supply chain, introducing

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counterfeit drugs into the supply chain with inefficient to no Active Pharmaceutical Ingredients (API), drugs produced in sub-standard conditions and even repackaging of expired medicines, can be easily done and is the cause for the majority of problems in the pharmaceutical industry [4]. Detecting such players in the supply chain with the malicious intent of introducing counterfeit drugs and penalizing them is a very difficult process considering this tangled supply chain [5]. One such recent incident Took place when Canadian drug wholesaler SB Medical has clearly shown sub-standard sanitary conditions where drugs lacking proper ventilation, temperature, humidity and security have been shipped to US doctors and medical clinics [6]. It is clearly visible, how the counterfeit drugs are acquired from unsafe locations and have been staged at different places for repackaging before introducing them into the US Pharmaceutical supply chain. It is estimated that 30% of drugs sold in 140 countries at an approximate cost of \$250 billion per year, are believed to be counterfeit [7].

A typical pharmaceutical supply chain is shown in Figure 1 [8, 9]. Manufacturers are the source for both brand name and generic drugs in the pharmaceutical supply chain. Based on the type of drug and active pharmaceutical ingredients, manufactures acquire the required raw materials from ingredient suppliers. Most of the brand drug manufacturers allocate some portion of the funds for research and development of new drugs. Any new drug formulated by these manufacturers has to undergo a series of tests in order to weigh the benefits compared to the risks along with the efficiency of the drug through drug trials. Once the drug information is gathered, the manufacturer contacts the Center for Drug Evaluation and Research (CDER) and the Food and Drug Administration (FDA) for approval before starting manufacturing and selling the drug. The FDA performs a series of tests and evaluates the CDER submitted to ensure that the drug is safe and outweighs related risks. Once this process is done labeling is proposed to the manufacturers along with approval to manufacture and sell. The manufacturers are also responsible for the distribution of drugs from facilities to retail pharmacies or to distributors for further reach in the supply chain. Another flow includes drugs flowing back from pharmacies to the manufacturer as a part of buy back programs which accommodates financial risks for pharmacies to stock up the resources.

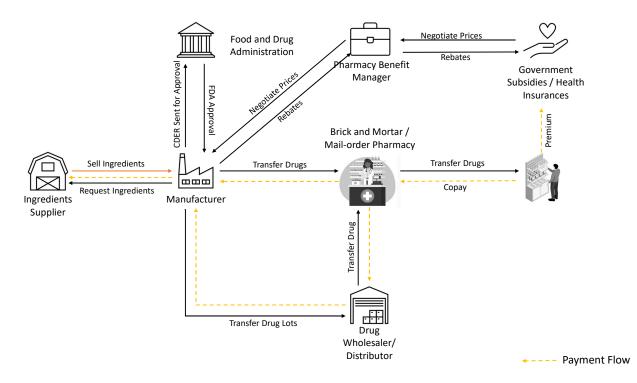


Fig. 1. Goods and Financial Transaction Flows in Pharmaceutical Supply Chain.

Wholesale distributors are responsible for purchasing the lots of manufactured medicine from manufacturers and distribute them to different customers including hospitals, pharmacies, care centers, etc. Wholesalers sell a variety of drugs and some wholesale distributors are specialized in particular drug distribution. Currently, wholesalers are also responsible for stocking and warehousing until there is demand and pharmacies place an order. The duties of pharmacies include ensuring that drugs reach consumers from wholesalers. They are also responsible for the safety of the stock and for fulfilling consumer demand along with providing safe usage of drugs. Pharmacy Benefit

Managers (PBM) are intermediaries responsible for negotiating prices from manufacturers and transfer those rebates to government agencies and insurance companies.

A. Problems with the Traditional Supply Chain in the Pharmaceutical Industry

Fragmentation of information is the first and foremost problem, as the pharmaceutical supply chain is comprised of many different active and passive entities [10, 11]. These entities become aware of different transactions between only when they are involved, resulting in blinded parties, as shown in Figure 2. Hence, there is no single source of truth for all entities to keep track of the goods within the supply chain. This results in discrepancies and difficulty in working collectively and delays supply chain operations. As the data is being introduced by multiple channels from different entities and information is fragmented, taking unilateral decisions by all entities in the supply chain will be difficult. This will also hinder manufacturers from forecasting demand and causes issues in inventory management. Drug shortages due to this incorrect forecast can lead to several public health problems [12]. It can be clearly seen during the COVID-19 pandemic how the shortage of vaccines and pharmaceuticals has caused unnecessary suffering and potential deaths [13].

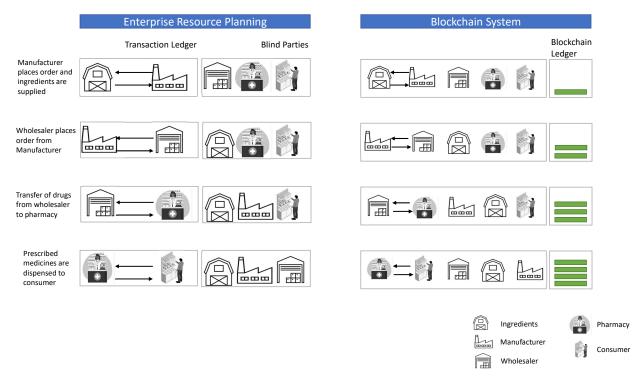


Fig. 2. Blind Parties in Conventional Record Keeping.

B. Distributed Ledger Systems Background

New technologies in the digital revolution such as the Internet-of-Things (IoT), Big data, New Information and Communication Technologies (ICTs), Distributed applications and Distributed Ledger Technologies (DLTs) have paved the path to improve and implement an efficient supply chain mechanism which can benefit the pharmaceutical industry. DLT is one of the latest technologies which has revolutionized financial systems by removing middlemen like banks and other financial institutions to implement fully decentralized peer-to-peer banking systems [14]. Advantages of DLT technologies arise in many different fields, including smart cities[15, 16], smart healthcare [17, 18, 19], smart farming [20, 21] and also in building efficient and transparent supply chains [22]. These transparent supply chains can benefit not only the manufacturers with acquiring raw materials and with the smooth production of goods and delivery but also customers will have confidence in product consumption as counterfeit detection and avoiding can be managed easily. A widely used and accepted distributed network is the blockchain but it is specialized for banking and payment systems. As one size doesn't fit all applications, several other DLTs came into the picture either to increase the scalability, to reduce resource utilization, to increase transaction speeds, provide smart contract

support, efficient consensus mechanisms, etc., based on different needs of applications. Based on different milestones, the development of these DLTs is classified into three generations.

The first generation started with the Bitcoin network which was developed by the pseudonymous developer Satoshi Nakamoto [23]. First generation Blockchains including Bitcoin, and Litecoin are mainly developed to address centralization issues in banking systems and to provide an efficient and fast peer-to-peer verifiable financial system to transfer digital assets. Bitcoin has made use of complex cryptographic techniques in order to achieve immutable, easily verifiable systems which can prevent double spending of digital assets and provides an immutable secure ledger. As the popularity of the blockchain has increased and its wide adoption is on the way, many different use cases of the technology in other fields has been unveiled, which needed further modifications. In the second generation, the main idea is to include the trust agreements along with keeping track of digital assets. This can help in automating and digitizing the contracts between untrusted parties which can be triggered and executed based on the events. These pieces of digitized agreements are termed "Smart Contracts" and were first implemented in Ethereum [24]. This has made blockchain processes available for incorporation into business processes, which made them more aligned with different business needs. Although blockchain technology has matured, there are still issues with scalability, transaction costs, energy utilization and specialized hardware requirements. In order to address these issues, many different platforms have been developed including Cardano, IOTA, Hedera Hashgraph, etc. These platforms are either updating the consensus mechanism behind the network or entirely changing the structure of the chain in order to increase transaction speeds, and provide more security along with increasing scalability.

1) Consenus Mechanism Overview

Distributed networks are prone to Byzantine fault and Sybil attacks. Byzantine fault arises due to the distributed nature of untrusted nodes in the network, as there is no centralized authority to authenticate the validity of data and communication between the nodes. Sybil attack also arises as an adversary node can create multiple zombie nodes which act as participants in the network in order to control the majority of the network to manipulate the data in the blockchain. In order to verify the authenticity of network and address the Byzantine fault and Sybil attack issues, a set of rules to process incoming transactions, called "Consensus Mechanism" is used. Nakamoto consensus is mainly used in major blockchain implementations like Bitcoin and Ethereum which include the longest chain selection algorithm and Proof-of-Work (PoW) consensus algorithm, as shown in figure 3.

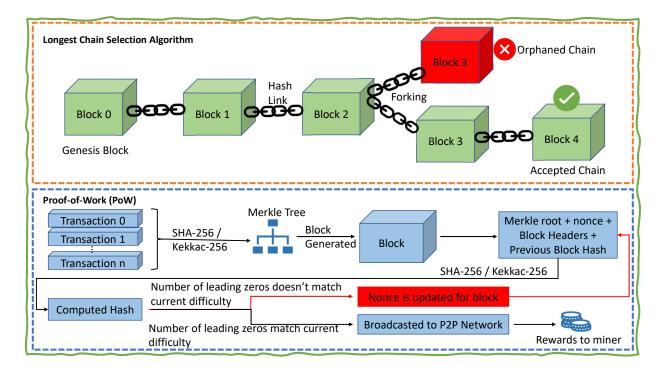


Fig. 3. Nakamoto Consensus for Avoiding Byzantine Fault and Sybil Attack.

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As the transactions are gathered and processed by distributed nodes, there is a chance of two miner nodes generating two blocks at the same time with different transactions. This will cause side-chains, which is called forking. In order to avoid this, the longest chain selection algorithm is implemented as part of the consensus mechanism. In this algorithm, the longest chain is elected for the next block to be added, ignoring the short side-chains leading to orphaned blocks but maintaining blockchain data consistency across the distributed nodes. Proof-of-Work (PoW) is part of the consensus mechanism which is designed to prevent Sybil attacks. In this algorithm, a hashcash problem is created [25] which requires significant computational power to solve but the solution is easy to verify. All the nodes will compete to solve this computationally hard problem, and whoever solves it first will be elected and is responsible for generating the new block in the blockchain. To engage and encourage the nodes to perform these mining processes, a reward and transaction fees are allocated to the winning node who is adding a new block.

2) Smart Contract Overview

The blockchain as such is built as a financial solution which keeps track of digital assets and prevents double spending. As discussed in the generations of blockchain, after identifying the usefulness of the blockchain in other fields, and in order to implement the business logic and make the blockchain relevant for other applications, smart contracts were invented. Smart contracts can be defined as simple pieces of code which can be automatically executed based on whether certain conditions are met. These are building blocks in creating a Decentralized Application (DApp). Each contract has the capability to interact with multiple smart contracts under the hood and to work collectively to achieve specified functions. For example, in a smart contract monitoring environmental parameters, events are triggered in case of temperature and humidity changes. Based on the threshold values, notifications can be triggered from smart contracts, as shown in figure 4.

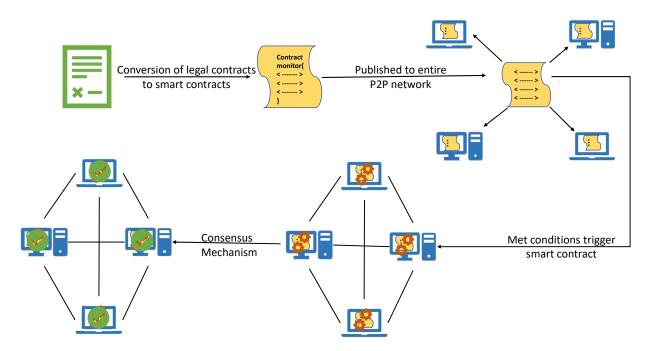


Fig. 4. Smart Contract Execution Steps.

Even though there are many advantages, there are also certain limitations for the usage of smart contracts in building DApps. Interacting with external world components to acquire data is a major problem. Smart contracts by design cannot perform HTTP requests to get real-time data. This is done to avoid the inconsistent values retrieved by the nodes which may lead the network to be unable to achieve consensus. This is one of the major problems for practical usage of smart contracts in real-world applications. For example, considering an insurance company with automating payouts to farmers based on loss due to weather conditions using smart contracts, all the nodes should receive the same data in order to achieve the consensus. Instead, as weather conditions change continuously, there will be inconsistencies to the data being seen by different nodes and the whole network will be jeopardized. To solve this problem "Oracles" are used in the proposed system for PSC.

The proposed PharmaChain makes use of DLTs combined with IoT technologies to address issues in the pharmaceutical industry. The rest of the paper is organized as follows: Section III presents the problems addressed in conventional enterprise resource planning systems which are centralized and novel contributions of the paper to address these problems. Section II discusses related research regarding proposed and existing DLT applications in the PSC. Section IV presents an architectural overview and the algorithmic foundation of the proposed systems. Section V includes the design implementation details of the proposed system. Section VI concludes the paper and presents possible future directions for research.

II. Related Prior Research

With very rapidly changing business needs and supply chain environments in the pharmaceutical industry, manufacturers have to redefine the supply chain in order to accommodate delivery of high quality products with low response times and high customer satisfaction. This can be achieved by incorporating multiple different technologies into the supply chain management (SCM). Regulations such as the Drug Supply Chain Security Act (DSCSA) require all entities participating in pharmaceutical supply chain to adopt technologies for reliable and secure transactions at each stage of the supply chain. This section critically analyzes the existing proposed solutions using blockchain as solution to build efficient supply chains to eliminate counterfeits and to provide the right amount of medicine to the right place, at the right time [26].

Crypto Pharmacy, proposed in [27], makes use of the New Economy Movement (NEM) blockchain in which Proof-of-Importance (POI) is implemented. An iOS application using Swift is built and integrated to the NEM network to make purchases using the XEM currency, which is the native coin for the NEM network. Modum.io AG, an application based on blockchain technology is proposed in [28]. It leverages smart contracts and the IoT to keep track of the medicines throughout the supply chain to ensure safe handling and delivery to consumer. This proposed architecture depends on the Ethereum platform and Proof-of-Work (PoW) consensus mechanisms. Environmental data from medicine lots is sensed using IoT sensors. The proposed architecture is robust and secure but the throughput of the network is low, as the Ethereum blockchain has bottlenecks at the current consensus mechanism. Taking into account that there will be a very large number of shipments which will be released into the supply chain of the huge pharmaceutical market, this will be a limiting factor along with the cost of operation. Another blockchain architecture proposed in [29] makes use of the Public Key Infrastructure (PKI) system to ensure the details are shared securely between manufacturer and consumer. As per the proposed architecture, every consumer who needs to access the details from the manufacturer has to share the public key, and upon approval from the manufacturer a QR code with information is encrypted using the requester public key and shared in the blockchain network, from where the intended consumer will be able to access details. It appears sufficient for secure sharing of information between two intended parties; however the main goal is to make the pharmaceutical supply chain transparent which will make information about drugs available to all participants in the network. Along with this problem there will be a large overhead for manufacturers as they have to approve every request from every customer who wants to access the information.

Drugledger, proposed in [30], makes use of an architecture similar to Bitcoin by assigning the drugs leaving or entering into a warehouse to Unspent Transactions (UTXO), thereby extending the tracking in multiple phases of packaging, distributing etc. Even though the Bitcoin network provides high security of data, it has a bottleneck for consensus mechanism which will be a major limiting factor in high transaction environment like PSC [28]. The blockchain architecture proposed in [31] is used for detecting the counterfeiting of drugs by using the Hyperledger Fabric platform to develop a private permissioned blockchain along with some chain-codes to implement business logic behind it.

A cloud based purchase management in the health sector model which makes use of IoT technology combined with blockchain implemented in Hyperledger, called "Composer" is proposed in [32], In this proposed architecture both ambient temperature and location were tracked throughout the supply chain. An Ethereum and smart contract based counterfeit tracking system for medicines is proposed in [33], however the scalability of the Ethereum platform is significantly reduced due the network congestion and consensus mechanism. Along with that, the cost of operations in the main network has increased significantly. Another such implementation which is also based on Ethereum platform and smart contracts can be seen in [34]. Ethereum is also used in [35] along with a proposed shipping container which can keep track of multiple ambient parameters for ensuring drug safety.

III. Novel Contributions

Presented here are the problems in current day Enterprise Resource Planning Systems which are considered for solution in the proposed-DLT based PSC.

A. Problems Addressed in the Current Paper

- Process delays in order management and shipment due to processing gaps between distributed systems.
- Information fragmentation and delay in auditing the problem orders in the traditional PSC.
- Existence of blind parties in traditional PSC causing delays and inconsistencies, along with difficulty in debugging issues with orders.
- The chance of entities participating in the network to include counterfeit drugs is very high and tracing of such entities is difficult.
- Safe recalling of drugs and ensuring safety at the point of administering the drug is another problem.
- Interacting with environmental parameter data is not very efficient and the chance of errors during recording data is high.

B. Novel Solutions Proposed in PharmaChain

- A distributed ledger based solution is proposed in the current PharmaChain implementation, which will ensure the faster processing of orders by avoiding communication gaps between entities in the PSC.
- The proposed distributed ledger will act as a single source of truth, accessible by all entities and addresses the information fragmentation problem.
- The proposed PharmaChain is a transparent supply chain where all the entities are aware of all the transactions between entities, thereby removing the blind parties which can equip them to take prompt actions.
- Counterfeiting can be removed as the entire trail of the drugs from manufacturing to the Point Of Sale (POS) can be tracked with non-repudiable ledger entries.
- As the transactions in the proposed PharmaChain are non-repudiable, accountability increases in the overall system, which can help in taking action against particular entities with malicious intent.
- Recalling of drugs will be an economical and easy process as there is a transparent drug traversing path in the supply chain, which will also ensure safe administering of these drugs.
- An automated process with IoT technology is implemented in PharmaChain which will help in extracting and communicating environmental parameter data with higher accuracy and which is less prone to human-errors.
- In order to solve the issue of smart contracts to interact with real world data, a process based on oracles is proposed in the current implemented PharmaChain.
- Automated warning systems can be built to notify entities about the status of the product in the supply chain.

IV. Overview of Proposed PharmaChain

An overview of the proposed system for PharmaChain is presented in this section. The whole proposed system can be divided into three logical components. The first component is API generation, which is responsible for integrating physical real-world data using sensors and creating an accessible API which will feed data into the next oracle component. For this, a cloud server-less configuration is used which will remove all the complex cloud infrastructure required and will be built by utilizing simple cloud functions. The sensing node proposed in the current implementation monitors minimal parameters like temperature and humidity along with GPS coordinates of the order in the supply chain. Data from the sensing node will be published to a topic using the Message Queuing Telemetry Transport (MOTT) lightweight publish-subscribe protocol. Whenever new sensing data is published on a topic, data will be consumed and IoT cloud custom rules are applied on the data to filter data with significance like temperature abnormalities. This filtered data is stored in a NoSQL database for auditing purposes and also a notification will be sent to the authorized authorities for taking prompt action to control environmental parameters of the shipment. Functions are used for transforming the JSON formatted data received from the MQTT topic to a dynamo DB compatible table item. Database logging will act as an event for another function which is going to make the data available for the API gateway end-point from which it is consumed by the oracle nodes while the hybrid smart contract is executed. The first component is shown in figure 5 and the interaction of different blocks in the proposed data source mechanism can be seen in Figure 6.

Oracles are third-party information services which act as data sources to provide real-time external data to smart contracts as they are not capable of interacting with real world systems and perform Application Programming Interface (API) calls. In distributed systems, multiple distributed nodes exist which are responsible for the working

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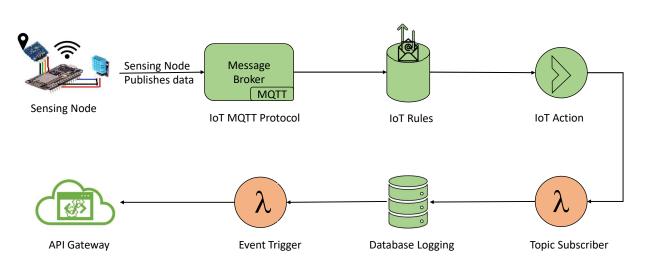


Fig. 5. Real-time Data Source for Oracle Component from Sensing Node in the Proposed PharmaChain.

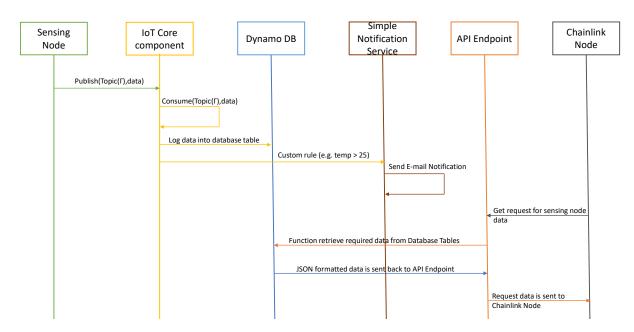


Fig. 6. Data Provider and Alert Notification Implementation using Sensing Node Data.

of the DApp. Some of these nodes are separated geographically and have no centralized authorities and are governed by a set of rules accepted across the network, called consensus mechanism. By calling the API directly in smart contracts, there is a chance for some nodes to receive the latest data but some not, because of the independent nature of these nodes. As there will be a discrepancy of data received from the API, this will cause the network to not reach consensus thus leading to the failure of the application. In order to avoid such situations, smart contracts are intentionally made not incapable of fetching data from external data sources. This will limit the interaction of smart contracts to real-world data thereby limiting the use-cases. Hence, decentralized data providers are designed to work with smart contracts and are named oracles. These oracles are classified into different types based on information sources, degree of decentralization and information flow direction. Based on the information source there are software oracles and hardware oracles. Software oracles mainly depend on the information gathered from online sources like websites, servers and other network databases. Such oracles can be mainly used in applications including the supply chain. A similar approach is followed in the proposed PharmaChain mechanism but the data to the online sources from hardware components like sensors, bar-code scanners, etc. Such oracles are perfect for designing systems like supply chains as they involve many electronic components to track and trace products throughout the supply-line. Another classification is based on the direction of the information flow with respective to smart contracts. Oracles providing data into the smart contracts by fetching data from external sources are called inbound oracles. On the other hand, oracles which are providing data to the external environment by fetching them from smart contracts are called outbound oracles. An inbound oracle to provide sensor data from sensing nodes to smart contracts is used in the proposed PharmaChain application. Based on the degree of decentralization, oracles can be classified as centralized oracles and decentralized oracles. If the data source is a single central entity, then it is called centralized oracle. On the other hand, if data providers are decentralized, it is called decentralized oracle. Centralized oracles are less effective and secure compared to decentralized oracles. The steps executed during the oracle interaction are shown below:

- A hybrid client smart contract which requires access to external data from any API will prepare a request with parameters job id (which depends on the type of data being retrieved from the API), destination address for the data, and the fulfillment function which needs to be executed after the data is fetched from the API.
- In next step, the oracle contract is going to publish an event with the given job id and will set parameter values along with the fee set by the client smart contract.
- All the nodes attached to the network will be notified by the event generated before with the requested job details.
- The node which has the job described will make use of all the parameters set by the client contract and execution is done.
- Once the execution is completed, the node which executed the job is going to send back the fulfillment results to the oracle smart contract.
- The oracle smart contract will then execute the fulfillment method and the desired data is accessed in the client smart contract.

In a decentralized oracle structure, multiple jobs are run instead of a single job and the result is aggregated to get a more reliable data source and making it more decentralized. An overview of the proposed blockchain architecture is shown in figure 7.

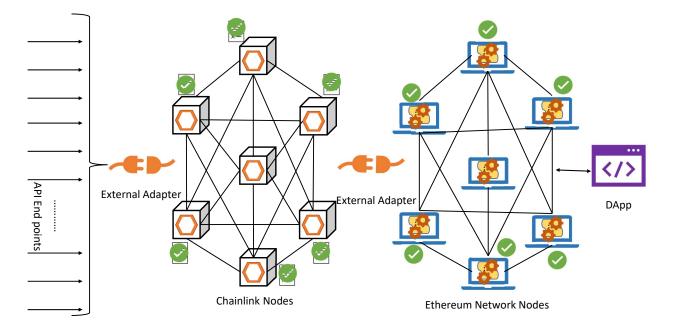


Fig. 7. Proposed Blockchain Architecture Components for PharmaChain.

The third logical component in the proposed system is the hybrid smart contract which is responsible for executing different activities in the supply chain based on an event. The result of the smart contract execution is made available to all participants in the network, thus producing a transparent ledger. Based on the proposed application, different entities have different roles and responsibilities to perform when a pharmaceutical shipment is moving through the supply chain. Hence, Role Based Access Control (RBAC) is proposed in the current system in which

the entity is assigned with roles and the corresponding responsibilities by smart contracts. For simplicity, fewer entities are considered while implementations and back orders/returns or payments are not considered. An entity with a manufacturer role has the activities of producing/manufacturing drugs, and selling manufactured drugs to the wholesaler/distributor. The distributor role has the activities of packaging and selling the product to the retailer. The retailer entity has the activity of selling packaged products to the consumer which is the end of the supply chain. The activity diagram is shown in figure 8. Role management is implemented in smart contracts by using function modifiers which will restrict the behavior of smart contract functions. These modifier conditions are checked prior to the function being executed. Different modifiers such as onlyManufacturer, onlyDistributor, onlyRetailer, onlyConsumer, etc., are defined to restrict the access of functions based on the role assigned to the user.

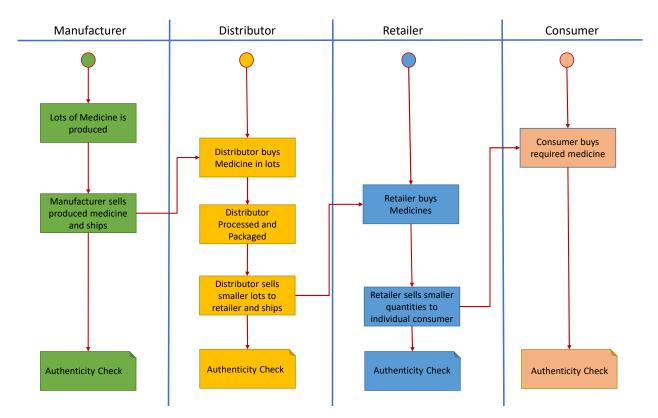


Fig. 8. Pharmaceutical Supply Chain Entity Activity Diagram considered in Implemented PharmaChain Solution.

V. Implementation and Validation of Proposed PharmaChain

A. Implementation of PharmaChain using Off-Shelf Components

A simple prototype of the proposed PharmaChain system is implemented using a single sensing node which is going to track one shipment from end-to-end in the supply chain. The sensing node is implemented using a NodeMCU ESP8266 module which is small form factor IoT platform with Wi-Fi capability and can integrate data from sensors to an IoT network easily through a wireless connection. For sensing environmental parameters around the shipment, a DHT11 sensor is used which is going to sense both ambient temperature and relative humidity around the shipment. A GPS Module GPS NEO-6M is used for keeping tracking of the shipment and sends latitude and longitude positions of the shipment during different stages in the supply chain. The implemented sensing node is shown in figure 9 and data published from the implemented sensing node is shown in figure 10. A secure communication channel is created by using RSA encryption keys to communicate between the sensing node and the cloud. The private key, and root CA certificate required to establish secure connection are loaded into the device's SPI Flash File System (SPIFFS) of NodeMCU. The WiFiClient library available for ESP8266 NodeMCU is used for loading the required security certificates before establishing the cloud connection. Once the secure connection is established, temperature and humidity data from the DHT11 sensor, GPS coordinates from the GPS module and other shipment information is populated into a JSON string using the ArduinoJson. This generated

JSON data is converted to a char array and published to a topic using the PubSubClient library. The sensing node algorithm is shown in algorithm 1.

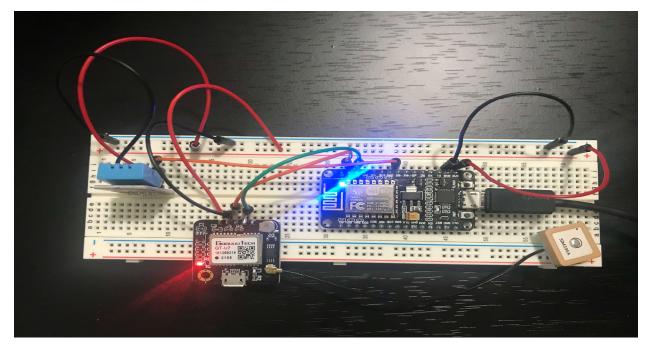


Fig. 9. Implemented Sensing Node for Proposed PharmaChain Application.

Data from the published topic is consumed by the cloud infrastructure which is going to process the received data based on a set of predefined rules and acts as data source for the oracle component. During the implementation of the prototype for an over the counter (OTC) available drug, for which storage temperature is less than 25° C, is considered as an example. Once the data is consumed, it is checked for any temperature abnormalities (i.e., in this case if Temperature > 25° C). If the temperature from the sensing node satisfies the rule, the data will be logged into a database for auditing and an immediate notification via e-mail is sent to the registered parties. The writing of data to the database and sending notifications is achieved by using cloud functions. Along with the abnormalities, the latest data is updated into another database table from which the data will be fetched when oracles make an API request. For fulfilling API requests, an API Gateway with required methods and path parameters is designed. As the data is only retrieved from the DB tables, an HTTP GET method, which will retrieve shipment details from the DB tables and functions can be seen in figure 11.

The proposed blockchain architecture is implemented using Chainlink for oracle services and Ethereum as the blockchain platform to build the smart contracts. Chainlink is an oracle service which will enable Web 3.0 applications to access data from existing real-world data sources [36]. This will help in developing hybrid smart contracts which can interact with real-world systems and to provide real-world solutions. Ethereum is an open-source blockchain platform and also provides a Turing complete language (Solidity) to build decentralized applications. The programming environment provided by Ethereum works based on the Ethereum Virtual Machine (EVM) which is a world computer whose state is maintained and agreed upon by all the participant nodes in the P2P network. Smart contracts are executed within the EVM and these executions will move state of the EVM from one finite state to another. All the nodes in the network will record and store the state changes of the EVM to form a ledger of transactions, thus forming the blockchain. Ethereum currently works on a Proof-of-Work (PoW) consensus mechanism where the miner is chosen by posing a computationally hard hash problem to miner nodes.

For building the prototype of the proposed PharmaChain, the KOVAN test network provided by Ethereum is used which works on Proof-of-Authority (PoA). Test networks are P2P networks which will simulate the main network of Ethereum in execution but utilize test tokens for all the payments and gas fees. Chainlink also provides different jobs in the KOVAN testnet which can be executed by utilizing a test LINK token instead of spending real money for deploying and testing decentralized applications. Smart contracts developed for PharmaChain can be logically divided into three components: smart contracts for implementing Role Based Access Control (RBAC), smart contracts

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	titude":33.21297,"longitude":-97.1575,"sku":2112101,"lot":547863250,"drugname":"Mucinex","temperature":21.5,"humidity":61}
23592	
	titude":33.21295,"longitude":-97.15753,"sku":2112101,"lot":547863250,"drugname":"Mucinex","temperature":21.4,"humidity":61}
23592	
	titude":33.21297,"longitude":-97.15757,"sku":2112101,"lot":547863250,"drugname":"Mucinex","temperature":21.4,"humidity":61}
23592	
	titude":33.21296,"longitude":-97.15765,"sku":2112101,"lot":547863250,"drugname":"Mucinex","temperature":21.4,"humidity":61}
23592	titude": 33.21296. "longitude": -97.15772. "sku": 2112101. "lot": 547863250. "drugname": "Mucinex". "temperature": 21.4. "humidity": 611
23592 sh message: {"timestamp":"11 / 01 / 2022 04 : 28 : 02 AM ","1at	titude":33.21296,"longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex","temperature":21.4,"humidity":61}
23592 sh message: {"timestamp":"ll / 01 / 2022 04 : 28 : 02 AM ","lat 23592	titude":33.21296,"longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex","temperature":21.4,"humidity":61}
: 23592 ish message: {"timestamp":"11 / 01 / 2022 04 : 28 : 02 AM ","1at : 23592 ish message: {"timestamp":"11 / 01 / 2022 04 : 28 : 02 AM ","1at	<pre>titude":33.21296,"longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex", "temperature":21.4, "humidity":61} titude":33.21296,"longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex", "temperature":21.4, "humidity":61}</pre>
23592 sh message: {"timestamp":"11 / 01 / 2022 04 : 28 : 02 AM ","lat 23592 sh message: {"timestamp":"11 / 01 / 2022 04 : 28 : 02 AM ","lat 23592	<pre>titude":33.21296,"longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex", "temperature":21.4, "humidity":61} titude":33.21296,"longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex", "temperature":21.4, "humidity":61} titude":33.21296, "longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex", "temperature":21.4, "humidity":61} titude":33.21296,"longitude:-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex", "temperature":21.4, "humidity":61}</pre>
22552 ib messages ("timestamp":"11 / 01 / 2022 04 : 28 : 02 AM ","lat 23552 ib messages ("timestamp":"11 / 01 / 2022 04 : 28 : 02 AM ","lat 23552 ib messages ("timestamp":"11 / 01 / 2022 04 : 28 : 28 AM ","lat	<pre>titude":33.21296,"longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex", "temperature":21.4, "humidity":61} titude":33.21296,"longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex", "temperature":21.4, "humidity":61} titude":33.21297,"longitude":-97.1575,"sku":2112101,"lot":547863250,"drugname":"Mucinex", "temperature":21.4, "humidity":61}</pre>
: 23592 isi message: {"timestamp":"11 / 01 / 2022 04 : 28 : 02 AM ","1at : 23552 isi message: {"timestamp":"11 / 01 / 2022 04 : 28 : 02 AM ","1at : 23552 isi message: {"timestamp":"11 / 01 / 2022 04 : 28 : 28 AM ","1at : 23552	<pre>titude":33.21296,"longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex","temperature":21.4,"humidity":61} titude":33.21296,"longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex","temperature":21.4,"humidity":61} titude":33.21297,"longitude":-97.1575, "sku":2112101,"lot":547863250,"drugname":"Mucinex","temperature":21.4,"humidity":61} titude":33.21297,"longitude":-97.1575, "sku":2112101,"lot":547863250,"drugname":"Mucinex","temperature":21.4,"humidity":61}</pre>
: 23592 1: bin message: ("timestamp":"11 / 01 / 2022 04 : 28 : 02 AM ","1at : 23592 1: bin message: ("timestamp":"11 / 01 / 2022 04 : 28 : 02 AM ","1at : 23592 1: bin message: ("timestamp":"11 / 01 / 2022 04 : 28 : 28 AM ","1at : 23592 1: bin message: ("timestamp":"11 / 01 / 2022 04 : 28 : 35 AM ","1at	<pre>titude":33.21296,"longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex", "temperature":21.4, "humidity":61} titude":33.21296,"longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex", "temperature":21.4, "humidity":61} titude":33.21297,"longitude":-97.1576,"sku":2112101,"lot":547863250,"drugname":"Mucinex", "temperature":21.4, "humidity":61} titude":33.21297,"longitude":-97.15767,"sku":2112101,"lot":547863250,"drugname":"Mucinex", "temperature":21.4, "humidity":61}</pre>
: 23592 ish message: ["timestamp":"11 / 01 / 2022 04 : 28 : 02 AM ","lat : 23592 ish message: ["timestamp":"11 / 01 / 2022 04 : 28 : 02 AM ","lat : 23592 ish message: ["timestamp":"11 / 01 / 2022 04 : 28 : 28 AM ","lat : 23592 ish message: ["timestamp":"11 / 01 / 2022 04 : 28 : 35 AM ","lat : 23592	titude":33.21296,"longitude":-97.15772,"sku":2112101,"lot":547863250,"drugname":"Mucinex","temperature":21.4,"humidity":61) titude":33.21297,"longitude":-97.1575,"sku":2112101,"lot":547863250,"drugname":"Mucinex","temperature":21.4,"humidity":61)

Fig. 10. Data Published from Implemented Sensing Node for PharmaChain.

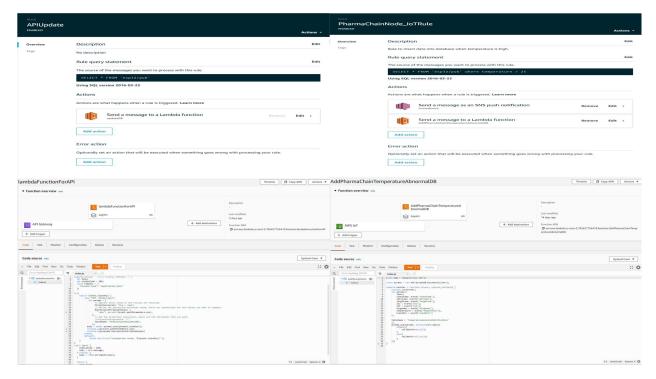


Fig. 11. Implemented Data Source Rules and Functions in Proposed PharmaChain.

	gorithm 1: Algorithm for Sensing Node in Proposed PharmaChain Implementation.	
Ou / *	<pre>put: Certificate and Keys for IoT thing, Ambient temperature (temp) and relative humidity (hum) from DHT11 Sensor, GPS latitude (lat) and longitude (lng) co-ordinates from GPS Module tput: Boolean status of message published to topic During Setup sensing node tries to setup wifi and MQTT connection and establish secure channel to communicate between sensing node and cloud ad required libraries for WiFi Connection, MQTT Communication, JSON for data manipulations</pre>	*/
2 wh 3 _	<pre>/* Sensing node tries to connect to Wi-Fi network using given SSID and PASSWORD strings if it not connected to network ile WiFi.status() == FALSE do WiFi.begin(ssid, password)</pre>	*/
	SPI Flash File Systems is mounted successfully	*/
4 wh 5 6 7 8	<pre>ile SPIFFS.status() == FALSE do SPIFFS.begin() /* Open Certificate and Key files in flash memory of sensing node SPIFFS.open("Root CA certificate(CA_{cert})") SPIFFS.open("Public Key (PuK)") SPIFFS.open("Private Key (PrK)")</pre>	*/
9 10 11	/* Load certificate and key files esp8266Client.loadCertificate(PuK) esp8266Client.loadPrivateKey(PrK) esp8266Client.loadCACert(CA _{cert})	*/
	Attempting MQTT connection	*/
12 WN	<pre>ile pubClient.connected() == FALSE do /* Trying to reconnect</pre>	*/
13	pubClient.connect()	
	Environmental and Geolocation inputs from sensors along with shipment overhead information is processed every 60 seconds <i>every 60 seconds</i> do /* JSON String is generetd including all environmental and geolocation data from	*/
	sensing node	*/
15	Publishing message MSG \leftarrow JSON(timestamp,lat,lng,sku,lot,drugName,temp,hum)	,
16	/* Data to be published is converted to char array $MSG+ \leftarrow JSON.toCharArray(MSG)$	*/
17	/* Converted MSG+ is published to topic τ_{pub} publicent.publish(τ_{pub} ,MSG+)	*/

to work with the Chainlink component and smart contracts for the core pharmaceutical supply chain mechanism. Smart contracts for RBAC will define different roles and will associate them with different Ethereum addresses. Responsibilities for each role are defined and controlled by defining modifiers in the Solidity language. Modifiers can also be defined to check some pre-defined conditions in the blockchain before executing the transactions submitted by a user. Before looking into defined modifiers, a shipment within the chain assumes different states which are enumerated as shown in Table I. Below are the smart contract functions and modifiers defined for RBAC in the implemented PharmaChain prototype.

function addManufacturer(address account) public onlyManufacturer: This function will add a new address to the list of manufacturers. It takes the address of the manufacturer as parameter and checks if the given address is not already associated with the role Manufacturer. If not associated, the address will be added to the list of manufacturers.

function renounceManufacturer() public: This function helps in removing the access privileges of a manufacturer. The address from which the transaction is generated (msg.sender) will be checked for manufacturer privileges; if assigned, the role will be revoked and the address will be removed from list of manufacturers.

function isManufacturer(address account) public view returns (bool): This is a view function defined to check if the given address has the manufacturer role assigned. If assigned, it will return true, and false otherwise.

Ā	lgorithm 2: Algorithm for Data Source API Generation in Proposed PharmChain Implementation.	
Īı	nput: Input JSON message from Sensing Node (MSG+), HTTP GET API Request from Oracle	
0	Output: Requested Shipment Data	
1	* Data published MSG+ on topic $ au_{pub}$ from previous algorithm is consumed by the IoT	2
	cloud component	*/
1 if	τ_{pub} .hasMessage() then	
2	$MSG \leftarrow IoTCore.consume(\tau_{pub}, MSG+)$	
	/* In the example drug taken temperature threshold is $25^{\circ}C$	*/
3	if <i>MSG.temperature</i> > 25) then	
	/* Update the audit table for temeprature abnormalities recorded during the	
	supply chain process AbnormalTempAuditTable.update(MSG)	*/
4	<pre>/* An E-mail notitification will be sent to the registered parties in case t</pre>	ho
	temperature exceeds the threshold	*/
5	SNSNotification.sendEmail(MSG)	^/
6	<pre>/* Update this table with latest information of the shipment ShipmentLatestUpdateTable.update(MSG)</pre>	*/
1		
7 W	hile HTTP GET Request (req) do	,
_	/* get request path parameters to geenrate the DB query	*/
8	pathParameter param = req.getParam()	
	<pre>/* Scan the DB Table to retreive requested shipment information and store resul */</pre>	t
9	result = ShipmentLatestUpdateTable.scan(filtering param)	
9	/* result from scan is mapped to http body	*/
10	http://www.http://www.http://www.http://www.http.mapBody(result)	*/
10	/* A http response is generated and sent back to the requested oracle	*/
11	httpResponse (resp) \leftarrow http.generateResponse(httpHeaders,httpBody)	27
12	http.ResponseSend(resp)	

function addDistributor(address account) public onlyDistributor: This function will add a new address to the list of Distributors. It takes the address of the distributor as parameter and checks if the given address is not already associated with the role distributor. If not associated, the address will be added to the list of distributors.

function renounceDistributor() public: This function helps in removing the access privileges of the distributor. The address from which the transaction is generated (msg.sender) will be checked for distributor privileges; if assigned, the role will be revoked and the address will be removed from list of distributors.

function isDistributor(address account) public view returns (bool): This is a view function defined to check if the given address has a distributor role assigned. If assigned, it will return true, and false otherwise.

function addRetailer(**address account**) **public onlyRetailer**: This function will add a new address to the list of retailers. It takes the address of the retailer as parameter and checks if the given address is not already associated with the role retailer. If not associated, the address will be added to the list of retailers.

function renounceRetailer() public: This function helps in removing the access privileges of the retailer. The address from which the transaction is generated (msg.sender) will be checked for retailer privileges; if assigned, the role will be revoked and the address will be removed from list of retailers.

function isRetailer(address account) public view returns (bool): This is a view function defined to check if the given address has retailer role assigned. If assigned, it will return true, and false otherwise.

function addConsumer(address account) public onlyConsumer: This function will add a new address to the list of consumers. It takes the address of the consumer as parameter and checks if the given address is not already associated with role consumer. If not associated, the address will be added to the list of consumers.

function renounceConsumer() public: This function helps in removing the access privileges of the consumer. The address from which the transaction is generated (msg.sender) will be checked for consumer privileges; if assigned the role will be revoked and the address will be removed from list of consumers.

function isConsumer(address account) public view returns (bool): This is a view function defined to check if the given address has consumer role assigned. If assigned, it will return true, and false otherwise.

modifier onlyOwner(): This modifier is defined to check the ownership of the contract. This function checks if the sender of the transaction (msg.sender) is the same as the one who deployed this contract and can be used to control transfer of ownership of contract from one user to another.

modifier verifyCaller(address _address): This modifier is used to compare the sender of a transaction (msg.sender) to a given Ethereum address parameter and returns a boolean true if they match, and false otherwise.

modifier producedByManufacturer(uint _upc): This modifier will take the Universal Product Code (UPC) of a shipment and will check the state of the product in the supply chain. If the state of the product is '0', as shown in Table I, it will return true and false otherwise.

modifier updateInventoryByManufacturer(uint _upc): This modifier will take the UPC as parameter and checks if the state of the manufactured product is moved to Update Inventory By Manufacturer. If the state matches '1', it returns true and false otherwise.

modifier purchasedByDistributor (uint _upc): This modifier will take the UPC of a shipment and will check the state of the product in the supply chain. If the state of the product is '2', as shown in Table I, it will return true and false otherwise.

modifier shippedByManufacturer (uint _upc): This modifier will take the UPC of a shipment and will check the state of the product in the supply chain. If the state of the product is '3', as shown in Table I, it will return true and false otherwise.

modifier receivedByDistributor (uint _upc): This modifier will take the UPC of a shipment and will check the state of the product in the supply chain. If the state of the product is '4', as shown in Table I, it will return true and false otherwise.

modifier processByDistributor (uint _upc): This modifier will take the UPC of a shipment and will check the state of the product in the supply chain. If the state of the product is '5', as shown in Table I, it will return true and false otherwise.

modifier packagedByDistributor (uint _upc): This modifier will take the UPC of a shipment and will check the state of the product in the supply chain. If the state of the product is '6', as shown in Table I, it will return true and false otherwise.

modifier forSaleByDistributor (uint _upc): This modifier will take the UPC of a shipment and will check the state of the product in the supply chain. If the state of the product is '7', as shown in Table I, it will return true and false otherwise.

modifier shippedByDistributor (uint _upc): This modifier will take the UPC of a shipment and will check the state of the product in the supply chain. If the state of the product is '8', as shown in Table I, it will return true and false otherwise.

modifier purchasedByRetailer (uint _upc): This modifier will take the UPC of a shipment and will check the state of the product in the supply chain. If the state of the product is '9', as shown in Table I, it will return true and false otherwise.

modifier receivedByRetailer (uint _upc): This modifier will take the UPC of a shipment and will check the state of the product in the supply chain. If the state of the product is '10', as shown in Table I, it will return true and false otherwise.

modifier forSaleByRetailer (uint _upc): This modifier will take the UPC of a shipment and will check the state of the product in the supply chain. If the state of the product is '11', as shown in Table I, it will return true and false otherwise.

modifier purchasedByConsumer (uint _upc): This modifier will take the UPC of a shipment and will check the state of the product in the supply chain. If the state of the product is '12', as shown in Table I, it will return true and false otherwise.

modifier onlyManufacturer(): This modifier is used to check if the sender of a transaction is associated with the role manufacturer. The function isManufacturer(address account) is called with address parameter msg.sender and a boolean result true is returned if the role is assigned or false otherwise.

modifier onlyDistributor(): Similar to the previous function isDistributor(address account) is called with address parameter from msg.sender and true is returned if the role is assigned or false otherwise.

modifier onlyRetailer(): This modifier is used to check if the sender of a transaction is associated with the role retailer. The function isRetailer(address account) is called with address parameter msg.sender and a boolean result true is returned if the role is assigned or false otherwise.

modifier onlyConsumer(): This modifier is used to check if the sender of transaction is associated with the role consumer. The function isConsumer(address account) is called with address parameter msg.sender and a boolean result true is returned if the role is assigned or false otherwise.

Pharmaceutical Shipment State	Description	Enumerated State Value
Produced By Manufacturer	Pharmaceutical Manufacturer acquires necessary	0
	RAW materials and drugs are manufactured	
Update Inventory By Manufacturer	Inventory is updated by manufactured and offered	1
	for sale	
Purchased By Distributor	Pharmaceutical Distributor purchases from inventory	2
	of manufacturer based on market demand	
Shipped By Manufacturer	After successful purchase product is shipped from	3
	manufacturer to distributor	
Received By Distributor	Purchased product is received by Distributor	4
Processed By Distributor	Inventory update at distributor	5
Packaged By Distributor	Processed product is sometimes re-packaged or	6
	branded by distributor	
For Sale By Distributor	Inventory is set up for sale by the distributor	7
Purchased By Retailer	Product is purchased by In-mail/ Brick and Mortar	8
	retailer pharmacies	
Shipped By Distributor	Once purchase is successful, product is moved from	9
	distributor to retailer locations	
Received By Retailer	Product is received by retailer pharmacies	10
For Sale By Retailer	Inventory update and offer it for sale either In-	11
	mail/physical pharmacy	
Purchased By Consumer	Consumer purchases required medication from pham-	12
	racies	

TABLE I	
ASSUMED SHIPMENT STATES IN PROPOSED PHARMACHAIN AND CORRESPON	DING ENUMERATION.

Functions implemented for pharmaceutical supply chain are discussed below and their associated access control modifiers can be seen in Table II.

requestTemperatureData(**string memory _sku**): This function helps to interact with the Chainlink oracle to retrieve the current ambient temperature around the shipment from the sensing node. It takes a SKU as a parameter and this value will be send to API Get request as path parameter, based on which the cloud DB tables will be queried to get the latest temperature value. This function can be called by any account with any role or no role attached.

requestHumidityData(**string memory _sku**): This function helps to interact with the Chainlink oracle to retrieve the current relative humidity around the shipment from the sensing node. It takes a SKU as a parameter and this value will be send to API Get request as path parameter, based on which the cloud DB tables will be queried to get the latest temperature value. This function can be called by any account with any role or no role attached.

requestLatitude(string memory _sku): This function helps to interact with the Chainlink oracle to retrieve the current latitude of the shipment from the sensing node. It takes a SKU as a parameter and this value will be send to API Get request as path parameter, based on which the cloud DB tables will be queried to get the latest temperature value. This function can be called by any account with any role or no role attached.

requestLongitude(**string memory _sku**): This function helps to interact with the Chainlink oracle to retrieve the current longitude of the shipment from the sensing node. It takes a SKU as a parameter and this value will be send to API Get request as path parameter, based on which the cloud DB tables will be queried to get the latest temperature value. This function can be called by any account with any role or no role attached.

verifyAuthenticityoFProduct(uint _upc): This function will verify the origin of the product along with the product transfer between manufacturer, distributor, retailer and consumer to prove the authenticity of the product. Authenticity can be verified at any stage of the supply chain, hence can be called from any account with or without any role attached.

produceItemByManufacturer(string memory _sku, string memory _drugName, uint _upc): This function is executed by a manufacturer for each newly produced product. Each product with assigned UPC and SKU along with drug name will be sent as parameters to this function. This will create a new product in the PSC. This function can only be executed by accounts with role manufacturer and generates an event "ProducedByManufacturer" to be recorded in the blockchain.

sellItemByManufacturer(uint _upc): This function is executed by the manufacturer after producing new products. Before executing this function, it will check if the caller of the function is manufacturer and really owns the product to sell. Once verified it will change the state of the product to available for selling and generates an event to record in the blockchain.

purchaseItemByDistributor(uint _upc): This function helps distributors to buy products offered for sale by manufacturers. Any account with distributor role can call this function and once the purchase is done, the status of the product along with the ownership of the product will change from manufacturer to purchased distributor.

shippedItemByManufacturer(uint _upc): After successful purchase, the manufacturer is responsible for shipping the product and this function is used to update the status of product to shipped and can be called by the manufacturer who is assigned as origin manufacturer of the product. This generates an event shipped to be recorded in the blockchain.

receivedItemByDistributor(uint _upc): This function is used by a distributor to update the shipment once received from manufacturer. This function verifies if the caller is a distributor and actually purchased the product before updating the status of shipment to "received". This function will generate a received shipment event to be recorded in the blockchain.

processedItemByDistributor(uint _upc): Once shipment is received by a distributor and processed, this function is used to update the status of the shipment to processed. This function can only be executed by the distributor who owns the shipment.

packageItemByDistributor(uint _upc): Sometimes distributors re-package the product before offering to sell it. Once repackaging is done this function is called by the distributor to change the state of the product to be re-packaged and ready to be sold. This function ensures the distributor who actually owns the product can make the function call.

sellItemByDistributor(uint _upc): This function helps a distributor to offer the product owned for sale. Only the distributor who owns the product can make a call to this function and if the status of the product is already packaged by distributor. A sell event is generated by this function to be recorded in the blockchain.

purchaseItemByRetailer(uint _upc): This function helps any user with retailer role assigned to make a purchase from a distributor. Once purchase is done, the owner of the product will be changed to the purchased retailer and an event is generated to be recorded in the blockchain.

shippedItemByDistributor(uint _upc): After the purchase by a retailer, the shipment is shipped from distributor to the retailer. This event is updated by using this function and a ship event is generated to be recorded in the blockchain.

receivedItemByRetailer(uint _upc): Once the shipment is received from the distributor, the retailer will execute this function to record the shipment received event in the blockchain.

sellItemByRetailer(uint _upc): Once a retailer receives the shipment and processes it, it is offered for sale by using this function. This function, before executing, will ensure that the caller of the function is the retailer who actually owns the product. An event is generated to be recorded in the blockchain.

purchaseItemByConsumer(uint _upc): This is the final step in the supply chain where the consumer can make the purchase of the product from a retailer. Any user with consumer role attached can execute this function which will change the ownership of the product from retailer to consumer along with generating a purchase event to be recorded in blockchain.

The implemented PharmaChain application sequence of execution steps is shown in figure 12. During the first step the manufacturer creates a new product which has its own SKU and UPC assigned by the manufacturer. Once the product is available, the manufacturer will update the inventory and will offer the product for sale. This will make the product available for purchase by any distributor. A distributor purchases the product which will change

Pharmaceutical Sup-	Modifiers	Event Generated
ply Chain Smart Contract Function		
requestTemperatureData(string memory _sku)	Any	None
requestHumidityData(string mem- ory _sku)	Any	None
requestLatitude(string memory _sku)	Any	None
requestLongitude(string memory _sku)	Any	None
verifyAuthenticityoFProduct(uint _upc)	Any	None
produceItemByManufacturer(string memory _sku, string memory _drugName, uint _upc)	onlyManufacturer()	ProducedByManufacturer(uint upc)
sellItemByManufacturer(uint _upc)	onlyManufacturer() producedByManufacturer(_upc) verifyCaller(items[_upc].ownerID)	UpdateInventoryByManufacturer(uin upc)
purchaseItemByDistributor(uint _upc)	onlyDistributor() updateInventoryByManufac- turer(_upc)	PurchasedByDistributor(uint upc)
shippedItemByManufacturer(uint _upc)	onlyManufacturer() purchasedByDistributor(_upc) verify- Caller(items[_upc].originManufactur	ShippedByManufacturer(uint upc)
receivedItemByDistributor(uint _upc)	onlyDistributor() shippedByManufacturer(_upc) verifyCaller(items[_upc].ownerID)	ReceivedByDistributor(uint upc)
processedItemByDistributor(uint _upc)	onlyDistributor() receivedByDistributor(_upc) verifyCaller(items[_upc].ownerID)	ProcessedByDistributor(uint upc)
packageItemByDistributor(uint _upc)	onlyDistributor() processByDistributor(_upc) verifyCaller(items[_upc].ownerID)	PackagedByDistributor(uint upc)
sellItemByDistributor(uint _upc)	onlyDistributor() packagedByDistributor(_upc) verifyCaller(items[_upc].ownerID)	ForSaleByDistributor(uint upc)
purchaseItemByRetailer(uint _upc)	onlyRetailer() forSaleByDistributor(_upc)	PurchasedByRetailer(uint upc)
shippedItemByDistributor(uint _upc)	onlyDistributor() purchasedByRetailer(_upc) verify- Caller(items[_upc].distributorID)	ShippedByDistributor(uint upc)
receivedItemByRetailer(uint _upc)	onlyRetailer() shippedByDistributor(_upc) verifyCaller(items[_upc].ownerID)	ReceivedByRetailer(uint upc)
sellItemByRetailer(uint _upc)	onlyRetailer() receivedByRetailer(_upc) verifyCaller(items[_upc].ownerID)	ForSaleByRetailer(uint upc)
purchaseItemByConsumer(uint _upc)	onlyConsumer()	PurchasedByConsumer(uint upc)

 TABLE II

 PHARMACHAIN SMART CONTRACT FUNCTIONS AND ASSOCIATED ACCESS CONTROL MODIFIERS AND EVENT GENERATIONS.

the ownership of the product from manufacturer to the distributor. The manufacturer then ships the product to the purchasing distributor and updates the status. Once the distributor receives the product, processes and packages the product before selling it to the retailer. The retailer will then make the purchase and the ownership of product will change from the distributor to retailer. Once the retailer receives the product, will offer it for sale to the customer. In the final step, consumer will be able to purchase it and also check the authenticity of the product by verifying the ownership changes between entities.

Manufactur	red Drug	Manufa	cturer		Distri	butor		Reta	iler		Cons	umer
*	produceItemByManufa	cturer()										
4	sellItemByManufactu	ırer()										
			purchas	eltemByDistrib	utor()							
			shipped	temByManufa	cturer()							
			received	ItemByDistrib	utor()							
	processedItemByDis	stributor()										
4	• •		package	ItemByDistrib	utor()							
	sellItemByDistributo	or()										
						purchas	eltemByRetailer()				
						shipped	ltemByDistributo	r()				
						received	lltemByRetailer()					
			sellite	mByRetailer()								
	•								purcha	seltemByConsu	imer()	
						verifyA	uthenticity()					
	•											

Fig. 12. Sequence Diagram of Implemented PharmaChain Showing Interactions Between Entities Using Smart Contract Functions.

Parameters	Value
Deployed Contract Ad-	0x9409739e0D68Cb459a41892a48F3E62A2ceb7eeF
dress	
Contract Owner Ad-	0xBc6251e4f6389117a8f08b9B465a0bd44bA15Ab4
dress	
Manufacturer Address	0x3eDe97Ea0DFF3EcD7320b1822E33f4a2764E8ed4
Distributor Address	0xE8FDEa8272393Ad05f004d2BF583D784b509f9D3
Retailer Address	0x81F66b24db9EA43f454636750922c4ef12c26b48
Consumer Address	0xD46c558431c9CA642A85885509FBf6dA4Ba60435
Contract Creation	0x0c99850a32d2f836023a8ada0ec3feab2ae34be2d16969c2d38537e51e77db7a
Transaction Hash	
Oracle Address for Un-	0xc57B33452b4F7BB189bB5AfaE9cc4aBa1f7a4FD8
signed Integer data	
Job ID for Unsigned	d5270d1c311941d0b08bead21fea7747
Integer data	
Oracle Address for	0x9C0383DE842A3A0f403b0021F6F85756524d5599
Signed Integer data	
Job ID for Signed Inte-	ba1d5d5070a247eaa7070f838a42bb03
ger data	

 TABLE III

 Deployed PharmaChain Smart Contract Details.

The User Interface (UI) is implemented using the web3 library to interact with published smart contracts from the KOVAN test network. Metamask is used as wallet provider to operate and utilize test currency for deploying and

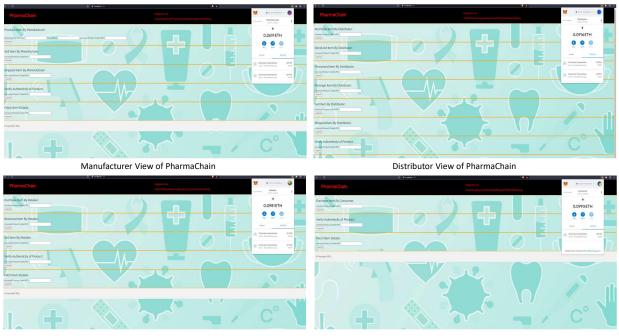
Action	Tx fees for	Tx hash	Tx fees for	Block
	Ethereum Blockchain		Chainlink Oracle	Time (s)
Contract Deploy-	0.00898 ETH	0x0c99850a32d2f836023a8ada0ec3feab2ae34be	0	8
ment		2d16969c2d38537e51e77db7a		
Add	0.00011 ETH	0xeae4963ac8b822d35f18bba2c20d5d01ee3ad4e	0	4
Manufacturer		b2451c6372c3d3f93e7b11b47		
Add Distributor	0.00011 ETH	0x23c223c177185e78e4e7ab90b8bcf217930ebc5	0	4
		7ebdcbce3431bfa366cd51bf9		
Add Retailer	0.00011 ETH	0x28aa318991ea876d5bd2bd7238577068adc690	0	8
		131af94ee06d9e61b01e41391d		
Add Consumer	0.00011 ETH	0xd6d91da84aa9f4d9346a7ab7ea088816dbde750	0	4
		d566e8eeeb7f23bdf4489b442		
Produce Item By	0.00151 ETH	0x28169fde8509c6f2996763eba72280b2b15133	0.5 LINK	4
Manufacturer		7d74143b5579e91a3ca8655aa7		
Sell Item By	0.00143 ETH	0x79048f114ea5c0ed737753f973f69c575a4cdbd	0.5 LINK	8
Manufacturer		206d2cd33c69db97c5098f19a		
Purchase Item By	0.00118 ETH	0xeab6ade50d16201b681a95a645499b0ae189f2d	0.4 LINK	4
Distributor		4738c670ab081cb2a39f34d42		
Shipped Item By	0.00106 ETH	0xa52f4a9629c69c40c6550c00f903d1e63133859	0.4 LINK	4
Manufacturer		6184e4552fa27841f8a9860a3		
Received Item By	0.00106 ETH	0x97eeb340ff9fb6b87cb006f4d4005043d7a809f	0.4 LINK	8
Distributor		0479de85ad84024ace520ce43		
Processed	0.00106 ETH	0x6ef2229a3947e4dde0dbe4f800158f78ff1fe9ed	0.4 LINK	4
Item By		e1300274f5c74b537218165a		
Distributor				
Packaged	0.00106 ETH	0xb11966bd5087d70c0baa6ead1e76e6b331b9d7	0.4 LINK	8
Item By		08aa43b8d8d82a997a2c185c3b		
Distributor				
Sell Item By	0.00107 ETH	0xf7c0c2cac67f765a4fadaac4f935bd088b907d1c	0.4 LINK	4
Distributor		ff4be0008328c0c1a68b6918		
Purchase Item By	0.00118 ETH	0x25961c30875f3d9387c27dcf38b570fbccfd7cdf	0.4 LINK	4
Retailer		c371404dc2f226c3d2d7d2fc		
Ship Item By	0.00106 ETH	0xb6d6fc63e6d5d21e53468f0d91193175f623c66	0.4 LINK	4
Distributor		d649e1f4df028d5769c0c536e		
Receive Item By	0.00106 ETH	0xd122e0c86688659a386a83a0a87c34aebf6db1a	0.4 LINK	8
Retailer		8a820676ef14413ae03b886cb		
Sell Item By	0.00106 ETH	0x13f1f0c73363b685445e54956d0f5667a7c0987	0.4 LINK	4
Retailer		aa1290bd1f26ef342d68ba275		
Purchase Item By	0.00118 ETH	0x3e522790ae54650b3880f5bfcbdf9c437f44b12	0.4 LINK	4
Consumer		a8298147fc34f8de417460094		

 TABLE IV

 Performance Evaluation of Implemented PharmaChain.

testing the functionality of the application. Based on roles, different entities have access to different functions within the supply chain, as shown in figure 13. The manufacturer has only access to "Produce Item By Manufacturer", "Sell Item By Manufacturer", and "Ship Item By Manufacturer". Similarly a distributor has access to "Purchase Item By Distributor", "Received Item By Distributor", "Processed Item By Distributor", "Package Item By Distributor", "Sell Item By Distributor" and "Shipped Item By Distributor". A retailer has access to "Purchase Item By Retailer", "Received Item By Retailer", and "Sell Item By Retailer". A consumer has access to "Purchase Item By Consumer". All the roles have access to "Fetch Item Details" and "Verify Authenticity of Product". Different transactions and transaction costs are evaluated in Table IV. These transaction costs can be avoided by implementing a private blockchain and private nodes for Chainlink.

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Retailer View of PharmaChain

Consumer View of PharmaChain

Fig. 13. User Interface for Implemented PharmaChain based on Different Roles.

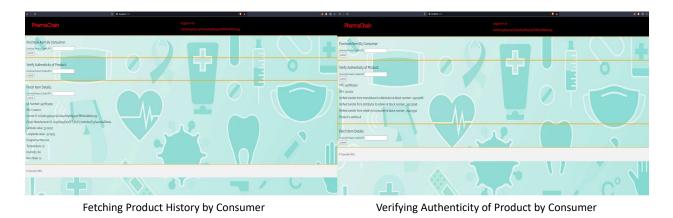


Fig. 14. Consumer Verifying Product Details and Authenticity in Proposed PharmaChain.

B. Validation of PharmaChain

Transaction time is major factor which needs to be evaluated for scalability of the proposed PharmaChain system. As can be seen in Table IV, the average transaction time is 5.6 sec. in the implemented prototype. Transaction times mainly depend on two components: mining times and Chainlink interactions, as the data has to be fetched from the given API. Mining times mainly depend on the transaction volume in the blockchain at that time, but it can be significantly reduced by adopting a private blockchain dedicated to process only transactions from the PSC. The prototype is implemented on a shared test network hence, the performance of the proposed application mainly depends on the performance of the API, as a blockchain test network cannot be controlled or fine tuned for performance. A load test is performed on the data source to determine the scalability and adaptability of the proposed system. To perform load tests, JMeter is used. A load test plan with 100 threads and loop count of 10 is used, which means that 1000 requests are sent to the cloud data source within a span of 2 seconds. From the results in Table V it can be seen that the number of failed transactions is 0 with an average response time of 285ms, which

is very low considering that a large number of oracle requests are sent within a short span. The graph in Figure 15 shows that a large portion of the requests are successful within 500ms, making the proposed architecture for data source scalable and adaptable in real-world PSC.

II IDEE	•
STATISTICS OF LOAD TESTING PERFORMED ON DATA SOU	JRCE IMPLEMENTED IN PROPOSED PHARMACHAIN.
Parameters	Value
Number of Oracle Requests sent	1000
Load Duration	2 Seconds
Failed Requests	0
Percentage of Error	0%

285.196 ms

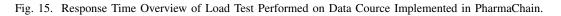
78 ms

16.66

1960 ms

TABLE V
STATISTICS OF LOAD TESTING DEPENDED ON DATA SOUDCE IMPLEMENTED IN DRODOSED DUADMACHAIN

10				
10				
10				
10				
10				
10				
10				
10				
10				
0	Requests having response time <= 500ms	Requests having response time > 500ms and <= 1,500ms	Requests having response time > 1,500ms	Requests in error
		Response times	s ranges	



VI. Conclusion and Future Research

Average Response Time(ms)

Throughput (Requests/Sec)

Minimum Response Time(ms)

Maximum Response Time(ms)

Along with providing a transparent pharmaceutical supply chain, which will enhance the processing times of shipments and smooth operation between different distributed entities, the proposed PharmaChain also equips consumers with very much needed tools to verify the authenticity of the purchased drugs before consumption. By building a transparent chain, the proposed PharmaChain is also enhancing the accountability of the entities involved in the supply chain, thereby making it easy to trace and take severe actions against entities with malicious intent. IoT technology was seamlessly integrated with the blockchain in the proposed system using oracles, along with real-time alerts to the stakeholders regarding different abnormal levels of environmental parameters like ambient temperature and relative humidity around the shipment. This helps in taking prompt actions to control the environment around the package and to ensure the safe handling of the product throughout the supply chain. A robust Role Based Access Control Mechanism was also implemented in the proposed PharmaChain using smart contracts. A proof-of-concept for the whole proposed system has been implemented and analyzed for scalability and reliability. The proof-of-concept was implemented using an Ethereum blockchain with Proof-of-Authority (PoA) as consensus mechanism and analysis of smart contract interactions has shown the average block time is 5.6sec which is acceptable for applications in the supply chain. To address the problem of smart contracts interacting with external APIs, oracles were used. A hybrid smart contract was built in PharmaChain which is capable of interacting with external IoT APIs and of consuming data into smart contracts which ensures the reliability of data being consumed in smart contracts. Scalability of the data source for oracles was also analyzed and the average response time is measured to be 285.196 ms hence proving the proposed PharmaChain application is reliable and scalable to be adopted as a solution to avoiding counterfeit medications in the PSU.

The current work can be extended further by introducing hardware security mechanisms into place like Physical Unclonable Functions (PUF) [37] instead of RSA encryption, which will increase the security of the IoT systems integrated with blockchain and will also reduce power consumption thereby reducing the cost of operations as there will be a very large number of sensing nodes needed throughout the supply chain. We have proposed the idea of PUFchain as an initial contribution in this regard [38]. Most of the supply chain actions are automated but still few

functions are not automated in the current implementation. An IoT mechanism will be implemented in future work to automate the entire handling of medical drugs through the supply chain. In this direction, we think integration of pharmaceutical supply chain in smart healthcare [39] as well as smart cities [40] will have great impact. Thus, research is necessary in these various directions.

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